

# Low detection rate of advanced neoplasia within 5 years after polypectomy of small serrated adenoma

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#### **ABSTRACT**

**Aim** Patients with small serrated adenomas (SAs) (<10 mm) often undergo surveillance colonoscopy before the routine recommended time. We aimed to determine the appropriate surveillance intervals following polypectomy of small SAs for symptomatic patients. **Method** We retrospectively reviewed the data of 638 patients, including 122 cases and 516 controls. Subjects in the case group had small SAs at baseline colonoscopy, including sessile SA/polyps and traditional SAs, while subjects in the control group had negative findings. All patients underwent at least one surveillance colonoscopy during the following 5 years.

**Results** There was no significant difference in the incidence rate of advanced neoplasia between the two groups over a 5-year duration (3.6% vs 2.6%. p=0.455). Moreover, both groups also showed a low prevalence of SA formation over 1-5 years (3.6% vs 1.0%, p=0.145). Patients with baseline SA tended to undergo the first surveillance colonoscopy earlier than those without adenoma ( $\leq 1$  year vs 1 to  $\leq 3$  years). Seventy-one (11.1%) of the total included subjects underwent inadequate initial colonoscopy, and 30 (42.3%) underwent early surveillance of adenoma formation within 1 year. Patients with a family history of colorectal cancer (OR 4.69, 95% CI 1.48 to 14.71, p=0.017) or inadequate baseline colonoscopy (OR 3.17, 95% CI 1.202 to 8.409, p=0.035) were at a higher risk of metachronous adenoma formation during the surveillance period.

**Conclusion** Patients with small SAs at baseline gain little benefit from follow-up of colonoscopy within 5 years after complete polypectomy.

#### INTRODUCTION

The serrated pathway is recognised as a molecular pathway to colorectal cancer (CRC), which accounts for 20%–30% of the population overall.<sup>1</sup> Unlike the chromosomal instability pathway<sup>2</sup> and the Lynch pathway, which develop through conventional adenomas, serrated lesions, especially sessile serrated adenoma/polyps (SSA/Ps) and traditional serrated adenomas (TSAs) with excessive methylcytosine in CpG islands, have been considered as hypermethylated pathways, which may devitalise tumour suppressor genes. 4-6 According to the WHO classification, serrated colorectal lesions can be divided pathologically into three subtypes: hyperplastic polyp (HP), SSA/P and TSA. The incidence of coexisting submucosal invasive carcinoma in patients with baseline HPs was reported as low

as 0.1%, so SSA/P and TSA require more attention than HP due to the higher malignant potential.

Endoscopic resection and endoscopic mucosal resection (EMR) contribute to en bloc resection of small serrated lesions (<10 mm) and have been widely used in the clinic for their feasibility and safety. 9-11 However, patients with a proximal serrated lesion were still at much greater risk of colorectal neoplasia in a follow-up study due to the absence of surveillance for serrated polyps after resection. 12 The serrated pathway is associated with interval cancers during the following 5 years after colonoscopy. 13 Moreover, residual neoplasia of serrated polyps after colonoscopic polypectomy is inevitable, and complete resection becomes increasingly difficult as the size of the lesion increases. 1 14 15 A systematic review and meta-analysis published between January 2006 and July 2015 summarised 11 observational studies (n=64 317) and found that patients with low-risk adenomas were at higher risk of advanced adenomas compared with those with a normal baseline colonoscopy (risk ratio [RR] 1.55, 95% CI 1.24 to 1.94, P=0.0001). 16 Hence, colonoscopy surveillance after complete resection of serrated polyps cannot be ignored.

After polypectomy, more than 75% of the patients did not meet the standard interval of endoscopic monitoring. 17 18 There are various recommended guidelines for endoscopic surveillance intervals after the baseline colonoscopy. Under general consideration, serrated polyps ≥10 mm or any serrated lesions with dysplasia necessitate post-polypectomy colonoscopy surveillance at an interval of 3 years. Patients with serrated polyps < 10 mm without any dysplastic changes are advised to prolong the follow-up interval from 5 years to 10 years. 7 19-22 Moreover, the British Society of Gastroenterology has suggested that regardless of the pathological pattern, no endoscopic surveillance is advised when all serrated polyps are smaller than 10 mm without any dysplastic progression except serrated polyposis syndrome.<sup>23</sup> Interestingly, according to the latest research update, patients with index SSA/Ps or TSAs are at high risk of large metachronous serrated polyps (RR 9.70, 95% CI 3.63 to 25.92) but not metachronous high-risk adenomas.<sup>24</sup> Nevertheless, due to the lack of data in the Asia-Pacific region, these criteria might not be the standard guideline for follow-up in these areas owing to ethnic differences.<sup>25</sup> Hence, we compared the incidence of advanced neoplasia (AN) in the following 5 years of observation between patients with <10 mm SAs and those without adenoma at



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baseline during the same period. This study aimed to investigate an appropriate surveillance interval for SA < 10 mm and to identify risk factors related to subsequent AN formation.

#### **METHODS**

# **Study population**

Consecutive patients were enrolled in this single-centre retrospective study. Patients enrolled in the study underwent colonoscopies from January 2010 to July 2017 at the Digestive Endoscopy Center of the General Hospital, Tianjin Medical University. Patients with SAs from 1 mm to 9 mm and those without any adenoma (hyperplastic polyp or no polyp) at routine baseline observation were recruited as a study population if they had undergone at least one or more follow-up colonoscopy surveillance visit within 5 years of the study. All SAs found at baseline were completely removed by biopsy forceps (for diminutive polyps 1-4 mm) or EMR or electrocautery (for larger ones 5-9 mm). Follow-up data were recorded in the database, and those who chose colonoscopic surveillance at other hospitals were followed up via telephone. The surveillance intervals of the patients' follow-up were divided into three groups after polypectomy: follow-up  $\leq 1$  year, 1 to  $\leq 3$  years and 3 to  $\leq 5$  years.

The following subjects were excluded: (1) patients with inflammatory bowel disease or intestinal tuberculosis; (2) patients with familiar hereditary polyposis syndromes, previous diagnosis of CRC or colon removal; (3) patients having a previous colonoscopy within 5 years before this baseline colonoscopy; (4) patients without complete follow-up data or pathological reports; (5) patients receiving complete polypectomy of SAs during the 3 months prior to study entry.

### Baseline colonoscopy review

All colonoscopic examinations were performed by certified endoscopists, and the endoscope used was an Olympus CF-Q260 (Olympus Optical, Tokyo, Japan). The withdrawal duration of every endoscopy performed was at least 6 min.<sup>26</sup> The general information of the patients was collected from the colonoscopy database (data from case history, colonoscopy report and pathology report), including (1) patients' gender, age and family history of CRC or personal history of polyps; (2) the indications for the initial colonoscopy; (3) the location, size, number and pathological classification of any polyps detected; and (4) the quality of the bowel preparation, which was assessed according to the Boston Bowel Preparation Scale<sup>27</sup> as adequate or inadequate. We used the term 'serrated adenoma' to search the colonoscopy database and finally selected 122 cases with serrated adenomas <10 mm in size. After that, we invited a senior pathologist to re-evaluate the diagnosis. If there was a dispute about the subtype of pathology, three pathologists jointly evaluated the final conclusion. The final diagnosis depended on the pathological outcome.

# First follow-up outcome

We found that most of the subjects included in our study underwent the first follow-up colonoscopic examination due to comprehensive reasons, such as sustained gastrointestinal discomfort, fear of cancerous progression and physicians' recommendations. Several definitions were taken under consideration before analysing the surveillance outcomes. Na AN was defined as having adenomas ≥10 mm, villous histology and high-grade dysplasia or carcinomas. Non-advanced adenomas (NAAs) were defined as having one or more tubular adenomas <10 mm. No adenomas were collectively defined as the presence

of hyperplastic polyps or absence of any polyps. It has been reported that both SSAs and TSAs have excessive methylcytosine in CpG islands with malignant potential. <sup>29 30</sup> SSA/P and TSA were collectively referred to as serrated adenomas (SAs). If more than one polyp was detected during the follow-up colonoscopy, the stratification depended on the most advanced lesion found. The most advanced lesion was defined with the largest diameter or the most severe pathological pattern.

#### STATISTICAL ANALYSIS

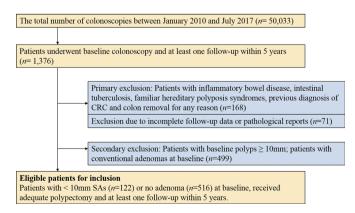
Continuous variables with normal distribution were calculated as the mean  $\pm$  SD, and variables with non-normal distribution were calculated as the median (IQR). Ratio and proportion were used to describe categorical data. The Kruskal-Wallis test followed by the Mann-Whitney U test was used to calculate non-normal data. Proportions were compared using Pearson's  $\chi^2$  test or Fisher's exact probability test. Quantitative data were compared using Student's t-test. Risk factors for subsequent adenoma formation were presented as ORs with 95% CIs. Statistical analyses were performed using SPSS V.17.0 software, and a two-tailed p value <0.05 was considered statistically significant.

#### **RESULTS**

# Patient characteristics at baseline

Our colonoscopy database included 50 033 colonoscopies performed from January 2010 to July 2017. During this period, 1137 patients underwent at least one follow-up colonoscopy within 5 years and met the research criteria described above. A total of 499 patients were excluded since they were found to have conventional adenomas at baseline. The number of patients included in our study was 638, including 122 patients (19.1%) who had small SAs (<10 mm) and 516 (80.9%) controls who had no adenoma (figure 1). A total of 587 cases of serrated lesions were retrieved from the database during the entire study period  $(587/50\ 033=1.17\%)$ . However, in our study, we focused on those with smaller sizes (<10 mm) and those who underwent surveillance colonoscopy in the following 5 years after complete removal. Moreover, patients whose integral information about case history, colonoscopy reports and pathology reports could not be tracked were excluded. Patients with hyperplastic polyps were also excluded. Finally, we included 20.8% individuals with serrated lesions in the current study (122/587).

The demographic and clinical characteristics of the study population are listed in table 1. A total of 37.7% (46/122) baseline SAs were found with low-grade dysplasia. In the whole cohort, 312 (48.8%) were men, and the mean age of the total population



**Figure 1** Inclusion and exclusion diagram of the study population. CRC, colorectal cancer; SA, serrated adenoma.

Table 1 Baseline characteristics of the included subjects who underwent at least one follow-up colonoscopy within 5 years

		Period of time un	Period of time until the follow-up (years)			
Characteristics	n (%)	≤1	1 to ≤3	3 to ≤5		
Total, n (%)	638 (100)	163 (25.5)	328 (51.4)	147 (23.0)		
Sex						
Male, n (%)	312 (48.9)	94 (30.1)	159 (51.0)	59 (18.9)		
Female, n (%)	326 (51.1)	69 (21.2)	169 (51.8)	88 (27.0)		
Mean age at baseline colonoscopy (years), mean±SD	53.2±12.6	53.8±13.0	53.7±12.7	51.7±12.0		
Reasons for baseline colonoscopy, n (%)*						
Screening colonoscopy	135 (21.2)	135 (21.2)	64 (47.4)	64 (47.4)		
Family history of CRC	16 (2.5)	3 (18.8)	10 (62.5)	3 (18.8)		
Previous diagnosis of polyps	87 (13.6)	35 (40.2)	36 (41.4)	16 (18.4)		
Positive FOBT, bleeding or anaemia	85 (13.3)	25 (29.4)	45 (52.9)	15 (17.6)		
Diarrhoea or/and constipation	179 (28.1)	42 (23.5)	102 (57.0)	35 (19.6)		
Other gastrointestinal symptoms†	323 (50.6)	82 (25.4)	152 (47.1)	89 (27.6)		
Baseline colonoscopy quality, n (%)						
Excellent preparation	262 (41.1)	63 (24.0)	142 (54.2)	57 (21.8)		
Good preparation	223 (35.0)	43 (19.3)	125 (56.1)	55 (24.7)		
Fair preparation	93 (14.6)	31 (33.3)	41 (44.1)	21 (22.6)		
Poor preparation	60 (9.4)	26 (43.3)	20 (33.3)	14 (23.3)		
Could not reach the caecum	22 (3.4)	10 (45.5)	11 (50.0)	1 (4.5)		
Poor preparation and could not reach the caecum	11 (1.7)	6 (54.5)	4 (36.4)	1 (9.1)		

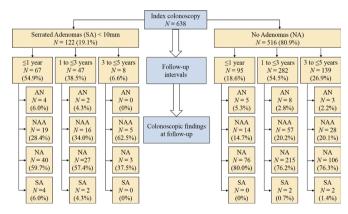
<sup>\*</sup>Patients might have more than one indication.

was 53.2±12.6 years. A total of 163 (25.5%) subjects underwent follow-up colonoscopy within 1 year, 328 (51.4%) within 1-3 years and 147 (23.0%) within 3-5 years. Patients with one, two, and three or more SAs at the initial colonoscopy underwent surveillance colonoscopy after 13 (3–63), 12 (6–42) and 11 (6–28) months, respectively (p=0.809). There was no significant difference in the follow-up interval in patients with a single SA between 1-5 mm and 6-9 mm observed at baseline (12 [3-63] vs 10 [3-48], p=0.129). Most of the baseline colonoscopies were performed to make a definitive diagnosis, including positive faecal occult blood test/lower gastrointestinal haemorrhage/anaemia (85 [13.3%]), changes in bowel habit (diarrhoea or/and constipation) (179 [28.1%]) and any other gastrointestinal symptoms (abdominal pain, distention and other abdominal discomfort) (323 [50.6%]). Additionally, some patients underwent baseline colonoscopies as they had a positive family history of CRC (16 [2.5%]), a previous finding of polyps (87 [13.6%]) or routine cancer screening (135 [21.2%]). Of the baseline colonoscopies, 71 (11.1%) patients underwent inadequate examinations, including poor bowel preparation (60 [9.4%]) and no caecum-reached operation (22 [3.4%]). Of the total baseline colonoscopies performed, 71 (11.2%) patients had insufficient visualisation of the gastrointestinal tract due to poor bowel preparation (60 [9.4%]) or an incomplete procedure due to the difficulty of the colonoscope in reaching the caecum (22 [3.4%]).

# Colonoscopic and pathological findings at baseline and follow-up

Of the 638 individuals, 171 (26.8%) had metachronous adenomas, among which 22 (3.4%) had AN, 139 (21.8%) had NAA and 10 (1.6%) had SA during the follow-up period, and the median follow-up time was 22 months (figure 2). Although none of the baseline adenomas were ever found in the control group, the number of patients with at least one follow-up adenoma was 119/516 (23.1%). Among them, 16 (3.1%) had AN (two of them were detected as high-grade intraepithelial neoplasia,

eight of them had tubular adenomas  $\geq 10$  mm and six of them had villous ingredients), 99 (19.2%) had NAA and 4 (0.8%) had SA. However, when taking only these detected baseline SAs into account, 42.6% (52/122) patients had adenoma formation at follow-up, of whom 6 (4.9%) had AN (none of them were detected as high-grade intraepithelial neoplasia, half of them had tubular adenomas  $\geq 10$  mm and the other half had villous ingredients), 40 (32.8%) had NAA and 6 (4.9%) had SA. We found that NAAs ranged in size from 1 to 8 mm, with 52.5% (21/40) associated with low-grade dysplasia. Nevertheless, there were no significant differences in the incidence of AN and SA during the 5 years of study between patients with baseline SA and without any baseline adenomas (AN, 3.6% vs 2.6% p=0.455; SA, 3.6% vs 1.0% p=0.145).



**Figure 2** Endoscopic findings at baseline and follow-up colonoscopy in the included subjects. AN, advanced neoplasia; NA, no adenoma; NAA, non-advanced adenoma; SA, serrated adenoma including sessile serrated adenoma/polyp and traditional serrated adenoma. Both groups, even those with no adenomas at baseline, underwent follow-up colonoscopies within 5 years for comprehensive reasons.

<sup>†</sup>Other gastrointestinal symptoms include abdominal pain, distention or any other discomfort.

CRC, colorectal cancer; FOBT, faecal occult blood test.

 Table 2
 Advanced neoplasia and serrated adenoma found at follow-up colonoscopy in included subjects

	Follow-up	Follow-up AN			Follow-up SA			
Baseline	≤1	1 to ≤3	3 to ≤5	P value*	≤1	1 to ≤3	3 to ≤5	P value*
Case group, n (%)	4 (6.0)	2 (4.3)	0 (0.0)	0.728	4 (6.0)	2 (4.3)	0 (0.0)	0.728
Control group, n (%)	5 (5.3)	8 (2.8)	3 (2.2)	0.481	0 (0.0)	2 (0.7)	2 (1.4)	0.402

<sup>\*</sup>Cohort 1-3 years vs 3-5 years.

AN, advanced neoplasia; SA, serrated adenoma.

Most of the patients (54.9%) in the SA group underwent the first surveillance colonoscopy within 1 year. Meanwhile, those without any adenoma at baseline would delay their surveillance until 1 to 3 years later (54.5%). In the whole population, the proportions of patients who had AN or SA at three intervals did not differ significantly. Nine patients (5.6%) had AN within a 1-year interval, ten patients (3.0%) had AN at 1-3 years and three patients (2.0%) had AN at 3-5 years (p=0.202). The risks of SA at the three intervals were 2.5%, 1.2% and 1.4% at 1 year, 1-3 years and 3-5 years, respectively (p=0.591). The risk of AN on the first surveillance colonoscopy was similar in the patients with intervals of 1-3 years (4.3%) and 3-5 years (0.0%), which all had SA at baseline (p=0.728) (table 2). SA also had a similar incidence in the two surveillance intervals (4.3% vs 0.0%, p=0.728). Likewise, in the cohort without any baseline adenomas, different surveillance intervals (1-3 years and 3-5 years) did not significantly influence the risk of AN and SA detected at follow-up (AN at 1-3 years and 3-5 years, 2.8% vs 2.2%, p=0.481; SA at 1-3 years and 3-5 years, 0.7% vs 1.4%, p=0.402). Hence, after adequate polypectomy, patients with SA < 10 mm at baseline might not be at a greater risk for subsequent AN or SA formation within 5 years.

### Risk factors affecting subsequent adenoma formation

A series of baseline characteristics that might influence follow-up adenoma formation are shown in table 3, which are presented as ORs with 95% CIs. By analysing the whole population, age over 50 years (OR 1.45, 95% CI 1.02 to 2.08), male gender

(OR 1.81, 95% CI 1.27 to 2.58) and a history of polyps (OR 2.73, 95% CI 1.71 to 4.34) predict the risk of any subsequent adenoma formation. Patients with a family history of CRC (OR 4.67, 95% CI 1.48 to 14.71) were at greater risk for subsequent AN than those without. The case group was also independently assessed the metachronous adenoma formation according to adenoma features at baseline. Adenoma features, including the number, maximum diameter and location, did not significantly influence subsequent adenoma formation within 5 years.

# Quality of baseline colonoscopy associated with follow-up outcome

Seventy-one (11.1%) patients underwent inadequate baseline colonoscopies, including inadequate bowel preparation (60, 9.4%), and the operation did not reach the caecum (22, 3.4%). Nearly half of them (30, 42.3%) had received the initial surveillance colonoscopies within 1 year. Table 4 shows the relationship between the quality of colonoscopy at baseline and subsequent adenoma detection. The demographic characteristics, including age and gender, did not influence the completion of examination at the initial colonoscopy. Inadequate baseline colonoscopies contributed to a lower adenoma detection rate (ADR) compared with adequate examination at the initial colonoscopy (16.9% vs 28.0, p=0.046). More AN and  $\geq$ 3 NAA could be found in patients with inadequate baseline colonoscopy than in those with adequate baseline examination (8.5% vs 2.8%, p=0.035; 11.3% vs 4.4%, p=0.014). Incomplete baseline colonoscopy

Baseline		Any adenoma in the follow-up			AN in the follow-up		
	Total, n	n (%)	OR (95% CI)	P value	n (%)	OR (95% CI)	P value
Age (years)							
Aged ≥50	349	105 (30.1)	1.45 (1.02 to 2.08)	0.040	11 (3.2)	0.82 (0.35 to 1.93)	0.652
Aged <50	289	66 (22.8)			11 (3.8)		
Sex							
Male	312	102 (32.7)	1.81 (1.27 to 2.58)	0.001	13 (4.2)	1.53 (0.65 to 3.64)	0.331
Female	326	69 (21.2)			9 (2.8)		
Family history of CRC							
With	32	12 (37.5)	1.60 (0.77 to 3.35)	0.206	4 (12.5)	4.67 (1.48 to 14.71)	0.017
Without	606	159 (26.2)			18 (3.0)		
History of polyps							
With	87	40 (46.0)	2.73 (1.71 to 4.34)	< 0.001	6 (6.9)	2.48 (0.94 to 6.51)	0.103
Without	551	131 (23.8)			16 (2.9)		
Adenoma features*							
≥2 adenomas	30	19 (63.3)	1.44 (0.67 to 3.10)	0.351	1 (3.3)	0.41 (0.05 to 3.64)	0.707
One adenoma	92	33 (35.9)			5 (5.4)		
6–9 mm adenoma	41	19 (46.3)	1.26 (0.59 to 2.68)	0.555	4 (9.8)	4.27 (0.75 to 24.37)	0.189
≤5 mm adenoma	81	33 (40.7)			2 (2.5)		
Distal adenoma	73	36 (49.3)	2.01 (0.95 to 4.26)	0.068	3 (4.1)	0.66 (0.13 to 3.40)	0.939
Proximal adenoma	49	16 (32.7)			3 (6.1)		

<sup>\*</sup>Adenoma features were compared among patients with baseline SA.

AN, advanced neoplasia; CRC, colorectal cancer.

**Table 4** Comparison of the characteristics of the included subjects according to the quality of baseline colonoscopy

	Baseline colonoscopy				
Characteristics, n (%)	Inadequate examination 71 (11.1%)	Adequate examination 567 (88.9%)	P value		
Aged ≥50	35 (49.3)	314 (55.4)	0.340		
Male	37 (52.1)	275 (48.5)	0.566		
ADR at baseline	12 (16.9)	159 (28.0)	0.046*		
Follow-up results					
AN	6 (8.5)	16 (2.8)	0.035*		
SA	2 (2.8)	8 (1.4)	0.369		
≥3 NAA	8 (11.3)	25 (4.4)	0.014*		
1-2 NAA	14 (19.7)	92 (16.2)	0.456		

<sup>\*</sup>Significant difference between the group with inadequate baseline examination and those with an adequate one.

ADR, adenoma detection rate; AN, advanced neoplasia; NAA, non-advanced adenoma; SA, serrated adenoma.

increased follow-up AN formation to a 3.17 OR risk (95% CI 1.202 to 8.409). Taken together, these findings strongly suggest that patients with inadequate baseline colonoscopy should have a shorter surveillance interval.

### **DISCUSSION**

A certain percentage of asymptomatic populations over 50 years have not undergone colonoscopy screening due to a lack of proper health awareness and financial crisis. On the other hand, frequent surveillance colonoscopy is suggested by physicians and endoscopists for low-risk patients, which possess considerable pressure due to limited available endoscopic resources. The current study, based on a representative Chinese cohort, assessed post-polypectomy colonoscopy surveillance of SAs < 10 mm during the following 5 years. During this study, we found no significant risk differences between the baseline SAs and those without any adenoma after complete polypectomy in the progression of AN within 5 years. A shorter surveillance interval is needed for patients with a family history of CRC or inadequate baseline colonoscopy. In Asian countries, there are no specific guidelines for surveillance colonoscopy after polypectomy. The study discussed here is both important and interesting owing to its contribution to the establishment of guidelines for colonoscopy surveillance after polypectomy of small SSA/Ps or TSAs.

Serrated lesions play a crucial role in the carcinogenesis of the colon. 31 32 The risk of CRC in patients with serrated polyps is strongly associated with the size of the polyps. Serrated polyps ≥10 mm have been considered to be an independent risk factor for synchronous AN (OR 3.24, 95% CI 2.05 to 5.13, p<0.0001).<sup>33</sup> Patients with at least one large proximal non-dysplastic serrated polyp (≥10 mm) at a screening colonoscopy have been reported to have synchronous AN (OR 3.37, 95% CI 1.71 to 6.65). 12 Moreover, large serrated polyps  $\geq$  10 mm were at a 2.5 OR risk to develop CRC compared with the population in average risk during an 11-year median follow-up.<sup>34</sup> Unlike above, the proportions of high-grade dysplasia coexisting with SSA/Ps and TSAs were 0.7% and 1.0%, respectively, which were much smaller than the proportion coexisting with high-grade dysplasia in conventional adenomas (5%).<sup>35</sup> The risk of AN formation in patients with proximal serrated polyps <10 mm at baseline was consistent with those with low-risk adenoma during surveillance. 12 The time period required for the progression of SSAs to carcinoma was assumed to be 10-15 years based on a

large cohort study including 2139 patients, which concluded that SSAs have a low cancerous progression rate. <sup>36</sup>

Interestingly, Maratt *et al*<sup>37</sup> identified several demographic and endoscopic characteristics, including family history of CRC and endoscopist screening ADR, in connexion with the high detection rate of SSA/Ps at baseline. Notably, a recent study found that baseline SA alone might be a risk predictor for the formation of a subsequent large serrated polyp, and synchronous baseline SA and high-risk adenoma put patients at high risk of cancer progression.<sup>38</sup> Further studies are necessary to determine the contradicting findings.

A meta-analysis found that baseline colonoscopy with excellent bowel preparation and adequate caecal intubation could maximally decrease the risk of interval cancer for identifying missed serrated lesions as efficiently as possible. <sup>39</sup> Low-quality colonoscopy screening, which has adherent mucus and faecal materials, would contribute to the formation of advanced neoplasia in short follow-up intervals. On the other hand, early AN formation from SA is partly attributed to incomplete resection. Research indicates that the incomplete resection rate of SSA (30%) is apparently higher than that of conventional adenoma (10%) because of its flat shape. <sup>14 40</sup> Fortunately, EMR following contrast injection under a high-definition colonoscope delineates the lesion edges of SAs and allows complete resection. <sup>141</sup>

The limitations of this study should not be ignored. First, the patients with SSAs and those with TSAs were sorted into the same group because of the limited sample size. Further research should be performed to assess the risk rate of progression of adenoma–carcinoma in different subsets of SAs. Second, in a representative cohort of the Caucasoid population including 2416 SSA/Ps in 2139 patients, only 14% were identified with cytological dysplasia.<sup>36</sup> In another cohort containing 1910 baseline colonoscopies, only 0.6% sessile serrated polyp (SSP) had dysplastic histology.<sup>42</sup> Both dysplastic and non-dysplastic serrated polyps smaller than 10 mm were included in the procedures. Sessile serrated polyps with cytological dysplasia are more advanced lesions than non-dysplasia serrated polyps, and TSAs are rare with uniform dysplasia.<sup>43</sup> Larger-scale cohorts are needed to singly determine the optional follow-up intervals for

## Main messages

- Patients with serrated adenomas (SAs) <10 mm at baseline seemed to have little benefit from follow-up colonoscopy within 5 years after complete polypectomy.
- Patients with a family history of colorectal cancer (CRC) or inadequate baseline colonoscopy need a shorter surveillance interval.
- ► Age over 50 years, male gender and a history of polyps predict the risk of any subsequent adenoma formation.

## **Current research questions**

- ▶ Do patients with serrated adenomas <10 mm with dysplasia need to undergo surveillance colonoscopy within a shorter follow-up interval compared with those without cytological dysplasia?
- ► When should patients with small SA with a family history of CRC undergo follow-up colonoscopy after complete removal?
- ► What is a better way to identify patients with inadequate baseline colonoscopy?

# Original article

SAs <10 mm with or without cytological dysplasia. Our research was conducted in a large medical centre. In the future, we will combine our data with several medical centres to further validate our conclusions and strengthen the long-term follow-up of patients with SAs.

**Contributors** DJ, HC and BW designed the research. DJ, DC, TL, WD and SW collected data. YZ was responsible for pathological analysis. DJ and YQ performed statistical analysis. DJ, BKS and HC wrote the paper. All authors were involved in the final approval of the manuscript.

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Competing interests None declared.

#### Patient consent for publication Obtained.

**Ethics approval** The study procedures were approved by the Institutional Review Board of the General Hospital, Tianjin Medical University, China.

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