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WJGO mainly publishes articles reporting research results and findings obtained in the field of gastrointestinal oncology and covering a wide range of topics including liver cell adenoma, gastric neoplasms, appendiceal neoplasms, biliary tract neoplasms, hepatocellular carcinoma, pancreatic carcinoma, cecal neoplasms, colonic neoplasms, colorectal neoplasms, duodenal neoplasms, esophageal neoplasms, gallbladder neoplasms, etc.

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Retrospective Cohort Study

## Trends in colorectal cancer incidence according to an increase in the number of colonoscopy cases in Korea

Ga Hee Kim, Yeong Chan Lee, Tae Jun Kim, Sung Noh Hong, Dong Kyung Chang, Young-Ho Kim, Dong-Hoon Yang, Chang Mo Moon, Kyunga Kim, Hyun Gun Kim, Eun-Ran Kim

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### Abstract

#### BACKGROUND

The incidence of colorectal cancer (CRC) and preinvasive CRC (*e.g.*, early colon cancer and advanced adenoma) is gradually increasing in several countries.

#### AIM

To evaluate the trend in incidence of CRC and preinvasive CRC according to the increase in the number of colonoscopies performed in Korea.

#### METHODS

This retrospective cohort study enrolled Korean patients from 2002 to 2020 to evaluate the incidence of CRC and preinvasive CRC, and assess the numbers of diagnostic colonoscopies and colonoscopic polypectomies. Colonoscopy-related complications by age group were also determined.

## RESULTS

The incidence of CRC showed a rapid increase, then decreased after 2012 in the 50-75 year-age group. During the study period, the rate of incidence of preinvasive CRC increased at a similar level in patients under 50 and 50-75 years of age. Since 2009, the increase has been rapid, showing a pattern similar to the increase in colonoscopies. The rate of colonoscopic polypectomy in patients aged under 50 was similar to the rate in patients over 75 years of age after 2007. The rate of complications after colonoscopy and related deaths within 3 mo was high for those over 75 years of age.

## CONCLUSION

The diagnosis of preinvasive CRC increased with the increase in the number of colonoscopies performed. As the risk of colonoscopy-related hospitalization and death is high in the elderly, if early lesions at risk of developing CRC are diagnosed and treated under or at the age of 75, colonoscopy-related complications can be reduced for those aged 76 years or over.

**Key Words:** Colorectal cancer; Preinvasive colorectal cancer; Colorectal polypectomy; Colonoscopy

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**Core Tip:** In Korea, colorectal cancer (CRC) showed the second highest incidence rate among all cancers in 2018. The volume of colonoscopies and colorectal polypectomies in Korea is also increasing due to the CRC screening program initiated in 2004, and rapid population aging. The purpose of this study was to evaluate the trend in incidence of CRC and preinvasive CRC according to the increase in the number of colonoscopies performed in Korea.

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## INTRODUCTION

Colorectal cancer (CRC) is the third most commonly diagnosed cancer with an estimated 1.8 million new cases in 2018, and is the second leading cause of cancer-related death in the world. In Korea, CRC showed the second highest incidence rate among all cancers in 2018, with an estimated 44.5 cases per 100000 persons per year[1,2]. Following a rapid decline during the 2000s, the CRC incidence rate in men and women has remained generally stable. This decline has been greatly affected by screening colonoscopy and the removal of colorectal precancerous lesions[3]. Most CRCs occur at the age of 50 to 70 years. Thus, several guidelines have recommended CRC screening colonoscopy for those over 50 years of age[4,5]. However, over the last few decades, the incidence of CRC has increased in younger patients aged < 50 years[6]. In Korea, due to the low cost of colonoscopy and availability of many skilled colonoscopists, access to screening colonoscopy from a young age is different to that in Western countries[7]. Several early CRCs, including precancerous lesions, have been detected by screening colonoscopy. Endoscopic resection procedures include polypectomy, endoscopic mucosal resection, and endoscopic submucosal dissection. Using these procedures, early diagnosis and treatment of early CRCs can be performed[8,9]. In Korea, the volume of colonoscopies and colorectal polypectomies in the elderly aged 75 years or older is also increasing due to the CRC screening program initiated in 2004 and rapid population aging[10]. In this context, we assessed national population-based data on CRC occurrence by age and gender. In particular, the purpose of this study was to evaluate the trend in incidence of CRC and preinvasive CRC according to age group and gender and to determine whether age adjustment for endoscopy during health examination would be necessary in the future. In addition, we attempted to confirm colonoscopy-related complications according to age and gender.

## MATERIALS AND METHODS

### Dataset

We used the National Health Insurance Service (NHIS) claims database to identify trends in CRCs and colorectal polypectomy in South Korea. We selected CRC patients and those who underwent colorectal polypectomy from the NHIS claims database that includes approximately 50 million persons, almost the entire population of South Korea. The database is deidentified with regard to gender, age, and medical history. The study population was followed up for 19 years from 2002 to 2020. We excluded patients who had data missing on gender, age, or imprecise data on death. The study protocol was approved by the Institutional Review Board of Samsung Medical Center (2019-08-140).



### Definition of outcome of interest

We defined CRC patients as those who were diagnosed with CRC [International Classification of Diseases version 10 (ICD-10) code C18-C21]. Preinvasive CRC patients were defined as those with ICD-10 codes D010, D011, and D012. Patients who underwent a colorectal polypectomy were defined as those with treatment codes of Q7701-03 and QX706. In addition, patients who underwent a colonoscopy without colorectal polypectomy were defined as those with the treatment code E7660.

### Study design for trends in CRC and colorectal polypectomy

To identify trends in CRC and colorectal polypectomy, an incident case was defined as a new case without diagnostic history for the target disease right before 2 years, while a prevalent case was defined as a case with the target disease every year. We classified subjects into three age groups: < 50 years, 50-75 years, and  $\geq 76$  years. The number of diagnostic colonoscopies was calculated as the sum of cases of colonoscopy alone and cases of polypectomy, minus the number of cases counted as one if both colonoscopy and polypectomy were performed within 1 year.

### Study design for identifying the associations between age, hospitalizations and deaths after colonoscopy

We analyzed whether patients who underwent colonoscopy from January 2002 to December 2020 were assigned a diagnostic code for colonoscopy-related complications. We determined whether patients undergoing colonoscopy were given diagnostic codes for gastrointestinal (GI)-related complications of 'intestinal perforation' [K63.1, (ICD-10)] and 'intestinal bleeding' [K92.0-92.2, (ICD-10)] and non-GI-related complications such as heart disease [I21 and I50.0, (ICD-10)] and brain disease [I60-64, (ICD-10)] that might contribute to mortality. We also assessed whether patients with these codes were hospitalized or died within 1 mo or 3 mo using trend analysis.

### Statistical analysis

We measured rates of prevalence, hospitalization, and death in those with GI- and non-GI complications after colonoscopy within 1 mo or 3 mo between 2002 and 2020. Then, we performed trend analyses using Cochran-Armitage trend tests to investigate the associations between age and complications. Statistical significance was considered when the *P* value was less than 0.05. Data processing and statistical analyses were performed with SAS software version 9.4 (SAS Institute, Cary, NC, United States) and R software (version 4.0.3; R Foundation for Statistical Computing, Vienna, Austria).

## RESULTS

### Incidence of CRC and preinvasive CRC between 2004 and 2020

A total of 430696 patients were diagnosed with CRC between 2004 and 2020 (Table 1). During the study period, the incidence of CRC was higher in men than in women over all periods. In particular, the incidence of CRC among men aged 50-75 years increased significantly from 2004 to 2012. The increase in CRC has slowed since 2012 in both men and women (Figure 1A). Between 2004 and 2020, a total of 52069 patients were diagnosed with preinvasive CRC (Table 2). Similar to the incidence of CRC, the incidence of preinvasive CRC during all periods was also more common in men. Since 2009, the incidence of preinvasive CRC has shown a sharp rise in both men and women, especially in men, and in those aged 50-75 years (Figure 1B). We also calculated the rate of incidence of preinvasive CRC (the ratio of the incidence of preinvasive CRC to the sum of the incidence of CRC and preinvasive CRC). Although the incidence of CRC continued to increase while the rate of its increase decreased from 2012, during the study period the rate of incidence of preinvasive CRC increased in all age groups. The rate of preinvasive CRC increase in patients under 50 years of age was similar to that in patients aged 50-75 years (Supplementary Table 1, Figure 2).

### Trends in the number of colonoscopies between 2002 and 2020

In this study, we identified the total number of patients who underwent diagnostic colonoscopy (colonoscopy without polypectomy) and colonoscopic polypectomy, respectively, from 2002 to 2020 in the study database (Table 3). The number of patients who underwent diagnostic colonoscopy increased since 2002 in all age groups. The number of patients in 2020 had increased by approximately fifteen-fold compared to that in 2002. In particular, it showed a gradual increase after 2002. It then showed a sharp increase after 2010 in the 50-75 years age group. This pattern also appeared in the age group of under 50 years. However, the total number of patients who underwent diagnostic colonoscopy was lower than that in the 50-75 years age group, the age group that is subjected to screening colonoscopy. The pattern of change in the number of patients who underwent a colonoscopic polypectomy was similar to that of diagnostic colonoscopy (Table 3, Figure 3A). The number of patients who underwent a colonoscopic polypectomy gradually increased since 2002 in all age groups. Since 2010, it has shown a sharp increase among those under 75, especially in the 50-75 years age group. For those aged 76 years or older, colonoscopic polypectomies increased significantly in 2020, compared to 2002. The rate of colonoscopic polypectomies (the ratio of colonoscopic polypectomies to the sum of diagnostic colonoscopies and colonoscopic polypectomies) also steadily increased during the study period (Table 3, Figure 3B). The rate of colonoscopic polypectomies in 2020 increased by approximately two-fold compared to that in 2002. The increase in the rate of colonoscopic polypectomies in the under 50 age group showed a relatively steep increase, compared to other age groups. After 2018, the rate of colonoscopic polypectomies for patients under 50 years was slightly higher than those in the other two age groups.

**Table 1 The incidence of colorectal cancer by age and gender between 2004 and 2020, *n* (%)**

	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020
Number of patients	15278	17090	20059	20750	21613	24179	24969	28342	29033	28480	27866	27624	29528	29452	29204	30025	27204
Age category (yr)																	
Male	8948	9999	11709	12283	12786	14347	14991	17112	17530	17086	16674	16462	17680	17497	17479	17877	16178
< 50	1613 (18.0)	1740 (17.4)	1834 (15.7)	1728 (14.1)	1830 (14.3)	1728 (12.0)	1834 (12.2)	2077 (12.1)	2032 (11.6)	1930 (11.3)	1741 (10.4)	1582 (9.6)	1677 (9.5)	1653 (9.4)	1592 (9.1)	1555 (8.7)	1491 (9.2)
50-75	6499 (72.6)	7306 (73.1)	8603 (73.5)	9167 (74.6)	9437 (73.8)	10730 (74.8)	11137 (74.3)	12608 (73.7)	12764 (72.8)	12257 (71.7)	11894 (71.3)	11635 (70.7)	12382 (70.0)	12151 (69.4)	11929 (68.2)	12234 (68.4)	10940 (67.6)
≥ 76	836 (9.3)	953 (9.5)	1272 (10.9)	1388 (11.3)	1519 (11.9)	1889 (13.2)	2020 (13.5)	2427 (14.2)	2734 (15.6)	2899 (17.0)	3039 (18.2)	3245 (19.7)	3621 (20.5)	3693 (21.1)	3958 (22.6)	4088 (22.9)	3747 (23.2)
Female	6330	7091	8350	8467	8827	9832	9978	11230	11503	11394	11192	11162	11848	11955	11725	12148	11026
< 50	1284 (20.3)	1350 (19.0)	1482 (17.7)	1357 (16.0)	1384 (15.7)	1460 (14.8)	1364 (13.7)	1559 (13.9)	1539 (13.4)	1413 (12.4)	1365 (12.2)	1261 (11.3)	1239 (10.5)	1358 (11.4)	1298 (11.1)	1261 (10.4)	1204 (10.9)
50-75	4302 (68.0)	4767 (67.2)	5624 (67.4)	5731 (67.7)	5929 (67.2)	6555 (66.7)	6596 (66.1)	7388 (65.8)	7430 (64.6)	7243 (63.6)	6980 (62.4)	6818 (61.1)	7092 (59.9)	6948 (58.1)	6630 (56.5)	6931 (57.1)	6208 (56.3)
≥ 76	744 (11.8)	974 (13.7)	1244 (14.9)	1379 (16.3)	1514 (17.2)	1817 (18.5)	2018 (20.2)	2283 (20.3)	2534 (22.0)	2738 (24.0)	2847 (25.4)	3083 (27.6)	3517 (29.7)	3649 (30.5)	3797 (32.4)	3956 (32.6)	3614 (32.8)

**Table 2 The incidence of preinvasive colorectal cancer by age and gender between 2004 and 2020, *n* (%)**

	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020
Number of patients	165	186	325	598	959	1462	2078	3045	3434	3531	3635	4428	5422	5476	6017	5886	5422
Age category (yr)																	
Male	96	125	202	380	601	968	1376	1979	2255	2320	2344	2862	3454	3481	3879	3726	3368
< 50	13 (13.5)	22 (17.6)	43 (21.3)	64 (16.8)	90 (15.0)	164 (16.9)	195 (14.2)	269 (13.6)	286 (12.7)	257 (11.1)	292 (12.5)	294 (10.3)	391 (11.3)	348 (10.0)	407 (10.5)	374 (10.0)	327 (9.7)
50-75	79 (82.3)	97 (77.6)	151 (74.8)	299 (78.7)	477 (79.4)	752 (77.7)	1104 (80.2)	1584 (80.0)	1808 (80.2)	1863 (80.3)	1829 (78.0)	2249 (78.6)	2694 (78.0)	2742 (78.8)	2988 (77.0)	2888 (77.5)	2593 (77.0)
≥ 76	4 (4.2)	6 (4.8)	8 (4.0)	17 (4.5)	34 (5.7)	52 (5.4)	77 (5.6)	126 (6.4)	161 (7.1)	200 (8.6)	223 (9.5)	319 (11.1)	369 (10.7)	391 (11.2)	484 (12.5)	464 (12.5)	448 (13.3)
Female	69	61	123	218	358	494	702	1066	1179	1211	1291	1566	1968	1995	2138	2160	2054
< 50	18 (26.1)	14 (23.0)	31 (25.2)	37 (17.0)	69 (19.3)	101 (20.4)	123 (17.5)	192 (18.0)	155 (13.1)	173 (14.3)	181 (14.0)	220 (14.0)	195 (9.9)	236 (11.8)	223 (10.4)	250 (11.6)	236 (11.5)
50-75	45 (65.2)	46 (75.4)	83 (67.5)	164 (75.2)	271 (75.7)	367 (74.3)	527 (75.1)	790 (74.1)	919 (77.9)	917 (75.7)	982 (76.1)	1167 (74.5)	1503 (76.4)	1466 (73.5)	1569 (73.4)	1538 (71.2)	1452 (70.7)
≥ 76	6 (8.7)	1 (1.6)	9 (7.3)	17 (7.8)	18 (5.0)	26 (5.3)	52 (7.4)	84 (7.9)	105 (8.9)	121 (10.0)	128 (9.9)	179 (11.4)	270 (13.7)	293 (14.7)	346 (16.2)	372 (17.2)	366 (17.8)

**Hospitalizations and deaths after colonoscopy between 2002 and 2020**

Among patients who underwent a colonoscopy, those who were additionally diagnosed with GI- and non-GI-related complications were classified by age group. We also determined whether patients were hospitalized or died within 1 mo and 3 mo (Supplementary Table 2). As age increased, the number of complications and related hospitalizations and deaths increased. In particular, the rate of complications after colonoscopy and related deaths within 3 mo were high for those aged 76 years or older.

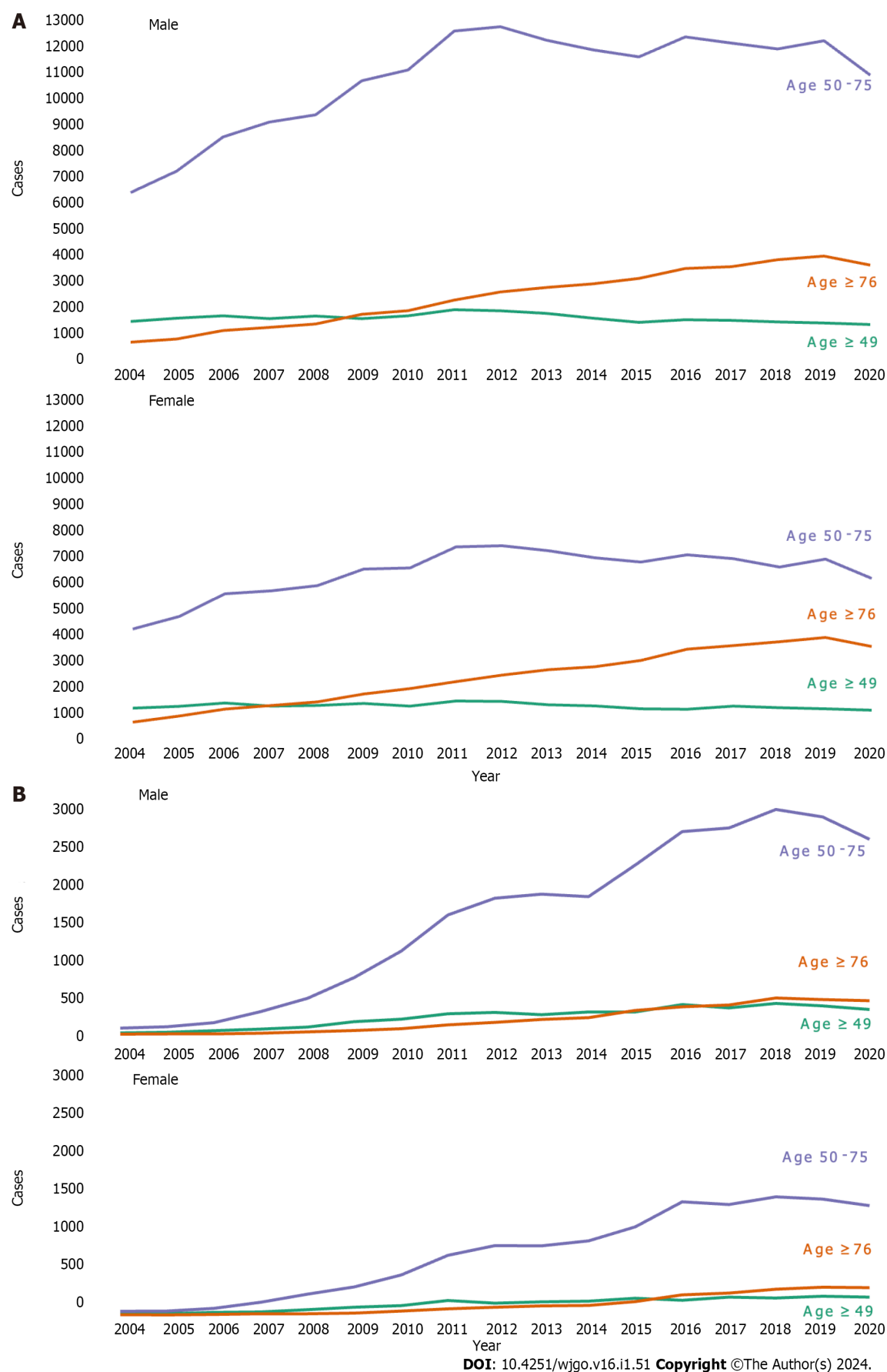


**Table 3** The number of diagnostic colonoscopies (colonoscopy without polypectomy), colonoscopic polypectomies, and the proportion of colonoscopic polypectomies by age between 2002 and 2020

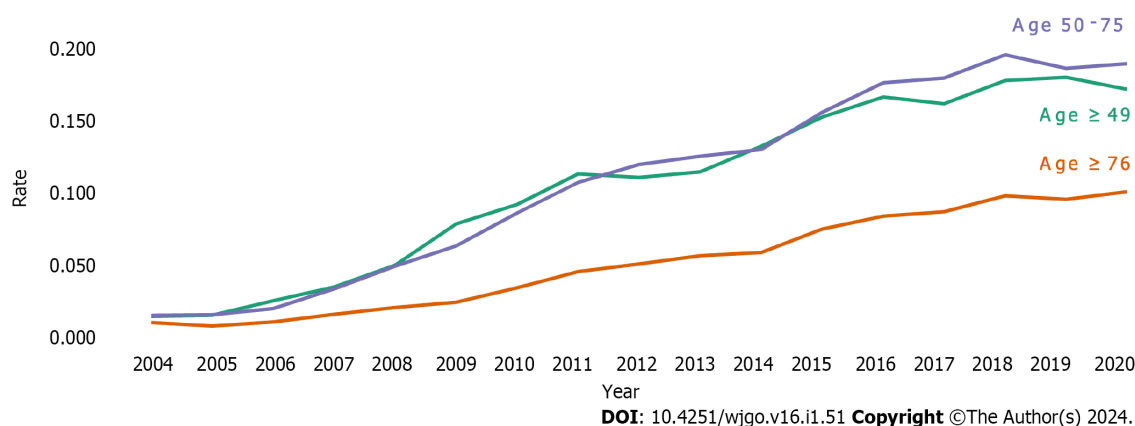
Diagnostic colonoscopy										
Age category (yr)	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011
< 50	29611	38996	42182	54209	66858	69898	75540	84326	78133	179675
50-75	48232	64589	74806	98356	126196	144205	167559	203449	225951	467005
≥ 76	2902	3619	4466	5932	7931	9063	10580	13188	15461	28063
Number of patients	80745	107204	121454	158497	200985	223166	253679	300963	319545	674743
Age category (yr)	2012	2013	2014	2015	2016	2017	2018	2019	2020	
< 50	181116	174704	174746	182356	200147	222661	238393	270541	272869	
50-75	540231	547652	553156	603368	688868	726703	791814	893072	902040	
≥ 76	34984	40693	44484	52028	61053	68676	79313	90198	92604	
Number of patients	756331	763049	772386	837752	950068	1018040	1109520	1253811	1267513	
Colonoscopic polypectomy										
Age category (yr)	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011
< 50	6452	7393	9466	14679	21307	26839	32929	38743	40388	96182
50-75	14373	17884	24041	36060	52212	65697	83504	105620	124745	267398
≥ 76	713	863	1156	1748	2769	3376	4444	5764	7393	14924
Number of patients	21538	26140	34663	52487	76288	95912	120877	150127	172526	378504
Age category (yr)	2012	2013	2014	2015	2016	2017	2018	2019	2020	
< 50	102224	102977	105681	112414	124598	147501	164400	196231	207971	
50-75	324912	337916	347156	382029	440815	484016	544244	635468	668497	
≥ 76	19942	24090	27010	32244	37831	44149	52080	61328	65287	
Number of patients	447078	464983	479847	526687	603244	675666	760724	893027	941755	
The rate of colonoscopic polypectomies										
Age category (yr)	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011
< 50	0.179	0.159	0.183	0.213	0.242	0.277	0.304	0.315	0.341	0.349
50-75	0.230	0.217	0.243	0.268	0.293	0.313	0.333	0.342	0.356	0.364
≥ 76	0.197	0.193	0.206	0.228	0.259	0.271	0.296	0.304	0.323	0.347
Age category (yr)	2012	2013	2014	2015	2016	2017	2018	2019	2020	
< 50	0.361	0.371	0.377	0.381	0.384	0.398	0.408	0.420	0.433	
50-75	0.376	0.382	0.386	0.388	0.390	0.400	0.407	0.416	0.426	
≥ 76	0.363	0.372	0.378	0.383	0.383	0.391	0.396	0.405	0.413	

## DISCUSSION

Using a nationwide, population-based database, we analyzed changes in the incidence of CRC according to age and the incidence rate of preinvasive CRC with endoscopic resection in Korea over a 19-year period. We found that the rate of endoscopically resectable preinvasive CRC continued to increase over time, with the 50-75 years age group having the highest proportion for both men and women. In addition, the rate of incidence of preinvasive CRC increased in a similar pattern in patients under 50 and 50-75 years of age. The incidence of preinvasive CRC showed an increased pattern similar to the increase in the number of colonoscopies, showing a gradual increase from 2005 to 2009, followed by a sharp increase after 2010. Unlike the incidence of preinvasive CRC, which steadily increased during the 17-year observation period, the incidence of CRC showed a slight decrease in the under 50 and 50-75 years age groups around the middle of the observation period. This suggested that the decrease in the incidence of CRC in the under 76 years age group might



**Figure 1** The incidence of colorectal cancer and preinvasive colorectal cancer by age and gender between 2004 and 2020. A: Colorectal cancer; B: Preinvasive colorectal cancer.



**Figure 2** The proportion of preinvasive colorectal cancer among all colorectal cancers by age between 2004 and 2020.

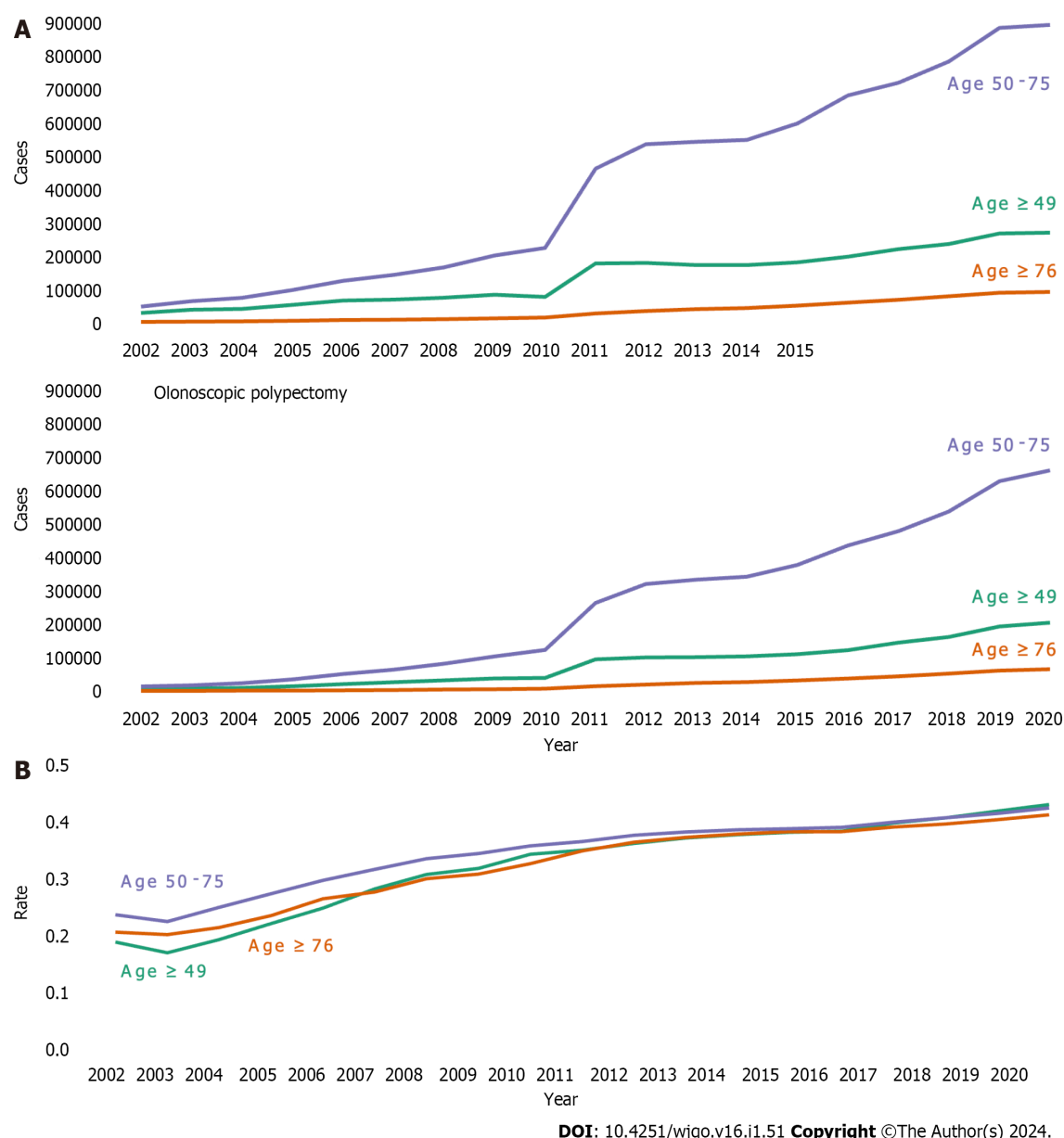
be related to the diagnosis, treatment, and subsequent periodic follow-up colonoscopy of preinvasive CRC in the younger age group. The increase in the incidence of CRC in patients aged 76 years or older was associated with an increase in the number of colonoscopies. The CRC screening guideline published in Korea in 2015 recommends starting CRC screening from age 50[11]. The total number of colonoscopies performed during the 19-year period showed a steady increase. In particular, it showed a sharp increase from 2010 to 2011. In addition to the explosive increase in colonoscopy performed for those in the 50-75 age group, there was a significant increase in the younger under 50 years age group. The new American Cancer Society guideline published in 2018 recommends extending the age of CRC screening colonoscopy to 45 years of age[12]. The trend in Korea for the past 17 years from 2004 to 2020 has shown a steady increase in CRC and preinvasive CRC diagnosis due to the increase in the number of colonoscopies and colonoscopic polypectomies in all age groups. In particular, the rate of colonoscopic polypectomies increased in those under 50 years of age. The rate of preinvasive CRC diagnosis in those under the age of 50 increased similar to that in those aged 50-75 years, the age group recommended for CRC screening. Therefore, these findings provide real-world evidence that colonoscopic polypectomy, especially preinvasive CRC resection, affects the occurrence of CRC. Several studies and guidelines have discussed extending the age of CRC screening from 45 to 75 years of age[4,13]. Previous studies have been conducted on how to determine the upper age limit for diagnostic colonoscopy in the elderly over 70 years of age[14-16]. Most studies recommended that the start and end time of CRC screening colonoscopy should be determined through a personalized discussion in consideration of the patient's underlying comorbidities and functional status as well as the patient's preferences[16,17]. In most countries, colonoscopy is mainly performed between the ages of 50 and 75[4,10,18]. Similarly, when subjects were classified into three age groups in this study, it was found that colonoscopy was mainly performed between 50 and 75 years of age. In this study, the frequency of colonoscopies and colonoscopic polypectomies increased in all age groups from 2002 to 2020. In particular, the number of colonoscopies increased in the elderly aged 76 years or older. We analyzed whether patients who underwent a colonoscopy from January 2002 to December 2020 were assigned a diagnostic code for colonoscopy-related complications[19,20], and determined hospitalizations and deaths that occurred within 1 mo and 3 mo after colonoscopy. As expected, the rate of hospitalization and death after colonoscopy in the elderly was high. The rate of complications and deaths after colonoscopy within 3 mo was high for those aged 76 years or older. This trend shows that older age and their comorbidities could increase the risk of these complications.

Rapid population aging in Korea is considered a major factor in the increased number of colonoscopies and colorectal polypectomies performed in the elderly[21]. We also found that the diagnosis of CRC and preinvasive CRC increased in the elderly aged 76 years or older with a high rate of complications after colonoscopy in these patients. Therefore, when endoscopists perform colonoscopy, they should consider each patient's individual information, rather than determining an upper age limit for screening colonoscopy.

This study has several limitations. The study population was obtained from the NHIS database, a database for the reimbursement of medical claims. In this study, CRC and preinvasive CRC were identified through ICD codes and the distribution level was confirmed by gender and age. In Korea, statistics on cancer patients are periodically checked through the national cancer registration statistics project. Data quality issues might exist. However, when data were compared by year, there was no significant difference. The trend in CRC was confirmed to be similar. In this study, NHIS data were used for further analysis. In addition, we were unable to assess several potential confounding factors[22], such as clinical findings, laboratory data, lifestyle or personal health behaviors (including smoking and alcohol intake) and physiological characteristics (such as body mass index). However, we minimized selection bias and referral bias by using large-scale administrative data obtained for the majority of the Korean population.

## CONCLUSION

Screening colonoscopy seems to have an important effect on the prevention of advanced CRC in the younger under 50 years age group as well as in the 50-75 year age group. Therefore, the role of colonoscopy as a screening test for CRC is very important even for those under the age of 50. As expected, the risk of colonoscopy-related hospitalization and death



**Figure 3** The total number of colonoscopies and colonoscopic polypectomies, and the proportion of colonoscopic polypectomies among the total number of colonoscopies and colonoscopic polypectomies by age, between 2002 and 2020. A: The total number of colonoscopies and colonoscopic polypectomies; B: The proportion of colonoscopic polypectomies among the total number of colonoscopies and colonoscopic polypectomies.

was high in the elderly. This suggested that if lesions at risk of developing CRC are diagnosed early and treated in the age group under 75 years of age, complications related to colonoscopy can be reduced in those over 75 years of age. In addition, before performing endoscopy for elderly patients, physicians should have sufficient consultation based on the patient's personalized information and make a decision considering the risk and benefit.

## ARTICLE HIGHLIGHTS

### Research background

In Korea, colorectal cancer (CRC) showed the second highest incidence rate among all cancers in 2018. The volume of colonoscopy and colorectal polypectomy in Korea is also increasing due to the CRC screening program initiated in 2004 and rapid population aging.

### Research motivation

We examined whether evaluating the trend in the incidence of CRC and preinvasive CRC by age group and gender would be helpful in determining the age group that is indicated for screening endoscopy in the future.

### Research objectives

This study aimed to evaluate the trend in incidence of CRC and preinvasive CRC according to the increase in colonoscopies performed in Korea.

### Research methods

Korean patients from 2002 to 2020 were identified to evaluate the incidence of CRC and preinvasive CRC and assess the number of diagnostic colonoscopies and colonoscopic polypectomies. Colonoscopy-related complications by age group were also determined.

### Research results

Over the study period, the rate of incidence of preinvasive CRC increased at a similar level for patients under 50 and 50-75 years of age. Since 2009, the increase in the rate of incidence has been rapid, showing a pattern similar to the increase in the number of colonoscopies.

### Research conclusions

The diagnosis of preinvasive CRC has increased with the increase in the number of colonoscopies performed. The role of screening colonoscopy for CRC has become very important even for those under the age of 50 years.

### Research perspectives

As the risk of colonoscopy-related complications is high in the elderly, if early lesions at risk of developing CRC are diagnosed and treated in the 75 years or under age group, complications related to colonoscopy can be reduced for those over 75 years of age.

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## FOOTNOTES

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**Co-first authors:** Ga Hee Kim and Yeong Chan Lee.

**Author contributions:** Kim TJ, Hong SN, Chang DK, Kim YH, Yang DH, Moon CM, Kim K, Kim HG, and Kim ER performed critical revision of the manuscript for important intellectual content; Kim GH and Lee YC acquired data from the National Health Insurance Service claims database and analyzed the data; Kim GH and Lee YC interpreted the data; Kim GH and Lee YC participated in the design of the study and performed the statistical analysis; the results shown in tables and figures were performed by Kim GH and Lee YC. The main text of the manuscript was written by Kim GH and Lee YC. Kim GH and Lee YC reviewed and edited the manuscript. Kim GH and Lee YC contributed equally to this work as co-first authors of this paper. Kim ER and Kim HG contributed the study concept and design. All authors have read and approved the final manuscript.

**Institutional review board statement:** The study was reviewed and approved for publication by our Institutional Reviewer.

**Informed consent statement:** In this retrospective cohort study, patients were not required to give informed consent to the study as the analysis used the National Health Insurance Service claims database which contains anonymous clinical data and is deidentified with regard to gender, age, and medical history.

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## REFERENCES

- 1 **Bray F**, Ferlay J, Soerjomataram I, Siegel RL, Torre LA, Jemal A. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. *CA Cancer J Clin* 2018; **68**: 394-424 [PMID: 30207593 DOI: 10.3322/caac.21492]
- 2 **Hong S**, Won YJ, Park YR, Jung KW, Kong HJ, Lee ES; Community of Population-Based Regional Cancer Registries. Cancer Statistics in Korea: Incidence, Mortality, Survival, and Prevalence in 2017. *Cancer Res Treat* 2020; **52**: 335-350 [PMID: 32178489 DOI: 10.4143/crt.2020.206]
- 3 **Siegel RL**, Miller KD, Jemal A. Cancer statistics, 2020. *CA Cancer J Clin* 2020; **70**: 7-30 [PMID: 31912902 DOI: 10.3322/caac.21590]
- 4 **Qaseem A**, Crandall CJ, Mustafa RA, Hicks LA, Wilt TJ; Clinical Guidelines Committee of the American College of Physicians, Forciea MA, Fitterman N, Horwitch CA, Kansagara D, Maroto M, McLean RM, Roa J, Tufte J. Screening for Colorectal Cancer in Asymptomatic Average-Risk Adults: A Guidance Statement From the American College of Physicians. *Ann Intern Med* 2019; **171**: 643-654 [PMID: 31683290 DOI: 10.7326/M19-0642]
- 5 **US Preventive Services Task Force**, Bibbins-Domingo K, Grossman DC, Curry SJ, Davidson KW, Epling JW Jr, García FAR, Gillman MW, Harper DM, Kemper AR, Krist AH, Kurth AE, Landefeld CS, Mangione CM, Owens DK, Phillips WR, Phipps MG, Pignone MP, Siu AL. Screening for Colorectal Cancer: US Preventive Services Task Force Recommendation Statement. *JAMA* 2016; **315**: 2564-2575 [PMID: 27304597 DOI: 10.1001/jama.2016.5989]
- 6 **Siegel RL**, Fedewa SA, Anderson WF, Miller KD, Ma J, Rosenberg PS, Jemal A. Colorectal Cancer Incidence Patterns in the United States, 1974-2013. *J Natl Cancer Inst* 2017; **109** [PMID: 28376186 DOI: 10.1093/jnci/djw322]
- 7 **Choi JH**, Cha JM, Yoon JY, Kwak MS, Jeon JW, Shin HP. The current capacity and quality of colonoscopy in Korea. *Intest Res* 2019; **17**: 119-126 [PMID: 30301340 DOI: 10.5217/ir.2018.00060]
- 8 **Tanaka S**, Kashida H, Saito Y, Yahagi N, Yamano H, Saito S, Hisabe T, Yao T, Watanabe M, Yoshida M, Kudo SE, Tsuruta O, Sugihara KI, Watanabe T, Saitoh Y, Igarashi M, Toyonaga T, Ajioka Y, Ichinose M, Matsui T, Sugita A, Sugano K, Fujimoto K, Tajiri H. JGES guidelines for colorectal endoscopic submucosal dissection/endoscopic mucosal resection. *Dig Endosc* 2015; **27**: 417-434 [PMID: 25652022 DOI: 10.1111/den.12456]
- 9 **Pimentel-Nunes P**, Dinis-Ribeiro M, Ponchon T, Repici A, Vieth M, De Ceglie A, Amato A, Berr F, Bhandari P, Bialek A, Conio M, Haringsma J, Langner C, Meisner S, Messmann H, Morino M, Neuhaus H, Piesseaux H, Rugge M, Saunders BP, Robaszkiewicz M, Seewald S, Kashin S, Dumoncean JM, Hassan C, Deprez PH. Endoscopic submucosal dissection: European Society of Gastrointestinal Endoscopy (ESGE) Guideline. *Endoscopy* 2015; **47**: 829-854 [PMID: 26317585 DOI: 10.1055/s-0034-1392882]
- 10 **Lee BI**, Hong SP, Kim SE, Kim SH, Kim HS, Hong SN, Yang DH, Shin SJ, Lee SH, Park DI, Kim YH, Kim HJ, Yang SK, Jeon HJ; Multi-Society Task Force for Development of Guidelines for Colorectal Polyp Screening, Surveillance and Management. Korean guidelines for colorectal cancer screening and polyp detection. *Clin Endosc* 2012; **45**: 25-43 [PMID: 22741131 DOI: 10.5946/ce.2012.45.1.25]
- 11 **Sohn DK**, Kim MJ, Park Y, Suh M, Shin A, Lee YH, Im JP, Cho HM, Hong SP, Kim BH, Kim Y, Kim JW, Kim HS, Nam CM, Park DI, Um JW, Oh SN, Lim HS, Chang HJ, Hahm SK, Chung JH, Kim SY, Lee WC, Jeong SY. The Korean guideline for colorectal cancer screening. *J Korean Med Assoc* 2015; **58**: 420-432 [DOI: 10.5124/jkma.2015.58.5.420]
- 12 **Wolf AMD**, Fontham ETH, Church TR, Flowers CR, Guerra CE, LaMonte SJ, Etzioni R, McKenna MT, Oeffinger KC, Shih YT, Walter LC, Andrews KS, Brawley OW, Brooks D, Fedewa SA, Manassaram-Baptiste D, Siegel RL, Wender RC, Smith RA. Colorectal cancer screening for average-risk adults: 2018 guideline update from the American Cancer Society. *CA Cancer J Clin* 2018; **68**: 250-281 [PMID: 29846947 DOI: 10.3322/caac.21457]
- 13 **Rex DK**, Boland CR, Dominitz JA, Giardiello FM, Johnson DA, Kaltenbach T, Levin TR, Lieberman D, Robertson DJ. Colorectal Cancer Screening: Recommendations for Physicians and Patients from the U.S. Multi-Society Task Force on Colorectal Cancer. *Am J Gastroenterol* 2017; **112**: 1016-1030 [PMID: 28555630 DOI: 10.1038/ajg.2017.174]
- 14 **Lin OS**. Performing colonoscopy in elderly and very elderly patients: Risks, costs and benefits. *World J Gastrointest Endosc* 2014; **6**: 220-226 [PMID: 24932373 DOI: 10.4253/wjge.v6.i6.220]
- 15 **García-Albéniz X**, Hsu J, Brethauer M, Hernán MA. Effectiveness of Screening Colonoscopy to Prevent Colorectal Cancer Among Medicare Beneficiaries Aged 70 to 79 Years: A Prospective Observational Study. *Ann Intern Med* 2017; **166**: 18-26 [PMID: 27669524 DOI: 10.7326/M16-0758]
- 16 **Nee J**, Chippendale RZ, Feuerstein JD. Screening for Colon Cancer in Older Adults: Risks, Benefits, and When to Stop. *Mayo Clin Proc* 2020; **95**: 184-196 [PMID: 31902414 DOI: 10.1016/j.mayocp.2019.02.021]
- 17 **Bénard F**, Barkun AN, Martel M, von Renteln D. Systematic review of colorectal cancer screening guidelines for average-risk adults: Summarizing the current global recommendations. *World J Gastroenterol* 2018; **24**: 124-138 [PMID: 29358889 DOI: 10.3748/wjg.v24.i1.124]
- 18 **Audibert C**, Perlaky A, Glass D. Global perspective on colonoscopy use for colorectal cancer screening: A multi-country survey of practicing colonoscopists. *Contemp Clin Trials Commun* 2017; **7**: 116-121 [PMID: 29696175 DOI: 10.1016/j.conctc.2017.06.008]
- 19 **Reumkens A**, Rondagh EJ, Bakker CM, Winkens B, Masclee AA, Sanduleanu S. Post-Colonoscopy Complications: A Systematic Review, Time Trends, and Meta-Analysis of Population-Based Studies. *Am J Gastroenterol* 2016; **111**: 1092-1101 [PMID: 27296945 DOI: 10.1038/ajg.2016.234]
- 20 **Kim SY**, Kim HS, Park HJ. Adverse events related to colonoscopy: Global trends and future challenges. *World J Gastroenterol* 2019; **25**: 190-204 [PMID: 30670909 DOI: 10.3748/wjg.v25.i2.190]
- 21 **Hyun KR**, Kang S, Lee S. Population Aging and Healthcare Expenditure in Korea. *Health Econ* 2016; **25**: 1239-1251 [PMID: 26085120 DOI: 10.1002/hec.3209]
- 22 **Wong MC**, Ding H, Wang J, Chan PS, Huang J. Prevalence and risk factors of colorectal cancer in Asia. *Intest Res* 2019; **17**: 317-329 [PMID: 31085968 DOI: 10.5217/ir.2019.00021]





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