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Editorial Board Member of *World Journal of Gastroenterology*, Dirk Jacob van Leeuwen, MD, PhD, FAASLD, Adjunct Professor, Section of Gastroenterology and Hepatology, Geisel School of Medicine at Dartmouth College, Hanover, NH 03756, United States. [dirk.j.vanleeuwen@gmail.com](mailto:dirk.j.vanleeuwen@gmail.com)

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## Retrospective Cohort Study

# Supply and quality of colonoscopy according to the characteristics of gastroenterologists in the French population-based colorectal-cancer screening program

Akoï Koïvogui, Catherine Vincelet, Gaëlle Abihsera, Hamou Ait-Hadad, Hélène Delattre, Tu Le Trung, Agnès Bernoux, Rachel Carroll, Jérôme Nicolet

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**Akoï Koïvogui**, Site de Seine-Saint-Denis, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Bondy 93146, France

**Catherine Vincelet**, Site des Yvelines, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Le Chesnay Cedex 78153, France

**Gaëlle Abihsera, Rachel Carroll**, Site du Val-de-Marne, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Joinville-le-Pont 94340, France

**Hamou Ait-Hadad**, Site de Seine-et-Marne, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Lieusaint 77763, France

**Hélène Delattre**, Site des Hauts-de-Seine, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Nanterre 92000, France

**Tu Le Trung**, site du Val-d'Oise, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Cergy Saint-Christophe 95800, France

**Agnès Bernoux**, Site de l'Essonne, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Fontenay-Les-Briis 91640, France

**Jérôme Nicolet**, Siège Paris, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), Paris 75015, France

**Corresponding author:** Akoï Koïvogui, MD, MHSc, MSc, Doctor, Site de Seine-Saint-Denis, Centre Régional de Coordination des Dépistages des Cancers en Ile-de-France (CRCDC-IDF), 41 Avenue de Verdun, Bondy 93146, France. [aakoivogui@live.fr](mailto:aakoivogui@live.fr)

## Abstract

### BACKGROUND

Since its complete roll-out in 2009, the French colorectal cancer screening program (CRCSP) experienced 3 major constraints [use of a less efficient Guaiac-test (gFOBT), stopping the supply of Fecal-Immunochemical-Test kits (FIT), and suspension of the program due to the coronavirus disease 2019 (COVID-19)] affecting its effectiveness.

**AIM**

To describe the impact of the constraints in terms of changes in the quality of screening-colonoscopy (Quali-Colo).

**METHODS**

This retrospective cohort study included screening-colonoscopy performed by gastroenterologists between Jan-2010 and Dec-2020 in people aged 50-74 living in Ile-de-France (France). The changes in Quali-Colo (Proportion of colonoscopies performed beyond 7 mo (Colo\_7 mo), Frequency of serious adverse events (SAE) and Colonoscopy detection rate) were described in a cohort of Gastroenterologists who performed at least one colonoscopy over each of the four periods defined according to the chronology of the constraints [gFOBT: Normal progress of the CRCSP using gFOBT (2010-2014); FIT: Normal progress of the CRCSP using FIT (2015-2018); STOP-FIT: Year (2019) during which the CRCSP experienced the cessation of the supply of test kits; COVID: Program suspension due to the COVID-19 health crisis (2020)]. The link between each dependent variable (Colo\_7 mo; SAE occurrence, neoplasm detection rate) and the predictive factors was analyzed in a two-level multivariate hierarchical model.

**RESULTS**

The 533 gastroenterologists (cohort) achieved 21509 screening colonoscopies over gFOBT period, 38352 over FIT, 7342 over STOP-FIT and 7995 over COVID period. The frequency of SAE did not change between periods (gFOBT: 0.3%; FIT: 0.3%; STOP-FIT: 0.3%; and COVID: 0.2%;  $P = 0.10$ ). The risk of Colo\_7 mo doubled between FIT [adjusted odds ratio (aOR): 1.2 (1.1; 1.2)] and STOP-FIT [aOR: 2.4 (2.1; 2.6)]; then, decreased by 40% between STOP-FIT and COVID [aOR: 2.0 (1.8; 2.2)]. Regardless of the period, this Colo\_7 mo's risk was twice as high for screening colonoscopy performed in a public hospital [aOR: 2.1 (1.3; 3.6)] compared to screening-colonoscopy performed in a private clinic. The neoplasm detection, which increased by 60% between gFOBT and FIT [aOR: 1.6 (1.5; 1.7)], decreased by 40% between FIT and COVID [aOR: 1.1 (1.0; 1.3)].

**CONCLUSION**

The constraints likely affected the time-to-colonoscopy as well as the colonoscopy detection rate without impacting the SAE's occurrence, highlighting the need for a respectable reference time-to-colonoscopy in CRCSP.

**Key Words:** Colorectal cancer screening; Screening colonoscopy; Faecal immunochemical test; Guaiac faecal occult blood test; Quality of colonoscopy; Severity of tumor lesions

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**Core Tip:** The study showed that the detection rate of colonoscopy dropped significantly in France during the years 2019 and 2020, probably due to the coronavirus disease health crisis. The risk of a long delay (> 7 mo) in performing the colonoscopy was twice as high in a public hospital compared to colonoscopies performed in a private endoscopy practice. The constraints likely affected the time to colonoscopy as well as the colonoscopy detection rate without impacting the occurrence of serious adverse events.

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**INTRODUCTION**

The impact of the Screening program on controlling colorectal cancer (CRC) morbidity and mortality has been widely proved[1-4]. But since its complete roll-out in France in 2009, the population-based colorectal cancer screening program (CRCSP) has continued to face constraints affecting its effectiveness. Despite the existence of the fecal immunochemical test (FIT) in certain European programs (*i.e.*, Italy, Czech Republic) when the program roll-out was completed in France[5], the health authority chose the Guaiac Hemoccult II test® (gFOBT). It later turned out that gFOBT only identified 50% of colorectal cancer (CRC) lesions and a third of adenomas[6], which led some GPs to be wary of it, at the

risk of seeing some of their patients fall through the cracks[6,7].

To consider this first constraint induced using a low sensitivity/specificity screening test, the health authority decided to replace gFOBT in 2015, with the FIT (Threshold set at 150 ng hemoglobin/mL of stool, “Institut National du Cancer”, [www.e-cancer.fr](http://www.e-cancer.fr)). While admitting an improvement in participation with FIT compared to gFOBT, most studies published in France have confirmed the high sensitivity (detection of advanced adenomas and CRC) of FIT and its better acceptability by the population and GPs[8-12]. This performance of the FIT inevitably leads to an increase in colonoscopy requests in the screened population and subsequently to an extension of the time to colonoscopy after a positive FIT result[13]. However, these analyses of the time to colonoscopy only considered the characteristics of the target population without any adjustment to the characteristics of the colonoscopy supply.

On April 25, 2018, the Paris Administrative Court cancelled, during an appeal session, the contract concluded in 2014 between the Health Insurance Agency and the Cerba-Daklapack® consortium ([www.slbc.fr](http://www.slbc.fr)). This contract, which related to the supply of screening test kits and the laboratory analysis of the tests carried out, had thus been cancelled only three years after the introduction of the FIT in CRCSP. This legal and administrative confusion led to a market shutdown between March and September 2019. In the Ile-de-France (IDF) region, this shutdown led to a drastic decrease in the number of tests carried out in 2019, compared to forecasts (annual activity report 2019).

Only a few months after the resumption of the test kits’ market, the World Health Organization (WHO) announced the pandemic of COVID-19[14]. This pandemic constraint required a relocation of health care resources to control this global health crisis. Screening programs, in particular the CRCSP, were suspended in many countries. The aim of this study was to describe the impact of the constraints listed above in terms of changes to the quality of screening colonoscopies (Quali-colo) in a cohort of gastroenterologists (GEs) practicing in IDF.

## MATERIALS AND METHODS

This retrospective cohort study included all screening colonoscopies, performed between 01/01/2010 and 31/12/2020 by GEs in the IDF region and collected by the eight sites (Paris, Seine-et-Marne, Yvelines, Essonne, Hauts-de-Seine, Seine-Saint-Denis, Val-de-Marne and Val-d’Oise) of the IDF CRCSP Coordination Centre (CRCDC-IDF). These screening colonoscopies were performed following a positive screening test in people aged 50-74, living in IDF, France.

Considering the chronology of the constraints in the CRCSP, four periods for carrying out the colonoscopy were distinguished (Figure 1). The first period (gFOBT) corresponded to the five years (2010-2014) of normal progress of the CRCSP using gFOBT. The second period (FIT) corresponded to the four years (2015-2018) of normal progress of the CRCSP using FIT. The third (FIT-STOP) corresponded to the year (2019) during which the CRCSP experienced the cessation of the supply of test kits and the fourth (COVID) corresponded to the program suspension due to the COVID-19 health crisis (2020).

The supply of screening colonoscopy was described by the number and type of practice of GEs practicing in IDF and having performed a screening colonoscopy in a person living in IDF. The Quali-colo was described in terms of time to colonoscopy, yield of colonoscopy and frequency of undesirable events (incidents/accidents, incomplete colonoscopy, refusal of 2<sup>nd</sup> colonoscopy).

Descriptive and evolutive analyses (supply and Quali-colo) were carried out between the periods (gFOBT, FIT, FIT-STOP, and COVID). These changes were first described according to the characteristics of the GEs who performed the screening colonoscopies. Secondly, the impact of constraints was described in terms of changes in Quali-colo indicators between the four periods, in a cohort of GEs (Cohort-GE) who performed at least one colonoscopy in each of the four periods.

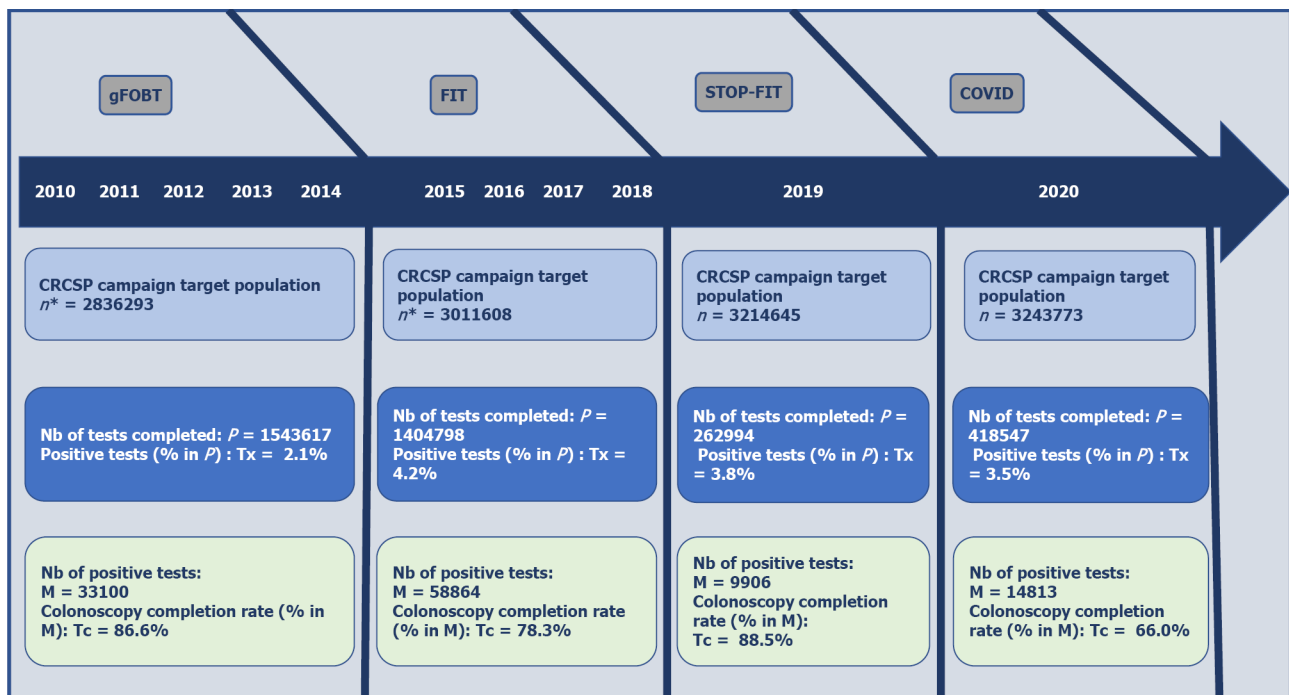
### Screening organization and study data collection

The National Council of the Order of Physicians (Research and Statistics Study Department) provided the medical demographic data. Screening data were extracted from CRCDC-IDF departmental databases. Over the study period, the CRCSP campaigns were organized following the CRCSP specifications[15,16]. As a preliminary to each campaign in each study department, an update of the files of eligible people was made after the transmission of individual data by the partners (Health Insurance plans, Medical Information Services of hospitals, Pathologists, GEs, Surgeons, GPs, patients). Anyone who had a screening test did not need a screening colonoscopy if the test result was negative. In case of a positive test result, the person was subsequently invited five years after a normal colonoscopy or excluded from the CRCSP after a positive colonoscopy result (polyp or CRC).

### Definition of variables

The screening colonoscopy (complete or incomplete) was considered completed only if the result was provided with or without a completion date. When the completion date was provided, the time to screening colonoscopy was expressed as the number of months between the date of completion of the screening test and the date of completion of the colonoscopy. In the cases where several colonoscopies





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**Figure 1** Evolution of the colorectal cancer screening program indicators (target population of the campaigns, number of tests carried out, number and proportion of positive tests, rate of completion of colonoscopy) over the 4 study periods (guaiac fecal occult blood test, fecal immunochemical test, STOP-fecal immunochemical test, and COVID). The asterisk (\*) is the target population at the start of the period. The colonoscopy completion rate was estimated based on data extraction as of January 31, 2022. gFOBT: Guaiac fecal occult blood test; COVID: Coronavirus disease; FIT: Fecal immunochemical test; Nb: Number; CRCSP: Colorectal cancer screening program.

were carried out to investigate the same positive test, the time to screening colonoscopy was that related to the first colonoscopy. The proportion of screening colonoscopies with an abnormally long time to access colonoscopy (Long-delay-colo) was estimated by the frequency of colonoscopies performed beyond a 7-mo delay among the screening colonoscopies for which the completion date was provided. This delay threshold considers the fact that the risk of colorectal cancer is increased by about 40% for any colonoscopy performed after a waiting period of 7-12 mo[17].

The screening colonoscopy was complete when the colon was examined until crossing the Bauhin valve. The reasons for an incomplete colonoscopy were: Insufficient preparation, Anatomical (dolichocolon, Presence of an obstructive lesion requiring a second colonoscopy or surgery). The accidents related to screening colonoscopy were: exterior hemorrhage with or without transfusion, perforation, death. Incidents related to anesthesia or general condition (cardiorespiratory disorders) were distinguished from those related to endoscopy (*i.e.*, difficulty crossing a cul-de-sac, placement of clips to stop bleeding after a polypectomy). The proportion of serious adverse events (SAEs) was estimated by the frequency of screening colonoscopies during which an incident/accident was notified.

The screening colonoscopy was classified as positive when a neoplasm (Polyp/adenoma/CRC) was discovered, negative if not. The screening colonoscopy detection rate (yield of colonoscopy) was estimated by the proportion of positive colonoscopies among the screening colonoscopies performed. The CRC and polyps/adenomas diagnoses were those coded C18-C20 and D12 according to the 10<sup>th</sup> version of the WHO International Classification of Diseases (ICD10)[18]. The CRC was considered "seen at colonoscopy" when an ulcerative-budding/ulcerative-necrotizing lesion was described by the GE. The high-risk polyps were adenomatous or scalloped polyps with a diameter of  $\geq 10$  mm (except hyperplastic polyps), high-grade dysplasia adenomas, villous or tubulo-villous adenomas. The TNM classification[19] has been used to define CRC severity. Any CRC  $\geq T3$  (subserous invaded) or  $\geq N1$  (at least one regional node invaded) or M1 (with metastasis) was considered severe CRC.

For each GE practicing in IDF region, having performed at least one screening colonoscopy, the factors studied were: (1) The existence of a gastroenterology consultation carried out before the screening colonoscopy completion date; (2) the annual number of screening colonoscopies performed (1, 2-30, 31-100, and  $> 100$  colonoscopies); (3) the place of performance of the screening colonoscopy (1-Private clinic in the IDF; 2-Private hospitals in the IDF; 3-Public hospital in the IDF including: The Public Assistance of Paris hospitals -APHP-, Other public hospitals in the IDF including army hospitals and municipal health centers). The colonoscopies performed by GEs practicing in  $\geq 2$  locations, the locations of which had not been specified ( $n = 2$ ), were attributed to the locations most frequented by these GEs over the period. Similarly, Colo for which the location was specified but for which the GEs were not

specified ( $n = 6$ ), were attributed to the GEs who performed the greatest number of colonoscopies on the location and over the period. Colo performed in a country other than France were classified as “Place Unspecified”. Colonoscopies performed in another region of France were classified “Outside-IDF”; (4) the annual number of colonoscopy locations (1 Location,  $\geq 2$  Locations); (5) the density of GEs in the municipality where the GE performed the screening colonoscopy. The density (D) of GEs was estimated as number of GEs/100000 inhabitants. Each colonoscopy year, with reference to a regional average density (M) and standard deviation (SD). Low density of GE was:  $D < M - SD$ , average-density of GE was:  $D$  in  $M \pm SD$ , high density of GE was:  $D > M + SD$ ; (6) the seniority of the GE (for any year “A”, the GE having no screening colonoscopy in the years prior to “A” was considered a new GE); (7) the residence of the CRCSP target patient treated by the GE (1-the Colonoscopy’s supply municipality, 2-other municipality in the Colonoscopy’s supply department, 3-other IDF departments). As a reminder, in 2018, The National Institute of Statistics and Economic Studies (INSEE) counted 1267 municipalities in IDF in addition to the city of Paris; and (8) the age of the CRCSP target patient treated by the GE (50-54, 55-59, 60-64, 65-69, and  $\geq 70$  years).

### Statistical analysis

The proportions (Colo performed within one month or after a waiting delay  $> 7$  mo, incomplete and redone Colo, incidents/accidents, positive Colo, high\_risk\_polyp, CRC seen at Colo, CRC with provided status, severe CRC) were described and compared between periods (gFOBT, FIT, FIT-STOP, and COVID) by the Pearson’s Chi-2 test. In the strata defined according to the characteristics of the cohort-GE, the time to perform the screening colonoscopy (in months) was analyzed in terms of average and confidence interval (CI) then, an analysis of variance (ANOVA on repeated measures) was used to compare the average delays between periods (gFOBT, FIT, FIT-STOP *vs* COVID). In the strata defined according to the characteristics of the cohort-GE, the proportions (colonoscopies performed after  $> 7$  mo delay, proportion of SAEs, yield of screening colonoscopy) were compared between periods (gFOBT, FIT, FIT-STOP *vs* COVID) by Cochran’s Q test.

The link between each dependent variable (binary variables 0/1: Long-delay-colo; SAEs, Yield of screening colonoscopy) and the predictive factors (annual number of screening colonoscopies performed, Place of performance of the screening colonoscopy, Annual number of colonoscopy locations, Density of GE, Residence of the patient, Age of the patient) was analyzed in a multivariate and two level (colonoscopy and GE) hierarchical regression model. The generalized linear model (family: Bernoulli, link: Logit) with mixed effect was preferred. This multivariate analysis was performed using a model with all covariates regardless of their relationship in univariate analysis. In addition, a strong correlation existed between several covariates (*i.e.*, annual number of screening colonoscopies and Place of performance, Annual number of screening colonoscopies and Municipal density of GEs, Annual number of screening colonoscopies and Period), the model was extended to these terms of interaction between covariates. Only the significant interaction terms ( $P < 0.05$  in univariate analysis) were kept in the final model evaluated by the likelihood ratio test. A biomedical statistician performed the statistical review. All the analyses were carried out at the 5% threshold with version 13 of the STATA software (College Station, TX, United States).

### Regulatory issues

Before analysis, all data were anonymized. The screening database had a favourable opinion from the institution that oversees the ethics of data collection (“Commission nationale de l’informatique et des libertés”: CNIL)[20]. According to the current French legislation, a study that does not change the care of patients did not require the opinion of the Clinical Research Centre’s Ethics Committee.

## RESULTS

### Descriptive and evolutive analyses

Out of a total of 1267 municipalities listed in the IDF region, only 155 municipalities had at least one GE in 2010. This number of municipalities having at least one GE falling from 155 in 2010 to 142 in 2020. In the municipalities having at least one GE, the average annual density of GEs fluctuated between a minimum of 6.3 (in 2014) and a maximum of 6.5 GE/100000 inhabitants over the study period (Table 1).

The gap between the number of GEs registered in the medical demographic database and the number of GEs having performed at least one screening colonoscopy, increased from 134 in 2010 (761 registered *vs* 627 having performed  $\geq 1$  screening colonoscopy), to 206 in 2015 (776 *vs* 570) before being reduced to 123 in 2019 (798 *vs* 675). The proportion of GEs performing screening colonoscopies at two or more locations varied from 20.6% in 2010 to 13.9% in 2015, then 21.8% in 2019. The proportion of new GEs decreased from 12.6% in 2011 to 7.7% in 2015, then increased to 13.5% in 2016 and further decreased to 4.7% in 2019. In 2016, a total of 727 GEs performed at least one colonoscopy. Among them, 97 GE performed only one screening colonoscopy and 8 GEs exceeded an annual number of 100 screening colonoscopies (Table 1).



**Table 1 Evolution of the regional offer in number of gastroenterologists and the number of gastroenterologists having performed at least one colonoscopy, by year of performance of the screening colonoscopy**

Year of colonoscopy	Nb of GE in IDF <sup>1</sup>		Number of gastroenterologists who performed a screening colonoscopy <sup>2</sup>												Total (n) of GE in IDF (% GE ≥ 2 location)
			Number of GE by seniority		Number of GE by density of GE in the municipality of practice of the GE			Number of GE by place of performance of the colonoscopy			Number of GE by annual number (A) of colonoscopies performed				
	Nb of GE (density) <sup>3</sup>	Nb of municipalities with GE	Senior	New (% in n)	Low	Average	High	Private clinic	Private Hop.	Public Hop.	A = 1	A = 2- 30	A = 30- 100	A > 100	
2010	761 (6.5)	155	627	-	134	85	493	415	114	214	119	473	35	-	627 (20.6)
2011	756 (6.4)	156	534	77 (12.6)	140	71	474	408	117	201	106	465	40	-	611 (17.2)
2012	759 (6.4)	155	538	57 (9.6)	116	79	473	383	112	206	117	454	24	-	595 (16.8)
2013	761 (6.4)	154	539	30 (5.3)	98	92	442	378	115	181	107	448	14	-	569 (16.5)
2014	757 (6.3)	155	522	63 (10.8)	129	75	451	384	106	193	123	448	14	-	585 (17.4)
2015	776 (6.4)	154	526	44 (7.7)	117	53	449	379	103	178	140	419	11	-	570 (13.9)
2016	784 (6.5)	154	629	98 (13.5)	128	65	628	432	143	312	97	447	175	8	727 (18.8)
2017	793 (6.5)	152	642	72 (10.1)	142	56	603	418	142	312	93	486	134	1	714 (19.9)
2018	799 (6.5)	149	665	64 (8.8)	141	51	626	424	151	312	100	488	139	2	729 (20.7)
2019	798 (6.5)	147	643	32 (4.7)	123	63	574	388	152	287	92	512	71	-	675 (21.8)
2020	802 (6.5)	142	619	76 (10.9)	147	50	582	412	162	265	124	475	96	-	695 (19.7)

<sup>1</sup>Number of gastroenterologists (GE) registered in the region (source: National Council of the Order of Physicians).

<sup>2</sup>Number of gastroenterologists who performed a screening colonoscopy during the calendar year (regardless of the type of test and regardless of the date of the screening test).

<sup>3</sup>Density in Number of GE/100000 inhabitants: Regional average density (5.5 à 7.5 GE/100000 inhabitants) Low density of GE (< 5.5 GE/100000 habitants) and high density of GE (> 7.5 GE/100000 inhabitants).

GE: Gastroenterologist; Hop: Hospital; IDF: Ile-de-France; Nb: Number.

In 2011, out of a total of 6428 colonoscopies performed in IDF, the proportion of colonoscopies performed by new GEs was 2.0%, the proportion of colonoscopies performed in a municipality with a high density of GEs was 62.2%, the proportion of colonoscopies performed in a public hospital was 12.5%. In 2016, 1041 screening colonoscopies were performed by the GEs having an annual volume of > 100 screening colonoscopies and 9148 (58.9%) screening colonoscopies were performed by the GEs having an annual volume of 30-100 screening colonoscopies. Compared to 2010 (1.7%), the proportion of screening colonoscopies performed outside the IDF region was significantly higher in 2020 (2.5%;  $P < 0.0001$ ). Similarly, compared to 2019 (16.8%), the proportion of screening colonoscopies performed in public hospitals decreased significantly in 2020 (13.0%,  $P < 0.0001$ ) (Table 2).

**Table 2 Evolution of the number of colonoscopies performed according to the characteristics of the gastroenterologist, by year of performance of the screening colonoscopy, *n* (%)**

Year of colonoscopy	Number of colonoscopies performed according to GE characteristics														
	Number of colonoscopies by seniority of GE		Number of colonoscopies by density of GE in the municipality of practice of the GE <sup>1</sup>			Number of colonoscopies by place of performance of the colonoscopy			Number of colonoscopies by GE's annual number (A) of colonoscopies performed				Total		
	Senior	New	Low	Average	High	Clinic	Private Hop.	Public Hop.	A = 1	A = 2-30	A = 31-100	A > 100	Nb (n) of Colo performed in IDF (average Nb of Colo by GE)	Nb of Colo with place specified (% outside IDF)	Nb of Colo (% Place unspecified)
2010	6059	-	1535	900 (14.9)	3624 (59.8)	4507	830 (13.7)	722 (11.9)	119	4493 (74.2)	1447 (23.9)	-	6059 (11)	6161 (1.7)	6441 (4.4)
2011	6300	128 (2.0)	1684	712 (11.1)	4032 (62.7)	4677	946 (14.7)	805 (12.5)	106	4578 (71.2)	1744 (27.1)	-	6428 (12)	6543 (1.8)	6928 (5.6)
2012	5355	76 (1.4)	1186	766 (14.1)	3479 (64.1)	3818	830 (15.3)	783 (14.4)	117	4284 (78.9)	1030 (19.0)	-	5431 (11)	5533 (1.8)	5852 (5.5)
2013	4309	47 (1.1)	1045	737 (16.9)	2574 (59.1)	3156	660 (15.2)	540 (12.4)	107	3725 (85.5)	524 (12.0)	-	4356 (9)	4409 (1.2)	4712 (6.4)
2014	4320	132 (3.0)	1104	611 (13.7)	2737 (61.5)	3199	650 (14.6)	603 (13.5)	123	3718(83.5)	611 (13.7)	-	4452 (9)	4515 (1.4)	4746 (4.9)
2015	3712	63 (1.7)	879	446 (11.8)	2450 (64.9)	2692	604 (16.0)	479 (12.7)	140	3198 (84.7)	437 (11.6)	-	3775 (8)	3818 (1.1)	4034 (5.4)
2016	15196	333 (2.1)	3406	1862 (12.0)	10261 (66.1)	10886	2527 (16.3)	2116 (13.6)	97	5243 (33.8)	9148 (58.9)	1041	15529 (25)	15811 (1.8)	16651 (5.0)
2017	11519	192 (1.6)	2876	1262 (10.8)	7573 (64.7)	7919	1937 (16.5)	1855 (15.8)	93	5370 (45.9)	6137(52.4)	111	11711 (18)	11920 (1.8)	12345 (3.4)
2018	12181	164 (1.3)	2758	1190 (9.6)	8397 (68.0)	8233	2300 (18.6)	1812 (14.7)	100	5331 (43.2)	6684 (54.1)	230	12345 (19)	12602 (2.0)	13057 (3.5)
2019	8189	98 (1.2)	1582	932 (11.3)	5773 (69.7)	5365	1532 (18.5)	1390 (16.8)	92	5261 (63.5)	2934 (35.4)	-	8287 (13)	8487 (2.4)	8767 (3.2)
2020	9103	158 (1.7)	2088	755 (8.2)	6418 (69.3)	6654	1900 (20.5)	1199 (13.0)	124	5049 (54.5)	4088 (44.1)	-	9261 (15)	9501 (2.5)	9793 (3.0)

<sup>1</sup>Density in Number of GE/100000 inhabitants: Regional average density (5.5 à 7.5 GE/100000 inhabitants), low density of GE (< 5.5 GE/100000 habitants) et High density of GE (> 7.5 GE/100000 inhabitants).

Colo: Screening colonoscopy; GE: Gastroenterologist; Hop.: Hospital; IDF: Ile-de-France; Nb: Number.

Overall, the time to screening colonoscopy was significantly longer over STOP-FIT (gFOBT:  $2.6 \pm 2.9$  vs FIT:  $3.0 \pm 3.0$ ; STOP-FIT:  $3.9 \pm 3.9$ , COVID:  $3.5 \pm 3.9$ ,  $P < 0.0001$ ). Over the gFOBT period, 3.1% of the 28679 colonoscopies performed were incomplete (20.7% were redone) for reasons: Anatomical (60.6%), insufficient preparation (16.1%). The proportion of incomplete and redone colonoscopies was significantly higher over FIT ( $P < 0.001$ ). Although one case of death was reported during the gFOBT period, the proportion of adverse events was not significantly related to the period (0.05). The proportion of cancers seen at colonoscopy was lower over FIT (gFOBT: 61.4%, vs FIT: 55.2% or STOP-FIT: 57.5% or COVID: 56.1%;  $P < 0.0001$ ) (Table 3).

**Table 3** Quality indicators and results of colonoscopies by period of performance of colonoscopy in people aged 50-74, residing in Ile-de-France, *n* (%)

Quality indicator	Period				
	gFOBT	FIT	STOP-FIT	COVID	<i>P</i> value <sup>1</sup>
Total number ( <i>n</i> ) of colonoscopies	28679	46087	8767	9783	
Existence of a GE consultation before colonoscopy					
Nb (A) colonoscopies with date of consultation	5267 (18.4)	1517 (3.3)	406 (4.6)	198 (2.0)	< 10 <sup>-3</sup>
Date of consultation ≠ Date of colonoscopy					
Nb colonoscopies of which date of consultation ≠ colon date (% in A)	298.4 (56.7)	883 (58.2)	402 (99.0)	191 (96.5)	< 10 <sup>-3</sup>
Time to colonoscopy					
Average (in mean ± SD)	2.6 ± 2.9	3.0 ± 3.0	3.9 ± 3.9	3.5 ± 2.9	< 10 <sup>-3*</sup>
Number of colonoscopies performed within one month	4957 (17.3)	4572 (9.9)	458 (5.2)	726 (7.4)	< 10 <sup>-3</sup>
Number of colonoscopies performed beyond 7 mo	1520 (5.3)	2949 (6.4)	1034 (11.8)	933 (9.5)	< 10 <sup>-3</sup>
Complete colonoscopy					< 10 <sup>-3</sup>
Nb colonoscopies without information on performance	1263 (4.4)	2360 (5.1)	410 (4.7)	432 (4.4)	
Number of complete colonoscopies	26530 (92.5)	41695(90.5)	8004 (91.3)	8981 (91.8)	
Nb (B) of incomplete colonoscopies	886 (3.1)	2032 (4.4)	357 (4.1)	376 (3.8)	
Reasons for incomplete colonoscopies					< 10 <sup>-3</sup>
Unspecified: <i>n</i> (% in B)	206 (23.3)	617 (30.4)	109 (30.5)	114 (30.3)	
Anatomical reason/Obstruction by lesion: <i>n</i> (% in B)	537 (60.6)	845 (41.6)	150 (42.0)	161 (42.8)	
Insufficient preparation: <i>n</i> (% in B)	143 (16.1)	570 (28.1)	98 (27.5)	101 (26.9)	
Redone incomplete colonoscopy					
Number of redone colonoscopies (% B)	183 (20.7)	960 (47.2)	158 (44.3)	163 (43.3)	< 10 <sup>-3</sup>
Frequency of incidents					0.14
No incidents reported: <i>n</i>	28873 (99.6)	45947 (99.7)	8740 (99.7)	9763 (99.8)	
Related to anaesthesia/general condition: <i>n</i>	18 (0.06)	24 (0.05)	3 (0.03)	2 (0.02)	
Related to endoscopy: <i>n</i>	88 (0.3)	116 (0.3)	22 (0.3)	18 (0.2)	
Frequency of accidents					0.17
No accidents reported: <i>n</i>	28589 (99.7)	45970 (99.8)	8749 (99.8)	9763 (99.8)	
Suspected complication: <i>n</i>	24 (0.08)	23 (0.05)	5 (0.04)	3 (0.03)	
Exterior bleeding: <i>n</i>	57 (0.2)	66 (0.1)	14 (0.2)	12 (0.1)	
Perforation: <i>n</i>	8 (0.03)	28 (0.06)	3 (0.03)	4 (0.04)	
Deaths: <i>n</i>	1 (0.0)	0	0	0	
Colonoscopies results					
Detection rate: Nb of lesions	14857 (51.8)	29843 (64.8)	5565 (63.5)	5967 (60.1)	< 10 <sup>-3</sup>
Nb Polyps (% HRP)	12947 (44.2)	26624 (56.4)	5040 (53.3)	5425 (51.8)	< 10 <sup>-3</sup>
Nb of CRC (% CRC seen at colonoscopy)	1910 (61.4)	3219 (55.2)	525 (57.3)	542 (56.1)	< 10 <sup>-3</sup>
% CRC with severity stage specified among Nb CRC <sup>2</sup>	90.3	80.5	74.3	72.3	< 10 <sup>-3</sup>
Nb CRC with severity stage specified (% severe CRC) <sup>2</sup>	1724 (50.7)	2592 (40.9)	390 (39.5)	392 (39.5)	< 10 <sup>-3</sup>

<sup>1</sup>Pearson's  $\chi^2$ /Fisher's exact test of proportion or Fisher's *F* test (ANOVA).<sup>2</sup>Any CRC ≥ T3 (subserous invaded) or ≥ N1 (at least one regional node invaded) or M1 (with metastasis) was considered severe colorectal cancer.

CRC: Colorectal cancer; FIT: Fecal immunochemical test; GE: Gastroenterologist; gFOBT: Guaiac fecal occult blood test; HRP: High risk polyps (advanced adenoma); Nb: Number.

### Changes in Quali-colo indicators between the four periods, in a cohort of GEs

The cohort of 533 GE achieved 21509 Screening colonoscopies over the gFOBT period, 38352 over FIT, 7342 over STOP-FIT and 7995 over the COVID period. In this cohort, the difference in time (months) to screening colonoscopy between periods was globally significant [gFOBT: 2.6 (2.5; 2.6) *vs* FIT: 3.0 (2.9; 3.0); STOP-FIT: 3.9 (3.8; 4.0) and COVID: 3.5 (3.4; 3.6);  $P < 0.0001$ ]. The average time to colonoscopy was longer in public hospitals compared to clinics or private hospital, regardless of the period. This average time was paradoxically shorter over the COVID period compared to the STOP-FIT period, regardless of the type of establishment [in STOP-FIT clinic: 3.7 (3.6; 3.7) *vs* COVID: 3.4 (3.3-3.5) in public hospitals STOP-FIT: 5.1 (4.7-5.9) *vs* COVID: 4.2 (3.8; 4.7)]. The average time to colonoscopy was significantly lower among GEs practicing in low-density areas of GEs compared to those practicing in high-density areas of GEs, over the gFOBT and FIT periods, conversely, depending on the density area the confidence intervals were not significant over the STOP-FIT and COVID periods (Table 4).

Regardless of the GE's characteristics, the proportion of screening colonoscopy performed in  $> 7$  mo delay was significantly higher over STOP-FIT ( $P < 0.001$ ). The proportion of colonoscopies performed in  $> 7$  mo delay was higher in public hospitals compared to clinics and private hospitals, regardless of the period ( $P < 0.001$  in each period). This proportion of colonoscopies performed in  $> 7$  mo delay decreased during the COVID period compared to the STOP-FIT period, regardless of the place of colonoscopy ( $P < 0.001$  for each place). The proportion of colonoscopies performed in  $> 7$  mo delay was higher in the 50-54 age group, regardless of the period ( $P < 0.001$  in each period) (Table 5).

Whatever the characteristics of the Cohort-GE, the decline in colonoscopy detection rate was significant between the FIT and COVID period (Table 6). The risk of having a long delay to colonoscopy was twice as high for screening-colonoscopy performed in a public hospital [adjusted odds ratio (aOR): 2.1 (1.3; 3.6)] compared to screening colonoscopy performed in a private IDF clinic. Except for the patient's age, the risk of adverse events was not related to any other predictive factor. Compared to patients aged 50-54, patients aged 70 had a 70% increased risk of neoplasm detection. The risk of neoplasm detection decreased by about 40% between the periods FIT [aOR: 1.6 (1.5; 1.7)] and COVID [aOR: 1.1 (1.0; 1.3)] (Table 7).

## DISCUSSION

The European guide for quality assurance of colorectal cancer screening recommends performing a colonoscopy within 31 d following a positive test result[21]. In our Cohort-GE, if the increase in the time to screening colonoscopy between the first and the second period was attributable to the introduction of FIT, its increase after the second period was attributable to the malfunction of the program due to the slowdown of the kit market and the COVID-19 health crisis. There is certainly no relationship between the kit market and the colonoscopy offer, but the unexplained increase in the time to perform colonoscopy during a year that saw a market slowdown can be explained factually by this market crisis. The hypothesis would be that general practitioners reacted to the market crisis by relaxing the program, in particular the follow-up of people who had a positive test. Indeed, in France, in addition to the distribution of the test kit, the training doctors are real facilitators of access to colonoscopy (helping the patient to make an appointment with a gastroenterologist, motivating the patient to have the colonoscopy). This hypothesis is confirmed by the slight decrease in the time to colonoscopy in 2020 compared to 2019, despite the COVID-19 health crisis. The year 2020 was moreover affected by this kit market crisis than by the COVID-19 health crisis. Indeed, after the resumption of the kit market in September 2019, several people who had a positive test during the last quarter of 2019 were inevitably the first to be affected by colonoscopy postponements at the start of the first confinement in March 2020. However, the improvement in the time to colonoscopy during the pandemic (compared to the STOP-FIT period) could also be linked to the fact that people have refocused their concerns on their health. Regardless of the characteristics of the Cohort-GE, the screening colonoscopy detection rate dropped significantly between the STOP-FIT and COVID periods, while the proportions of SAEs stayed unchanged.

The long delay to access colonoscopy observed on the gFOBT and FIT periods converges with the results of another French study[22], although it is clearly higher than those observed elsewhere[23,24]. The definition of a reference delay and the obligation of compliance with it by all GEs taking part in CRCSP would effectively reduce the delay in France. This reframing is necessary, especially since the number of GEs is large, but with an increased disparity in terms of the number of screening colonoscopies performed by GEs.

Despite this longer waiting time to colonoscopy, the proportion of colonoscopies during which a SAE was reported did not change between periods. Although high, the frequency of perforations remains lower than that (1.1%) found in Alsace[25]. In the program, there was no nationally standardized forms for collecting screening colonoscopy data. Information concerning the date of consultation before the colonoscopy, or the progress of the examination can sometimes be missed or be considered irrelevant during this collection. Therefore, the low frequency of SAEs reported in this study could be the consequence of under-reporting.

**Table 4 Average time (in months) to colonoscopy according to the characteristics of the cohort of gastroenterologists who performed at least one colonoscopy over each of the four periods (guaiac fecal occult blood test, fecal immunochemical test, STOP-fecal immunochemical test, and COVID)**

Characteristics of the cohort of gastroenterologists	Nb of GE	Average time (in months) to colonoscopy, by period								P <sup>1</sup>
		gFOBT		FIT		STOP-FIT		COVID		
		Nb of Colo	Average, 95%CI	Nb of Colo	Average, 95%CI	Nb of Colo	Average, 95%CI	Nb of Colo	Average, 95%CI	
Overall	533	21509	2.6 [2.5; 2.6]	38352	3.0 [2.9; 3.0]	7342	3.9 [3.8; 4.0]	7995	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
Annual Nb of Colo										
1	201 <sup>2</sup>	304	3.1 [2.8; 3.5]	150	3.3 [2.7; 3.8]	38	4.3 [3.1; 4.8]	51	4.0 [3.5; 4.6]	0.08
2-30	481 <sup>2</sup>	16819	2.6 [2.5; 2.6]	15970	3.0 [3.0; 3.1]	4887	3.9 [3.8; 4.0]	4211	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
31-100	44 <sup>2</sup>	4386	2.4 [2.3; 2.5]	21137	3.0 [2.9; 3.0]	2817	3.8 [3.6; 3.9]	3733	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
> 100	0	0		1095	2.5 [2.3; 2.6]	0		0		
Place of S-colo performance										
Clinic	355 <sup>2</sup>	15745	2.4 [2.4; 2.5]	27003	2.9 [2.8; 2.9]	5039	3.7 [3.6; 3.7]	5560	3.4 [3.3; 3.5]	< 10 <sup>-3</sup>
Private hospital	125 <sup>2</sup>	3041	2.5 [2.4; 2.6]	6500	2.9 [2.8; 3.0]	1359	3.6 [3.5; 3.7]	1621	3.4 [3.3; 3.5]	< 10 <sup>-3</sup>
Public hospital	235 <sup>2</sup>	2723	3.3 [3.2; 3.4]	4849	3.8 [3.7; 3.9]	940	5.1 [4.7; 5.9]	795	4.2 [3.8; 4.7]	< 10 <sup>-3</sup>
Average density of GE (GE/100000iHbts)										
Low	127 <sup>2</sup>	4643	2.4 [2.3; 2.5]	8419	2.9 [2.8; 2.9]	1519	3.9 [3.8; 4.1]	1800	3.4 [3.3; 3.5]	< 10 <sup>-3</sup>
Average	108 <sup>2</sup>	3245	2.5 [2.4; 2.5]	4314	2.9 [2.8; 2.9]	810	4.1 [3.8; 4.4]	781	3.6 [3.4; 3.9]	< 10 <sup>-3</sup>
High	467 <sup>2</sup>	13621	2.6 [2.6; 2.7]	25619	3.0 [3.0; 3.1]	5009	3.8 [3.7; 3.9]	5395	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
Annual Nb of Colo locations										
1 location	483 <sup>2</sup>	14437	2.6 [2.6; 2.7]	24851	3.0 [3.0; 3.1]	4763	3.8 [3.7; 3.9]	5160	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
≥ 2 locations	153 <sup>2</sup>	7072	2.4 [2.4; 2.5]	13501	2.9 [2.9; 3.0]	2575	4.0 [3.9; 4.1]	2816	3.5 [3.4; 3.7]	< 10 <sup>-3</sup>
Residence of the patient										
Colonoscopy's supply municipality	338 <sup>2</sup>	4947	2.5 [2.4; 2.5]	7775	2.9 [2.9; 3.0]	1502	3.9 [3.7; 4.1]	1530	3.5 [3.3; 3.6]	< 10 <sup>-3</sup>
Other municipality in Colonoscopy's supply department	480 <sup>2</sup>	13259	2.5 [2.5; 2.6]	23754	3.0 [3.0; 3.1]	4401	3.9 [3.8; 4.0]	4982	3.5 [3.4; 3.6]	< 10 <sup>-3</sup>
Other departments in IDF	419 <sup>2</sup>	3303	2.7 [2.6; 2.8]	6823	2.9[2.9; 3.0]	1435	3.8 [3.6; 4.0]	1464	3.6 [3.5; 3.8]	< 10 <sup>-3</sup>
Age (in yr) of the patients										
50-54	485 <sup>2</sup>	4995	2.7 [2.6; 2.8]	8018	3.1 [3.0; 3.2]	1695	4.1 [3.9; 4.2]	1616	3.8 [3.7; 4.0]	< 10 <sup>-3</sup>
55-59	452 <sup>2</sup>	4669	2.6 [2.5; 2.7]	7355	3.1 [3.0; 3.1]	1446	3.9 [3.7; 4.1]	1560	3.6 [3.4; 3.7]	< 10 <sup>-3</sup>
60-64	466 <sup>2</sup>	4889	2.5 [2.4; 2.6]	7851	3.0 [2.9; 3.0]	1478	3.7 [3.5; 3.9]	1531	3.5 [3.4; 3.7]	< 10 <sup>-3</sup>
65-69	464 <sup>2</sup>	3766	2.5 [2.4; 2.5]	8511	2.9 [2.8; 2.9]	1403	3.8 [3.6; 4.0]	1590	3.3 [3.1; 3.5]	< 10 <sup>-3</sup>
≥ 70	431 <sup>2</sup>	3190	2.4 [2.3; 2.5]	6617	2.9 [2.8; 3.0]	1316	3.7 [3.5; 3.8]	1679	3.4 [3.3; 3.6]	< 10 <sup>-3</sup>

<sup>1</sup>(Prob > F) ANOVA.<sup>2</sup>This is the number of gastroenterologists (GEs) having performed a colonoscopy over the guaiac fecal occult blood test period, the same GE can be present in all of the modalities, for example the same GE having performed colonoscopies at 3 different sites corresponding to each of the density zones (low average high).

95%CI: 95% confidence interval; Colo: Screening colonoscopy; FIT: Fecal immunochemical test; GE: Gastroenterologist; gFOBT: Guaiac fecal occult blood test; iHbts: Inhabitants; IDF: Ile-De-France; Nb: Number.

The high proportion of incomplete colonoscopies due to insufficient preparation should alert to the need to set up a specific preparation protocol for screening colonoscopy. To date, it is impossible to evaluate with relevance the preparation of a colonoscopy in outpatients, who are not hospitalized at the time of the preparation. Similarly, there is no standard preparation scheme imposed in the French



**Table 5** Proportion of colonoscopies performed beyond 7 mo and proportion of serious adverse events, according to the characteristics of the cohort of gastroenterologists who performed at least one colonoscopy in each of the three periods (guaiac fecal occult blood test, fecal immunochemical test, STOP-fecal immunochemical test, and COVID)

Characteristics of the cohort of gastroenterologists	Proportion of colonoscopies performed beyond 7 mo by period					Proportion of serious adverse events by period				
	gFOBT	FIT	STOP-FIT	COVID	<i>P</i> <sup>1</sup>	gFOBT	FIT	STOP-FIT	COVID	<i>P</i> <sup>1</sup>
	Nb of Colo (% > 7 mo)	Nb of Colo (% > 7 mo)	Nb of Colo (% > 7 mo)	Nb of Colo (% > 7 mo)		%EI	%EI		%EI	
Overall	21509 (5.3)	38352 (6.2)	7342 (11.3)	7995 (9.2)	< 10 <sup>-3</sup>	0.3	0.3	0.3	0.2	0.10
Annual Nb of Colo										
1	304 (10.1)	150 (11.8)	38 (18.4)	51 (12.0)	< 10 <sup>-3</sup>	0.3	0	0	0	0.67
2-30	16819 (5.5)	15970 (6.9)	4887 (11.8)	4211 (9.4)	< 10 <sup>-3</sup>	0.3	0.4	0.3	0.1	0.09
31-100	4386 (4.1)	21137 (5.7)	2817 (10.6)	3733 (9.0)	< 10 <sup>-3</sup>	0.4	0.2	0.3	0.2	0.25
> 100		1095 (3.7)					0.2			
Place of Colo performance										
Clinic	15745 (4.8)	27003 (5.4)	5039 (10.4)	5560 (8.3)	< 10 <sup>-3</sup>	0.3	0.3	0.3	0.1	0.16
Private hospital	3041 (5.1)	6500 (5.9)	1359 (9.6)	1621 (9.5)	< 10 <sup>-3</sup>	0.3	0.2	0.2	0.2	0.71
Public hospital	2723 (8.5)	4849 (10.6)	940 (18.8)	795 (15.1)	< 10 <sup>-3</sup>	0.6	0.6	0.2	0.4	0.48
Average density of GE (GE/100000 iHbts)										
Low	4643 (5.1)	8419 (5.8)	1519 (11.5)	1800 (9.1)	< 10 <sup>-3</sup>	0.3	0.2	0.3	0.1	0.59
Average	3245 (4.9)	4314 (6.5)	810 (11.0)	781 (9.9)	< 10 <sup>-3</sup>	0.2	0.3	0.1	0.5	0.48
High	13621 (5.5)	25619 (6.2)	5009 (11.3)	5395 (9.2)	< 10 <sup>-3</sup>	0.4	0.3	0.3	0.1	0.04
Annual Nb of Colo locations										
1 location	14437 (5.6)	24851 (6.3)	4763 (10.6)	5160 (9.5)	< 10 <sup>-3</sup>	0.4	0.3	0.3	0.2	0.17
≥ 2 location	7072 (4.8)	13501 (6.0)	2575 (12.7)	2816 (8.7)	< 10 <sup>-3</sup>	0.2	0.2	0.2	0.1	0.80
Residence of the patient										
Colonoscopy's supply municipality	4947 (4.9)	7775 (5.9)	1502 (11.6)	1530 (8.0)	< 10 <sup>-3</sup>	0.2	0.3	0.3	0.0	0.18
Other municipality in Colonoscopy's supply department	13259 (5.1)	23754 (6.0)	4401 (11.0)	4982 (9.6)	< 10 <sup>-3</sup>	0.4	0.3	0.2	0.2	0.02
Other departments in IDF	3303 (6.6)	6823 (6.9)	1435 (12.1)	1464 (9.2)	< 10 <sup>-3</sup>	0.3	0.4	0.4	0.3	0.80
Age (in yr) of the patients										
50-54	4995 (6.7)	8018 (7.0)	1695 (12.2)	1616 (11.0)	< 10 <sup>-3</sup>	0.2	0.2	0.1	0.1	0.43
55-59	4669 (5.7)	7355 (6.6)	1446 (11.8)	1560 (9.9)	< 10 <sup>-3</sup>	0.3	0.3	0.4	0.2	0.79
60-64	4889 (4.7)	7851 (5.9)	1478 (10.3)	1531 (8.9)	< 10 <sup>-3</sup>	0.4	0.3	0.5	0.3	0.76
65-69	3766 (4.4)	8511 (5.5)	1403 (11.7)	1590 (8.2)	< 10 <sup>-3</sup>	0.4	0.3	0.3	0.1	0.20
≥ 70	3190 (4.6)	6617 (5.7)	1316 (10.6)	1679 (8.2)	< 10 <sup>-3</sup>	0.5	0.4	0.2	0.2	0.49

<sup>1</sup>Cochran Q test.

Colo: Screening colonoscopy; Colo+: Positive screening colonoscopy; FIT: Fecal immunochemical test; GE: Gastroenterologist; gFOBT: Guaiac fecal occult blood test; iHbts: Inhabitants; IDF: Ile-De-France; Nb: Number.

screening program, each GE proposing the method of his choice to the patient. However, although a non-superiority of a preparation scheme (Enema *vs* Oral preparation) was argued[21], studies admitted that a short time (1-6 h *vs* > 8 h) between the colic preparation and colonoscopy is associated with a

**Table 6 Neoplasm detection rate at colonoscopy, according to the characteristics of the cohort of gastroenterologists who performed at least one colonoscopy in each of the three periods (guaiac fecal occult blood test, fecal immunochemical test, STOP-fecal immunochemical test, and COVID)**

Characteristics of the cohort of gastroenterologists	Neoplasm detection rate at colonoscopy by period				P <sup>1</sup>
	gFOBT	FIT-1	STOP-FIT	COVID	
	Nb of Colo (% Colo+)	Nb of Colo (% Colo+)	Nb of Colo (% Colo+)	Nb of Colo (% Colo+)	
Overall	21509 (52.3)	38352 (65.0)	7342 (63.3)	7995 (60.1)	< 10 <sup>-3</sup>
Annual Nb of Colo					
1	304 (50.7)	150 (62.0)	38 (71.1)	51 (51.0)	0.02
2-30	16819 (53.6)	15970 (64.5)	4887 (63.7)	4211 (59.6)	< 10 <sup>-3</sup>
31-100	4386 (47.6)	21137 (65.8)	2817 (62.5)	3733 (61.8)	< 10 <sup>-3</sup>
> 100		1095 (58.6)	0	0	
Place of S-colo performance					
Clinic	15745 (52.4)	27 003 (64.9)	5039 (62.8)	5560 (60.1)	< 10 <sup>-3</sup>
Private hospital	3041 (54.3)	6500 (65.0)	1359 (64.8)	1621 (62.7)	< 10 <sup>-3</sup>
Public hospital	2723 (49.8)	4849 (65.6)	940 (63.7)	795 (59.5)	< 10 <sup>-3</sup>
Average Density of GE (GE/100000iHbts)					
Low	4643 (53.0)	8419 (64.5)	1519 (61.7)	1800 (58.4)	< 10 <sup>-3</sup>
Average	3245 (53.1)	4314 (64.1)	810 (65.2)	781 (61.1)	< 10 <sup>-3</sup>
High	13621 (51.9)	25619 (65.4)	5009 (63.4)	5395 (61.2)	< 10 <sup>-3</sup>
Annual Nb of S-colo locations					
1 location	14437 (52.8)	24851 (64.8)	4763 (63.0)	5160 (60.8)	< 10 <sup>-3</sup>
≥ 2 locations	7072 (51.3)	13501 (65.6)	2575 (63.8)	2816 (60.2)	< 10 <sup>-3</sup>
Residence of the patient					
Colonoscopy's supply municipality	4947 (53.4)	7775 (65.3)	1502 (61.1)	1530 (59.5)	< 10 <sup>-3</sup>
Other municipality in Colonoscopy's supply department	13259 (51.7)	23754 (65.0)	4401 (64.2)	4982 (61.2)	< 10 <sup>-3</sup>
Other departments in IDF	3303 (53.1)	6823 (64.9)	1435 (62.8)	1464 (59.6)	< 10 <sup>-3</sup>
Age (in yrs) of the patients					
50-54	4995 (44.9)	8018 (56.5)	1695 (55.2)	1616 (52.9)	< 10 <sup>-3</sup>
55-59	4669 (50.8)	7355 (63.1)	1446 (61.3)	1560 (58.9)	< 10 <sup>-3</sup>
60-64	4889 (54.2)	7851 (67.9)	1478 (65.5)	1531 (63.1)	< 10 <sup>-3</sup>
65-69	3766 (57.1)	8511 (68.9)	1403 (68.3)	1590 (65.3)	< 10 <sup>-3</sup>
≥ 70	3190 (57.5)	6617 (69.2)	1316 (67.9)	1679 (62.8)	< 10 <sup>-3</sup>

<sup>1</sup>Cochran Q test.

Colo: Screening colonoscopy; Colo+: Positive screening colonoscopy; FIT: Fecal immunochemical test; GE: Gastroenterologist; gFOBT: Guaiac fecal occult blood test; iHbts: Inhabitants; IDF: Ile-De-France; Nb: Number.

better quality of colonic preparation[26].

Compared to gFOBT, the high proportion of 2<sup>nd</sup> colonoscopies over the FIT period would confirm the literature on the performance of FIT in screening for precancerous lesions[27], which most often only require endoscopic resection. However, in addition to a high proportion of obstructive lesions, the proportion of severe cancers was significantly higher over the gFOBT period.

Several study results converge on a link between the long delay in access to colonoscopy and the CRC risk. Forbes *et al*[28] propose that wherever possible, colonoscopy should not be delayed beyond 6 mo of positive fecal testing as an aspirational target (with 9 mo as an upper limit). In the Kaiser Permanente (California) health plan members, the risk of CRC was increased by about 40% for any colonoscopy

**Table 7** Multivariate analysis of the relationship between each dependent variable (binary variables 0/1: Screening colonoscopy performed beyond 7-mo; Serious adverse events, Yield of neoplasm at screening colonoscopy) and the predictive factors

Characteristics of the cohort of gastroenterologists	Colo performed beyond a 7-mo risk analysis		Serious adverse events risk analysis		Neoplasms risk analysis	
	OR <sub>a</sub> , 95%CI	P <sup>1</sup>	OR <sub>a</sub> , 95%CI	P <sup>1</sup>	OR <sub>a</sub> , 95%CI	P <sup>1</sup>
Annual Nb of Colo (Ref: 1 Colo)						
2-30	0.7 [0.6; 1.0]	0.002	2.5 [0.3; 18.2]	0.37	0.9 [0.7; 1.1]	0.41
> 30	0.7 [0.3; 0.9]	0.008	2.9 [0.9; 23.0]	0.05	0.8 [0.6; 1.1]	0.32
Place of S-colo performance (Ref: Clinic)						
Private hospital	1.2 [0.9; 1.6]	0.18	0.7 [0.3; 1.8]	0.47	1.1 [0.9; 1.3]	0.41
Public hospital	2.1 [1.3; 3.6]	0.001	1.6 [0.3; 8.7]	0.60	1.1 [0.8; 1.4]	0.20
Density of GE (Ref: Low)						
Average	0.9 [0.8; 1.0]	0.05	1.2 [0.6; 2.2]	0.59	1.0 [0.9; 1.1]	0.76
High	1.0 [1.0; 1.2]	0.28	1.2 [0.6; 2.3]	0.65	0.9 [0.8; 1.0]	0.04
Annual Nb of S-colo locations (Ref: 1 location)						
≥ 2 locations	1.1 [0.8; 1.5]	0.11	1.6 [0.5; 4.4]	0.41	1.0 [0.8; 1.3]	0.84
Residence of the patient (Ref: Colonoscopy's supply municipality)						
Other municipality in Colonoscopy's supply department	1.0 [0.9; 1.0]	0.30	1.0 [0.5; 2.1]	0.97	1.0 [0.9; 1.0]	0.31
Other departments in IDF	1.2 [1.1; 1.3]	< 10 <sup>-3</sup>	1.2 [0.3; 5.2]	0.81	0.9 [0.8; 1.0]	0.13
Age (yrs) of the patients (Ref: 50-54 yr)						
55-59	0.9 [0.8; 1.0]	0.03	1.6 [1.0; 2.6]	0.04	1.3 [1.2; 1.4]	< 10 <sup>-3</sup>
60-64	0.8 [0.7; 0.9]	0.001	2.0 [1.2; 3.1]	0.006	1.6 [1.5; 1.6]	< 10 <sup>-3</sup>
65-69	0.7 [0.7; 0.8]	< 10 <sup>-3</sup>	1.9 [1.2; 3.0]	0.01	1.7 [1.6; 1.8]	< 10 <sup>-3</sup>
≥ 70	0.7 [0.6; 0.8]	0.003	2.1 [1.3; 3.4]	0.002	1.7 [1.6; 1.8]	< 10 <sup>-3</sup>
Period (Ref.: gFOBT)						
FIT	1.2 [1.1; 1.2]	< 10 <sup>-3</sup>	0.8 [0.4; 1.5]	0.11	1.6 [1.5; 1.7]	< 10 <sup>-3</sup>
STOP-FIT	2.4 [2.1; 2.6]	< 10 <sup>-3</sup>	0.8 [0.5; 1.3]	0.27	1.3 [1.1; 1.5]	< 10 <sup>-3</sup>
COVID	2.0 [1.8; 2.2]	< 10 <sup>-3</sup>	0.5 [0.3; 0.9]	0.02	1.1 [1.0; 1.3]	0.08

<sup>1</sup>P > |z|.95%CI: 95% confidence interval; Colo: Screening colonoscopy; FIT: Fecal immunochemical test; GE: Gastroenterologist; gFOBT: Guaiac fecal occult blood test; IDF: Ile-De-France; OR<sub>a</sub>: Adjusted odds-ratio.

performed after a waiting period of 7-12 mo[17]. A recent meta-analysis shows that the risk of colorectal cancer is increased by 42%, and that the risk of cancer at an advanced stage was multiplied by 2 or even more, when colonoscopy was performed more than 6 mo after a positive test[29]. In this study, the time to access colonoscopy as well as its lengthening, induced first by the change of the test and then by the health crisis, had no impact in terms of the CRC severity, probably because of the discriminatory approach prioritizing patients with already existing symptoms. As a reminder, the French Society of Digestive Endoscopy had made, in mid-April 2020, the specific recommendation to postpone by 6 wk any colonoscopy following a positive screening test result, if there was no clinical nor biological sign of CCR[30]. In addition, since FIT was introduced in 2015 in a population screened biannually with gFOBT, the severe CRC screened by FIT are likely to be those not detected at an early stage by gFOBT. This hypothesis is confirmed by the drop in the colonoscopy detection rate and by the proportion of severe CRC over STOP-FIT and COVID periods.

To celebrate the tenth anniversary of the first atlas of medical demography, the National Council of the Order of Physicians focused on the gradual transfer from liberal activity to salaried activity. The focus also mentioned the widening of territorial inequalities to the detriment of regions and departments already in difficulty in terms of medical density[31]. Although the number of GEs is unevenly distributed over the 1268 IDF municipalities, the density of GEs in the IDF region was well

above the range (4.2 to 4.9) of the national average observed in 2017[31].

Each GE participating in a CRCSP must perform at least 300 colonoscopies per year[21]. Despite the superiority of the regional offer compared to the national average, the annual number of colonoscopies per GE stays very disparate and below 300, especially for GEs in public hospitals. The main limitation of this study is the fact that it only gives an opinion on screening colonoscopies. Indeed, screening colonoscopies only represented 5.5% of all colonoscopies performed in France in 2012 (gFOBT-period) and about 10% in 2016 (FIT-period)[32]. Since the patient base of a GE is not limited to the population of the region of practice, several GEs in the IDF region could reach or exceed this recommended annual number, in particular GEs practicing in a private clinic. The other limit of the study would come from the fact that the measurements of the indicators cannot be generalized over the whole of France. Indeed, the density of gastroenterologists and the types of practice (clinical hospital, *etc.*) may vary from one municipality (or department or region) to another. Only access to databases for the reimbursement of colonoscopy procedures could allow the exhaustive evaluation of such a quality indicator.

## CONCLUSION

Although GEs are unevenly distributed over the municipalities of the IDF region, the supply of colonoscopies has remained almost constant between 2010 and 2020. The increase in colonoscopy requests induced by the change of the test kit has led to an increase in the average annual number of colonoscopies performed by GEs at the start of the FIT period. This very disparate annual average number between GEs fell over the STOP-FIT and COVID periods, due to the decrease in demand induced by the shutdown of the test kit market and the COVID-19 health crisis. The definition of a reference time and the obligation to respect it by all GEs would effectively reduce the time to access screening colonoscopy in France. The increase in the time to colonoscopy between the first and the second period was attributable to the introduction of the FIT, its increase after the second period was probably attributable to the malfunction of the program due to the slowdown of the kit market and the COVID-19 health crisis. Regardless of the characteristics of the GEs, the colonoscopy detection rate dropped significantly between the STOP-FIT and COVID periods, while the proportions of SAEs remained unchanged. However, the time to colonoscopy as well as its lengthening induced by the constraints had no impact in terms of CRC severity, probably because of a discriminatory approach prioritizing patients with existing symptoms.

## ARTICLE HIGHLIGHTS

### Research background

The impact of the Screening program on controlling the colorectal cancer (CRC) morbidity and mortality has been proved. But since its complete roll-out in 2009, the French population-based colorectal cancer screening program (CRCSP) experienced 3 major constraints [use of a less efficient Guaiac-test (gFOBT), Stopping the supply of Faecal-Immunochemical-Test kits (FIT), Suspension of the program due to the coronavirus disease 2019 (COVID-19)] affecting its effectiveness.

### Research motivation

At this time when all the spotlights are focused on the impact of the health crisis linked to COVID-19, our motivation was to warn of the continued deterioration in the quality of screening colonoscopies in France.

### Research objectives

To describe the impact of the constraints in terms of changes to the quality of screening colonoscopies.

### Research methods

This retrospective cohort study included screening colonoscopies performed by the gastroenterologists between January 2010 and December 2020 in people aged 50-74 Living in Ile-de-France (France). The changes to the quality of screening colonoscopy (proportion of colonoscopies performed beyond 7 mo, Frequency of serious adverse events and the colonoscopy detection rate) were described in a cohort of Gastroenterologists who performed at least one colonoscopy over each of the four periods defined according to the chronology of the constraints [gFOBT: Normal progress of the CRCSP using gFOBT (2010-2014); FIT: Normal progress of the CRCSP using FIT(2015-2018); STOP-FIT: Year (2019) during which the CRCSP experienced the cessation of the supply of test kits; COVID: program suspension due to the COVID-19 health crisis (2020)]. The link between each dependent variable (Colo\_7 mo; SAE Occurrence, Neoplasm detection rate) and the predictive factors was analyzed in a two-level multivariate hierarchical model.

### Research results

The retrospective cohort was made up of 533 gastroenterologists. These 533 gastroenterologists achieved 21509 screening colonoscopies over the gFOBT period, 38,352 over FIT, 7342 over STOP-FIT and 7995 over the COVID period. The frequency of serious adverse events did not change between periods (gFOBT: 0.3%; FIT: 0.3%; STOP-FIT: 0.3%, and COVID: 0.2%;  $P = 0.10$ ). The risk of colonoscopies performed beyond 7 mo doubled between FIT [adjusted-odds-ratio (aOR): 1.2 (1.1; 1.2)] and STOP-FIT [aOR: 2.4 (2.1; 2.6)], then decreased by 40% between STOP-FIT and COVID [aOR: 2.0 (1.8; 2.2)]. Regardless of the period, this Colo\_7 mo's risk was twice as high for screening colonoscopy performed in a public hospital [aOR: 2.1 (1.3; 3.6)] compared to screening-colonoscopy performed in a private clinic. The neoplasm detection, which increased by 60% between gFOBT and FIT [aOR: 1.6 (1.5; 1.7)], decreased by 40% between FIT and COVID [aOR: 1.1 (1.0; 1.3)].

### Research conclusions

The study showed that the constraints likely affected the time-to-colonoscopy as well as the colonoscopy detection rate without impacting the occurrence of the serious adverse events, highlighting the need for a respectable reference time-to-colonoscopy in CRCSP.

### Research perspectives

At the end of this study, we initially aim to develop, evaluate, and validate a standard form for collecting data from screening colonoscopies in France. In a second step, we will evaluate the impact of the patient's motivation by the attending physician on the time taken to perform the colonoscopy.

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## FOOTNOTES

**Author contributions:** Koivogui A, Vincelet C, Abihsera G, Ait-Hadad H, Delattre H, Le Trung T, and Bernoux A are the doctors in charge of coordinating the screening program in each department; Nicolet J is the medical director of the CRCDC-IDF; Koivogui A conceptualized and designed the project; all doctors in charge of coordinating the screening program collected the field data; Koivogui A, Vincelet C, and Abihsera G analyzed the data, interpreted the results, and drafted the manuscript; all the authors revised the manuscript, read, and approved the final version of this manuscript.

**Institutional review board statement:** This study is co-signed by the heads of the structures involved, as such, no further Institutional Review Board was required.

**Informed consent statement:** Patients were not required to give informed consent to the study because the analysis used anonymous data that was obtained after each patient agreed to participate in screening campaigns.

**Conflict-of-interest statement:** The authors declare that they have no conflict of interest.

**Data sharing statement:** Data and materials are available when requested by e-mail. However, each request will be processed following French legislation on the availability of research data.

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**Country/Territory of origin:** France

**ORCID number:** Akoï Koivogui 0000-0001-9097-3006.

**S-Editor:** Chen YL



L-Editor: A

P-Editor: Chen YL

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