

Screening for colorectal cancer in a factory-based population with Fecatest

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Summary This report concerns a screening programme for colorectal cancer using Fecatest, a relatively sensitive test for faecal occult blood, in a factory-based population. A total of 2420 workers between 40 and 65 years of age returned kits for testing following suitable dietary restriction. In Factory A, in which screening was offered as part of an annual medical examination, 989 men agreed to participate, a compliance rate of 45%. In Factory B, in which screening was offered in their weekly pay packets, 1431 men participated, a compliance rate of 22%. An overall positivity rate of 5.8% was found, 4.6% in Factory A and 6.6% in Factory B. Five cancers were found, two of the rectum, one at the recto-sigmoid junction, one in the sigmoid colon and one in the transverse colon. Two of these lesions were at Dukes Stage A, one at Stage B and two at Stage C. In addition, 25 adenomatous polyps were found in 17 men. In 13 these were characterized as tubular adenomata and in 4 as tubulovillous adenomata. The mean age at diagnosis in the polyp cases was 52.6 years and in the cancer cases 58.8 years. Increased specificity is obtained by excluding participants below 46 years of age. The detection rate in the 46-65 year range is ~1 in 100 for adenomatous polyps and 1 in 300 for cancer in the population screened. This detection rate is higher than most comparable studies using a less sensitive Guaiac test on older populations.

The high incidence of colorectal cancer in Western communities (Office of Population 1975; Cancer Facts & Figures 1977) has led to the introduction of screening programmes with the aim of increasing the proportion of localised (Dukes A & B) cancer at presentation. Two studies utilizing sigmoidoscopy and polyp removal suggested that such a programme might reduce the incidence of colorectal cancer although no control observations were made (Hertz *et al.*, 1960; Gilbertsen, 1973). Dales *et al.* (1973) demonstrated reduced mortality from colorectal cancer utilizing sigmoidoscopy in a screened group compared with controls. Gregor (1971) popularised the Guaiac test for faecal occult blood as a screening test to select a proportion of asymptomatic individuals for fuller investigation of the large bowel. Subsequently, the Haemoccult II test has been widely used in screening programmes with the finding of ~1 carcinoma per 500-1000 persons screened (Glober & Peskoe, 1974; Miller and Knight 1977; Bralow, 1979). In addition, in recent programmes in which colonoscopy was employed ~3 times as many adenomatous polyps as cancers were discovered (Winawer *et al.*, 1977; Frühmorgen & Demling, 1980). It is hoped that removal of polyps by snare polypectomy may also reduce the incidence of subsequent cancer in view of the known tendency for such lesions to pass from a benign to a malignant phase (Grinnell & Lane 1958; Morson, 1966). A number of screening

programmes have been carried out and early results of two controlled studies (Winawer *et al.*, 1980; Hardcastle *et al.*, 1983) indicate benefit for the screened population in terms of increased diagnosis of localized lesions (Dukes A & B) with corresponding improved prognosis (Bussey, 1963; Gill & Morris, 1978). Problems remaining relate to identification and compliance of the population to be screened, the sensitivity and specificity of the tests involved and the cost/benefit of the programme compared with any disadvantage from physical and psychological injury involved in the investigative programme. This study reports the results of a screening programme for colorectal cancer in a factory-based population using Fecatest.

Methods

Fecatest

The Fecatest paper is fitted between 2 plastic compartments, the patient applying one small specimen of faeces to one side of the paper and the test is completed on the other side. Fecatest has a sensitivity just above what is regarded as normal blood loss $-2.5-5.0 \text{ ml day}^{-1}$ (Adlercreutz *et al.*, 1978). Two factory groups were studied, the programme being aimed at male workers aged between 40 and 65 years.

Factory A

Employees were offered screening for colorectal

cancer as part of their annual medical examination. Nurses from the Occupational Health Department gave written and verbal instructions with regard to the faecal occult blood tests. Completed kits were returned to the factory Medical Department where they were interpreted by trained laboratory technicians.

Factory B

The same screening programme was offered to employees by written invitation in their pay packets. They attended the factory Medical Department to collect the kits from nursing staff who gave verbal and written instructions. Completed kits were sent by post to the Department of Gastroenterology, Victoria Hospital, Blackpool, for interpretation.

Participants were invited to complete a questionnaire relating to family history of cancer, and current drug usage, notably antirheumatic and analgesic agents.

As recommended by the manufacturers, the kits were used on 3 successive stools, or days on which stools were passed, following a 3-day period of dietary restriction. Red meat and some vegetables and fruit contain peroxidase activity and for more specific results individuals should take a diet excluding red meat, liver, bananas, swede, tomatoes and turnips. Fish and chicken are allowed. A diet containing vegetables and wholemeal bread is recommended. Vitamin C may produce false negative tests and patients are advised to exclude such preparations.

Workers with one or more positive test were investigated by flexible sigmoidoscopy, barium enema and colonoscopy in the Gastroenterology and Radiology Departments, Victoria Hospital. Polyps were removed by snare polypectomy.

Where appropriate, statistical differences between groups were calculated using the Chi square test.

Results

Compliance

In Factory A, in which screening was offered to workers as part of an annual medical examination, 45% returned the test kits. In contrast, in Factory B, in which screening was offered in the pay packet, response was only 22% of those approached. ($P < 0.001$).

Positivity rate

The combined positivity rate for the two factories was 5.8%, for Factory A it was 4.6% and for Factory B 6.6%. This difference is significant

($P < 0.05$). The majority of Fecatest positive participants completed the investigation programme. Only 8 men withdrew, 6 after flexible sigmoidoscopy and 2 after the barium enema examination.

Pathological findings

The findings in the total group and the 2 factories are shown in Table I. Five cancers were found, 2 at Dukes Stage A, one at Stage B and 2 at Stage C. Details of these are shown in Table II. Twenty-five adenomatous polyps were found in 17 men. In 13 men the lesions were characterized as tubular adenomata and in 4 as tubulovillous adenomata. Four men were found to have occult inflammatory

Table I Results of screening programme in total participants and Factories A & B

	Total	Factory A		Factory B
Number screened	2,420	989		1,431
Fecatest positive	140(5.8%)	45(4.6%)	$P < 0.05$	95(6.6%)
Adenomatous polyps	17	3	NS	14
Cancer	5	3	NS	2
I.B.D.	4	1	NS	3
Diverticular disease	23	8	NS	15
Perianal disease	50	15	NS	35
Nil ("false positive")	52(32%)	17	NS	35

(Some men had more than one lesion).

Table II Cases of colorectal cancer

Case	Age	Site	Surgical procedure	Details	Dukes grade
C1	60	Rectum	Abdomino-perineal resection	Lymph node involved 0/6	A
C2	61	Recto-sigmoid 15 cm from anus	Anterior resection	Into M. propria Lymph node involved 0/0	B
C3	54	Rectum	Abdomino-perineal resection	6 cm diam. into M. propria Lymph node involved 1/6	C
C4	58	Sigmoid 35 cm from anus	Resection	1 × 1.5 cm diam. into submucosa Lymph node involved 1/6	C
C5	61	Mid transverse colon	Resection	Polypoid. No invasion. Lymph node involved 0	A

bowel disease. In addition, a proportion of men were found to have perianal disease or diverticular disease although the significance of these as causes of occult blood loss is dubious. The remaining so-called false positive cases were not investigated for upper gastrointestinal disease, prior consent not having been obtained. No patient had reported gastro-intestinal symptoms or rectal bleeding and none had anaemia.

Table III shows the age at diagnosis in the 17 men in whom adenomatous polyps were found and the size of the largest polyp. One man had 3 polyps, 6 men had 2 and 10 had only 1 polyp. Twelve patients had polyps >1 cm in diameter. All except 1 polyp were removed by snare polypectomy. The exception had a small (0.4 cm) polyp in an area of diverticular disease with a tortuous narrow sigmoid colon which, in spite of two attempts, could not be successfully snared.

Table III Diameter^a of adenomatous polyps and age at diagnosis

Age range— years	No.	<1 cm	1–2 cm	>2 cm
Under 46	2	0	1	1
46–50	5	2	3	0
51–55	5	1	1	3
56–60	3	2	1	0
61–65	2	0	1	1
All	17	5	7	5

^aWhere more than one polyp was present, the diameter of the largest is given.

Surgical resection of the area was carried out without complication. All the polyps and 4/5 cancers were found distal to the splenic flexure. One large malignant polyp was situated in the mid-transverse colon. The mean age at diagnosis of the 17 polyps was 52.6 years and of the 5 cancers 58.8 years. It may reasonably be speculated that removal of the benign lesions may be associated with prevention of malignant disease later. The predictive value of the screening test is improved by excluding individuals under 46 years of age (Table IV). The detection rate in the 46–65 years age range is 1 in 100 for adenomatous polyps and 1 in 300 for cancer in the population screened.

In the 22 men in whom polyps and cancers were found, digital examination of the rectum revealed two cancers. At flexible sigmoidoscopy 13 polyps and 4 cancers were visible and on barium enema examination 8 polyps were noted, but 3 of these had not been visible at sigmoidoscopy. Colonoscopy revealed one polyp and one cancer not visible within the range of the flexible sigmoidoscopy nor on barium enema. The cancer was not seen on a single contrast examination but was visible in the transverse colon when the double contrast technique was used after colonoscopy.

Discussion

The decision to use Fecatest, a more sensitive Guaiac test for faecal occult blood than the previously more popular Haemocult, was prompted by an earlier finding of Fecatest positive

Table IV Screening for colorectal cancer in the United Kingdom: Details and analysis of studies

Reference	Population	Age range	Guaiac test	Positive tests no. screened	Percentage positive	CA + polyp (CA)	CA + polyp(CA) %/no. screened	CA + polyp(CA) %/no. F.O.B. positive. (predictive value)
Hardcastle & Balfour (1980)	General practice	>45	H*	27/713	3.8	6 (2)	0.84 (0.22)	22.2 (7.4)
Farrands <i>et al.</i> (1981)	General practice	>40	H*	124/2439	5.0	12 (4)	0.49 (0.16)	9.6 (3.2)
Million <i>et al.</i> (1982)	General practice	>45	H*	37/1646	2.3	7 (2)	0.43 (0.12)	18.9 (5.4)
Lee (1983)	Factory based	40–65	F**	140/2420	5.8	22 (5)	0.9 (0.21)	15.7 (3.6)
Lee (1983)	Factory based	46–65	F**	92/1575	5.8	20 (5)	1.27 (0.32)	21.7 (5.4)

H*—Haemocult.

F**—Fecatest.

Haemocult negative colorectal cancers (Lee & Costello 1982). Several reports indicate that an undetermined number of colorectal cancers are Haemocult negative (Griffith *et al.*, 1981; Ribet *et al.*, 1980; Gnauck, 1980) and a false negative rate of 76% for rectosigmoid polyps has been reported (Winawer *et al.*, 1977). Positive Haemocult tests can convert to negative after a few days' storage (Fleisher *et al.*, 1977; Heinrich *et al.*, 1980) and this is relevant where participants send in their specimens by post. The positivity rate for Haemocult can be increased by rehydration but some loss of specificity may occur (Fleisher *et al.*, 1977).

Although the majority of studies to date involve Haemocult, there have been suggestions that a more sensitive test would be useful (Vellacott *et al.*, 1981; Williams *et al.*, 1982). The use of Fecatest in this study has been associated with higher detection rates for colorectal cancer compared with other U.K. series based on general practice populations without a lowering of the predictive value of the basic screening test (Table IV).

Using a more sensitive test may result in an increase in false positivity and one problem here relates to the possibility that upper gastro-intestinal disease may be responsible for positive faecal occult blood tests in asymptomatic patients. With a sensitive test such as Fecatest the importance of proper dietary restriction should be stressed. This requires explanation to participants and a strong motivation to co-operate. It may be that these requirements are more likely to be met in screening a relatively young population, such as that based on place of work, rather than in surveys based in general practice, which usually include a higher proportion of elderly participants.

Although flexible sigmoidoscopy is useful in surveillance it cannot replace double contrast barium enema or colonoscopy since up to 50% of lesions may be beyond the range of the instruments. In this series 17/22 cancers and polyps (77%) were visible on flexible sigmoidoscopy. This high yield may reflect the relatively young population involved, since the incidence of right sided lesions

increases with age (Slater *et al.*, 1982). In one large series, 95% of malignant polyps were located distal to the splenic flexure (Appel, 1982) and in another large colonoscopic study only 34% of polyps and 36% of cancers were located beyond the range of the flexible sigmoidoscope (Tedesco *et al.*, 1980). Clearly examination of the whole colon in screening programmes is needed, but it seems that the diagnostic return is likely to be less in the right colon, the younger the population studied. Fork (1981) has compared the diagnostic accuracy of double contrast barium enema as assessed by colonoscopy and concluded that when both techniques are used only few polyps will be missed. The need for careful radiographic techniques is stressed.

Our finding of a co-operation rate of 45% in Factory A compared with 22% in Factory B coincides with previous observation that acceptance is more likely in groups attending for regular medical examination (Glober & Peskoe, 1974; Winawer *et al.*, 1977). Hardcastle *et al.* (1980) noted that in a general practice population compliance was highest in the age range 50–65 years declining sharply over 75 years of age. There was an overall compliance rate of 45% in their study compared with 27% in the "Frome experience" (Farrands *et al.*, 1981) and 28% in Salford (Million *et al.*, 1982). If screening programmes are demonstrated to be beneficial in terms of early diagnosis and prevention of colorectal cancer, more research is needed into optimum recruitment procedures (Halper *et al.*, 1980; Hardcastle *et al.*, 1983).

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