

Sensitivity and Specificity of Colorectal Cancer Mass Screening Methods: A Systematic Review of the Literature

Allameh Z¹, Davari M², Emami MH³

Abstract

Background: Colorectal cancer is the second cause of the cancer related mortalities in the world. Screening can effectively reduce the mortality and morbidity rate of this cancer. Sensitivity and specificity of screening methods play an important role in their efficacy. The aim of this study is to review sensitivity and specificity of selected colorectal cancer screening methods systematically.

Methods: This study conducted a systematic review of sensitivity and specificity of five common colorectal cancer screening methods. The sites Pubmed, Cochrane library and the center for review and dissemination (CRD database) were searched systematically in Jan 2009. Key questions for this search were focused on the sensitivity and specificity of the 5 screening methods.

Results: In these databases 2713 articles were matched well with our subject. Of these 130 articles were selected with specified inclusion-exclusion criteria. The mean \pm standard deviation per patient sensitivities of colonoscopy, sigmoidoscopy, double contrast barium enema, CT colonography and fecal occult blood test for detecting colorectal cancer were respectively $94.7 \pm 4.6\%$, $82.0 \pm 9.3\%$, $82.3 \pm 8.7\%$, $95.7 \pm 5.9\%$ and $45.7 \pm 26.5\%$. Specificities of these tests were respectively $99.8 \pm 0.2\%$, 83.9% , $92.4 \pm 14.7\%$, $98.5 \pm 1.3\%$ and $87.6 \pm 11.4\%$.

Conclusions: Based on available evidences, colonoscopy has the highest sensitivity and specificity among the selected screening methods and fecal occult blood test has the lowest sensitivity. Almost all of the tests except stool exams have acceptable sensitivity and specificity for detecting colorectal cancer.

Key words: Sensitivity; Specificity; Colorectal neoplasm; Mass screening

Please cite this article as: Allameh Z, Davari M, Emami MH. Sensitivity and Specificity of Colorectal Cancer Mass Screening Methods: A Systematic Review of the Literature. *Iran J Cancer Prev*. 2011; Vol4, No2, P88-105.

1. Dept. of Clinical Pharmacy, Faculty of Pharmacy, Shahid Beheshti University of Medical Sciences, Tehran, Iran
2. Dept. of Clinical Pharmacy, Faculty of Pharmacy, Isfahan University of Medical Sciences, Isfahan, Iran
3. Dept. of Gastroenterology, Faculty of Medicine, Isfahan University of Medical Sciences, Isfahan, Iran

Corresponding Author:
Zahra Allameh, Pharm D
Tel: (+98) 311 27 36 097
Email: zahraallameh@gmail.com

Received: 8 Dec. 2010
Accepted: 25 Mar. 2011
Iran J Cancer Prev 2011; 2:88-105

Introduction

Colorectal cancer has the greatest mortality rate after lung cancer among all malignancies and it has caused many problems in western countries [1]. Over 95% of colorectal cancers result from adenomatous polyps. These polyps are removable. This means that we can reduce the burden of this cancer by screening [2]. Screening is considered as a second type prevention. It means that it can detect patients that have a silent problem (with no symptoms), so they would go under treatment and their disease have no time to develop.

Some diseases are more suitable for screening. Colorectal cancer is one of them. It can be diagnosed

in early stages and it can be treated more effectively. Screening colorectal cancer will prevent it from becoming advanced [1, 3].

There are several methods for screening colorectal cancer. Each method has its own performance characteristics. Performance characteristics are the characteristics that can be used to describe the quality and usefulness of a test. Accuracy can be expressed through sensitivity and specificity, positive and negative predictive values, or positive and negative diagnostic likelihood ratios.

A proper test for screening should have four properties. It should be simple and not expensive [1]; should not have any danger to patients or healthy individuals [4]; should be acceptable for patients [1];

and it should be available [5]. It is obvious that the disease to be screened must be also treatable [4].

Some of the characteristics of the screening methods for colorectal cancer are explained as follows:

Colonoscopy

Colonoscopy has become increasingly popular for screening. There is a strong biologic argument that colonoscopy, with direct visualization of the colonic mucosa and the ability to biopsy or excise polyps and localized cancers can prevent colorectal cancers and deaths. Furthermore, colonoscopy has the potential to detect proximal lesions that would be missed by screening sigmoidoscopy [1]. This procedure has a low (0.35%) risk of serious complications [6]. A recent study published in the *Annals of Internal Medicine* [7] implies that colonoscopy screening prevents approximately two thirds of the deaths due to colorectal cancers on the left side of the colon, and is not associated with a significant reduction in deaths from right-sided disease.

Sigmoidoscopy

The 60 cm flexible sigmoidoscope can reach to the splenic flexure. Primary care physicians, nurse practitioners and physicians assistants can become proficient in the technique of sigmoidoscopy with proper training. The patient preparation is less onerous than for colonoscopy or CT colonography, and the procedure may be performed without sedation in the office.

Computed Tomographic Colonography

"Virtual Colonoscopy" is performed via computed tomography (CT), sometimes called a CAT scan, or with magnetic resonance imaging (MRI) [8]. About 1 in 10 patients will not have a complete right colon (cecum) evaluation completed with conventional colonoscopy [9]. It also takes less time than either a conventional colonoscopy or a lower GI series. VC provides a secondary benefit of revealing diseases or abnormalities outside the colon [10]. Patients prepare for CTC with aggressive bowel preparation, the same as used for optical colonoscopy. Virtual Colonoscopy performed with CT exposes the patient to ionizing radiation; however some research has demonstrated that ultra-low dose VC can be just as effective in demonstrating colon and bowel disease due to the great difference in x-ray absorption between air and the tissue comprising the inner wall of the colon.

Fecal Occult Blood Test

Several guaiac reagents are marketed; Hemocult-SENSA is more sensitive than Hemocult, Hemocult-II or Hemocult-R. Newer, high-sensitivity modern tests look for globin, and are now recommended as best practice. Conventional guaiac-based fecal occult blood tests which look for heme are a less-preferred option. If colon cancer is suspected in an individual (such as in someone with an unexplained anemia) fecal occult blood tests are typically not warranted. If a doctor suspects colon cancer, more rigorous investigation is necessary, whether or not the test is positive.

The test is often false-positive. (This is often due to recent ingestion of under-cooked meats, and a patient is generally advised to keep a meat-free diet for several days before handing in the feces sample. False negatives may result if the patient has been taking vitamin C supplements. The test is more sensitive if the sample is hydrated before testing. However, the specificity is decreased in this method.

Fecal Immunological Test (FIT)

FIT tests detect the globin in feces rather than heme. By detecting globin the tests are both more sensitive and specific for lower gastrointestinal bleeding. The superior FIT tests are now recommended in place of the traditional annual standard guaiac FOBT, which is now identified as a "less-preferred option."

Fecal immunochemical tests (FIT or iFOBT) for hemoglobin are more specific than guaiac tests because they respond only to human globin, and do not detect upper gastrointestinal bleeding (since the globin is digested in transit) or foods with peroxidase activity.

Fecal DNA Tests

Colorectal neoplasm shed DNA in the stool where it can be isolated and tested for the presence of changes acquired during carcinogenesis. Not all genetic abnormalities associated with colorectal cancer can be included in the stool DNA test (sDNA), however, and false negative results occur. The test is expensive. Fecal DNA test (PreGen-Plus) is more sensitive than fecal occult blood in one study (51.6% vs. 12.9%) [11].

Double-contrast Barium Enema

Patients must undergo bowel preparation. Sedation is not ordinarily given. Patients experience some cramping during the procedure but can return to work after the examination. There is less radiation

exposure than an abdominal CT. Most experts feel that the risk is low compared with the benefits.

Optical colonoscopy is taken as the "gold standard" for colorectal cancer screening by the vast majority of the medical and research communities. Some radiologists recommend VC as a preferred approach to colorectal screening. However, optical colonoscopy is considered the gold standard by some professionals because it permits complete visualization of the entire colon, hence providing the opportunity to identify precancerous polyps and cancer, and then to do diagnostic biopsies or therapeutic removal of these lesions, as soon as possible.

The aims of this study is to undertaken a systematic review of sensitivity and specificity of the five selected colorectal screening methods including colonoscopy, flexible sigmoidoscopy, CT colonography, double contrast barium enema and fecal occult blood test.

Materials and Methods

To review the sensitivity and specificity of each method of screening systematically 4 steps undertaken:

Step 1-Pubmed, Cochrane library and CRD database searched systematically in 24/1/2009.

Step 1-1: First the appropriate terms describe effectiveness and represent the accuracy of methods of screening were extracted from articles, books and especially from "MESH" database in the site "Pubmed". These terms were as follows:

Sensitivity, Specificity, Validity, Value, Accuracy, Efficacy, Effectiveness, Detection rate, Detectability, Positivity, Positive predictive value and Negative predictive value

Step 1-2: Then the terms were searched separately. The results then combined to each other by the logical operator "OR" to expand the research field appropriately.

We used the same strategy to search for "colon", "Cancer", and their synonyms separately. The terms used in this section were as follows:

Colon, Rectum, Colorectal, Bowel, Large intestine and Colonic

Step 1-3: The results then combined to each other by the logical operator "OR" to expand the research field appropriately.

Cancer, Carcinoma, Malignancy, Malignant, Tumor, Polyp, Adenoma, Neoplasia, Dysplasia, Hyperplasia and Neoplasm

Step 1-4: These results also were combined to each other by the logical operator "OR".

Then the name of each method of screening and their synonyms were searched systematically. The terms used in this section were as follows:

- Colonoscopy OR Colonoscopic
- Sigmoidoscopy OR Proctosigmoidoscopy
- FOBT OR "FOB test" OR "OB" OR Fecal occult blood test OR Guaiac OR "FOB"
- DCBE OR Double contrast barium enema OR Air contrast barium enema
- Virtual colonoscopy OR CT colonography OR Computed tomographic colongraphy

Step 1-5: The results of all these stages (steps 1-2, 1-3, 1-4 and the results for the names of each method of screening) put together with the operator "AND" for each method of screening separately to limit the search results reasonably.

Step 2-Reviewing the titles of the articles.

In this step the titles of articles were reviewed fully and proper ones were selected based on the inclusion and exclusion criteria.

Inclusion Criteria

All of the articles that directly measured sensitivity or specificity of each method of screening.

Exclusion Criteria

1-Studies that were not directly about measuring sensitivity or specificity of each method of screening and it is not the scope of their study.

2-Studies that were in the form of a review article, meta analysis or cost effectiveness analysis. The reason for putting these articles in exclusion criteria is that in this study we didn't use any "limit" and this study includes all of the studies that were used in review articles.

3-Because adenomatous and large polyps increase the risk of colorectal cancer, studies that only measured sensitivity of screening methods for small benign polyps, were omitted.

Step 3-Reviewing the abstracts of selected articles.

In this step the abstracts of selected articles were reviewed carefully and, according to inclusion and exclusion criteria, appropriate papers were selected to be reviewed by researchers.

Step 4-Reviewing the full text of the selected articles

In this step the full texts of selected papers reviewed and all relevant information were extracted.

Sensitivities/specificities were reported for different polyp sizes. Most of the reported sensitivities were for polyps ≥ 10 mm or cancer. In this study sensitivities that were reported for polyps ≤ 5 mm deleted; because these polyps do not indicate colorectal cancer [1].

The studies were conducted in different countries by different experts, equipments and in different times.

Results

In this study "the sensitivity for large polyps" means both the sensitivities reported for polyps ≥ 10 mm and polyps ≥ 6 mm and normally detecting the larger polyps is easier and the reported sensitivity/specificity is higher.

The results of studying articles and selecting proper ones using inclusion/exclusion criteria in the Pubmed, Cochrane library and CRD databases are presented in tables 1,2 and 3 respectively (Tables 1,2 , 3).

The summary of the sensitivity/specificity results extracted from the articles are presented in table 4.

Tables 5 and 6 summarized the results extracted from the publications based on the screening method and the criteria of the sensitivity.

Discussion

Diagnostic or screening tests should be valid, easy, with high compliance for patients and also cost-effective. The results show that colonoscopy has the highest sensitivity and specificity. In some methods of screening like sigmoidoscopy, there is actually some extent of lack of good evidences but in some of them like fecal occult blood test and CT colonography there are as many evidences as we can conclude a definite result from analyzing them. On the other hand

increasing evidences is not so good in some aspects. For example the sensitivity of guaiac test varies widely among different articles, but in most of them, this test has a very low sensitivity. So for this test and the tests like it, doing a meta analysis on the results is encouraged.

Some reasons of this wide range of sensitivity are followed:

Sensitivities of screening tests are reported in different ways in articles. For example per-polyp sensitivity indicates the number of polyps detected with a method out of 100 polyps. Per patient sensitivity is the number of patients with polyps that are detected out of 100 patients with polyps. In the articles studied, per patient sensitivities in most of the cases were more than per polyp sensitivities. It could be because of the fact that the numbers of polyps that are missed during screening are more than the number of patients that are missed. But the difference between these sensitivities was not so much that affect the results.

Sensitivities/specificities were reported for different polyp sizes. Most of the reported sensitivities were for polyps ≥ 10 mm or cancer. [In this study sensitivities that were reported for polyps ≤ 5 modulated. Because these polyps does not indicate colorectal cancer [2]. In this study "the sensitivity for large polyps" means both the sensitivities reported for polyps ≥ 10 mm and polyps ≥ 6 mm and normally detecting the larger polyps is easier and the reported sensitivity/specificity is higher.

The studies were conducted in different countries by different experts, equipments and in different times. This study did not use any limit for publication date.

In FOBT method, the sensitivity depends on the number of tests performed. For example doing the test in 6 days gives more sensitivity than doing 3 times. In most of the articles it was done 3 times.

Although CT colonography is used widely in these days, still colonoscopy has a high sensitivity and still can be the gold standard.

Among all types of sensitivities, sensitivity for cancer in all methods is the highest. It is because of the fact that when the lesion is in cancerous state it's much easier to detect.

Table1. The search results of the Pubmed database

Method of screening	The initial number of articles	Number of articles after reviewing the titles	Number of articles after reviewing the abstracts	Final number of articles
Colonoscopy	1239	56	24	20
Sigmoidoscopy	377	83	15	12
FoBt	539	101	47	38
BE	135	42	27	26
CT colonography	323	113	67	64

Table2. The search results of the Cochrane library

Method of screening	The initial number	Number of articles after reviewing the titles	Number of articles after reviewing the abstracts
Colonoscopy	123	8	1
Sigmoidoscopy	56	2	0
FoBt	31	4	1
CT colonography	24	6	0

Table3. The results of studying articles of CRD site

Method of screening	The initial number of articles	Number of articles after reviewing the titles	Number of articles after reviewing the abstracts
Colonoscopy	135	2	0
Sigmoidoscopy	65	0	0
FoBt	74	1	0
CT colonography	29	1	0

Table 4. The initial results of studying articles about sensitivity/specificity of colorectal cancer [12-139]

Colonoscopy

Authors and year	Sample size	Gold standard	Sensitivity (percent)			specificity	Ref. Number
			For large polyps	For cancer/ carcinoma	over all		
1 Heresbach D, Barrioz T(2008)	294	Same-day back-to-back video colonoscopy	74-91	100			12
2 Hosokawa O, Hattori M(2007)	59162	Cancer registry			88.9		13
3 Bressler B, Paszat LF(2007)	12487	Cancer Registry	94.1-97.7				14
4 Menardo G.(2004)			94				9
5 Rockey DC, Paulson	614	Air contrast barium enema,	98-99			99.6	15

E(2005)				CTcolonography,Colonoscopy			
6	HoppeH, P(2004)	Netzer	100	CT colonography,Colonoscopy	94		16
7	Rex DK, CS(1997)	Cutler	183	Back-to-back colonoscopies	87-94	76	17
8	Warneke J, N(1992)	Petrelli	235	Pathologic findings of the surgical specimen	96		18
9	Gelfand DW, MY(1992)	Chen	77	Barium enema examination		85	19
10	Byrd RL, Jr(1989)	Boggs HW	429	Postoperative pathologic specimen reports		97	20
11	Kalra L, AN(1988)	Hamlyn	154	Known cases	88-100		21
12	Walleser S, A(2007)	Griffiths	-		95	96	99.7-99.8
13	Cotton PB, VL(2004)	Durkalski	615	CT colonography,colonoscopy	99-100		100
14	Iannaccone R, Catalano C(2005)		88	Colonoscopy	84-90		100
15	Pickhardt PJ, JR(2003)	Choi	1233	CT colonography	87.5-92.3		25
16	Smith GA, PJ(2001)	O'Dwyer	1081	Follow up after 1-2 years	91.4	97.5	26
17	Rex DK, EY(1997)	Rahmani	2193	Medical records		95	27
18	Reiertsen O, A(1988)	Bakka	303	Known cases		90	28
19	Thoeni RF, A(1982)	Petras	53	Known cases		78	29
20	Fork FT(1981)		250	Known cases		91	30

Sigmoidoscopy

	Authors and year	Sample size	Gold standard	Sensitivity (percent)			Specificity	Ref. Number
				for cancer	for large polyps/ad enomas	over ally		
1	Soon MS, Kozarek RA(2005)	4859	Colonoscopy	67.5-79.3			31	
2	Kalra L, Hamlyn AN(1988)	154	Known cases	81-90			21	
3	Graser A, Stieber P(2009)	307	CT colonography colonoscopy			83.3	32	
4	Sung JJ, Chan FK(2003)	505	Colonoscopy			77.8	83.9	
5	Schoenfeld P, Lipscomb S(1999)	3028	Repeat sigmoidoscopy, performed by a gastroenterologist			79	34	
6	Saito Y, Slezak P(1989)	675	DCBE		88.8-100		35	
7	Dubow RA, Katon RM(1985)	258				98	36	
8	Farrands PA, Vellacott KD(1983)	227	Colonoscopy DCBE	94.2	98		37	
9	Hardcastle JD, Farrands	20525	Known cases		97.7		38	

PA(1983)							
10	Jensen J, Kewenter J(1990)	853	Follow up after 1-2 years			86	39
11	Rozen P, Ron E(1987)	1176	Colonoscopy	80	95	93.8	40
12	Jensen J, Kewenter J(1986)	458	DCBE	82			41

CT colonography

	Authors	Pub year	Sample size	Gold standard	Sensitivity (percent)			Specificity			Ref number
					for large polyps	for cancer	over ally	For large polyps	For cancer	over ally	
1	Rockey DC, Paulson E	2005	614	Air contrast barium enema,CT colonography, Colonoscopy	51-59			96			15
2	Hoppe H, Netzer P	2004	100sym ptomatic	CT colonography, Colonoscopy	61-95			88-98			16
3	Graser A, Stieber P	2009	307	CT colonography colonoscopy				96.7			32
4	Johnson CD, Toledano AY	2003	341	Colonoscopy and pathology reports				75		73	73
5	Walleser S, Griffiths A	2007	-	-	63	89		95	97		22
6	Chen SC, Lu DS	1999	23	Colonoscopy	95						74
7	Suzuki K, Yoshida H	2008	73	Colonoscopy		96.4					75
8	White TJ, Avery GR	2009	150 (High risk)	Colonoscopy	91	100				99.2	76
9	Selçuk D, Demirel K	2006	48(High risk)	Colonoscopy	85-100		86			98	77
10	Cotton PB, Durkalski VL	2004	615	CT colonography, colonoscopy	39-55			90.5-96			23
11	Ng CS, Doyle TC	2002	1031	Pathological and cancer registration records, together with colonoscopy +BE			85			91	78

12	Wong BC, Wong WM	2002	71	Colonoscopy	92	59	100	88	79
13	Laghi A, Iannaccone R	2002	66	Colonoscopy	84.6-92.8	93.7		94.1	80
14	Spinzi G, Belloni G	2001	99	Colonoscopy		57.8		92.6	81
15	Mendelson RM, Foster NM	2000	100	Colonoscopy	73				82
16	Miao YM, Amin Z	2000	201	Colonoscopy	73	100	94	99	83
17	Kay CL, Kulling D	2000	38	Colonoscopy	66.7-90		75-82.1		84
18	Fenlon HM, Nunes DP	1999	100	Colonoscopy	82-91		84	100	85
19	Beaulieu CF, Jeffrey RB Jr	1999	-	Panoramic endoscopic volume-rendered studies	68				86
20	Johnson CD, Chen MH	2008	2531	Colonoscopy and histologic review	78-84	90		86	87
21	Kim YS, Kim N	2008	241	Colonoscopy	60.4-86.7				88
22	Summers RM, Handwerker LR	2008	104	Colonoscopy	82.1-97.6				89
23	Juchems MS, Ernst A	2009	79	Colonoscopy	58.3-60.4			75	90
24	Jensch S, de Vries AH	2008	168	Colonoscopy	70-82		79-97		91
25	Johnson CD, Chen MH	2008	114	Colonoscopy	53-93			71-100	87
26	Roberts-Thomson IC, Tucker GR	2008	202	Colonoscopy	71	50	67	48	92
27	Yun JY, Roh HJ	2007	399	Colonoscopy	89-91				93
28	Arnesen RB, von Benzon E	2007	231	Colonoscopy	66-81		91-98		94
29	Baker ME, Bogoni L	2007	30	Computer-aided detection		81			95
30	Chaparro Sánchez M, del Campo Val L	2007	50	Colonoscopy	75-80		75-80	94	96

31	Sallam BM, Pilch-Kowalczyk A	2007	77	Colonoscopy and/or barium enema and pathomorphological examinations	100		98		97
32	Purkayastha S, Athanasio u T	2007	-	Colonoscopy			96		100 98
33	Kalra N, Suri S	2006	42	Colonoscopy	97-100		83-100		99
34	Reuterskiö Id MH, Lason A	2006	111	Colonoscopy	86-91	100	92	45	100
35	Dehmeshki J, Halligan S	2007	138	Computer assisted detection	87.8-95.2				101
36	Duff SE, Murray D	2006	112	Follow up after 1 years			87.5		97.1 102
37	Wessling J, Fischbach R	2006	-	Colon phantom	87-100				103
38	Juchems MS, Fleiter TR	2006	21	Colonoscopy	56.3-100				104
39	Summers RM, Yao J	2005	792	Colonoscopy	89.3				105
40	You YT, Chang Chien CR	2006	434	Colonoscopy, surgical finding, and clinical follow-up	100			83	106
41	Abdel Razek AA, Abu Zeid MM	2005	32	Colonoscopy			86.7		100 107
42	Iannaccone R, Catalano C	2005	88	Colonoscopy	84-86		62		82 24
43	Halligan S, Altman DG	2005	-	-	86-93		86-97		108
44	Wessling J, Domagk D	2005	78	Colonoscopy	81-100	100			86 109
45	Anupindi S, Perumpillichira J	2005	7	Colonoscopy	66.7				110
46	Mulhall BP, Veerappan GR	2005	-	Colonoscopy	70-85		93-97		111
47	Arnesen	2005	41	Colonoscopy	67-75		84-95		112

	RB, Adamsen S										
48	Park SH, Ha HK	2005	56	Colonoscopy	75-79						113
49	Iannaccon e R, Laghi A	2004	203	Colonoscopy	89.9-95.5				92.2		114
50	Pickhardt PJ, Choi JR	2004	1233	Colonoscopy	85.7-92.2						115
51	Van Gelder RE, Nio CY	2004	249	Colonoscopy	75-84			92			116
52	Johnson CD, MacCarty RL	2004	837	Colonoscopy +DCBE	56-79			96-99			117
53	Macari M, Bini EJ	2004	68	Colonoscopy	52.9-100			89.7			118
54	Pickhardt PJ, Choi JR	2003	1233	Colonoscopy	88.7-93.8			79.6-96			25
55	Iannaccon e R, Laghi A	2003	158	Colonoscopy	83.3-100	100	96		96.6		119
56	Sosna J, Morrin MM	2003	-	-	62-88			95			120
57	Munikrishn an V, Gillams AR	2003	80	Colonoscopy	83-100	97	74	98	96		121
58	Johnson CD, Harmsen WS	2003	703	Colonoscopy	43.8-50.5				86- 98		122
59	Gluecker T, Dorta G	2002	50	Colonoscopy	33-82				90		123
60	Yee J, Akerkar GA	2001	300	Colonoscopy	80.1-90	100	90.1		72		124
61	Spinzi G, Belloni G	2001	99	Colonoscopy				57.8	92.6		81
62	Britton I, Dover S	2001	50	Pathology, colonoscopy, barium enema ERCP and clinical follow-up	100	82		94			125
63	McFarlan d EG, Brink JA	2001	-	Colon phantom	61-100			89-92	-72 83		126
64	Civelli EM, Gallino G	2000	53	Known cases		100			78.7		127

Barium Enema

	Authors	Published year	Sam- ple size	Gold standard	Sensitivity (percent)		Specificity	Ref .Number
					For large polyps	For over cancer ally		
1	Rockey DC, Paulson E	2005	614	CT colonography, Colonoscopy	35-48		90	15
2	Kalra L, Hamlyn AN	1988	154	Known cases		79-81		21
3	Thorson AG, Christensen MA	1986	176	Colonoscopy	42-58	68-78		128
4	McPherson A, Payne JE	1983	108	Colonoscopy			90	57 129
5	Ribet A, Escourrou J	1980	603	FS			41	72
6	Wolff WI, Shinya H	1975	500	Colonoscopy	58	75		130
7	Saito Y, Slezak P	1989	675	Colonoscopy	64.7-83.3			35
8	Farrands PA, Vellacott KD	1983	227	Colonoscopy	74	76.5		37
9	Hardcastle JD, Farrands PA	1983	361 3	Known cases	62	75		38
10	Johnson CD, MacCarty RL	2004	837	Colonoscopy +CTC	39-56		99- 100	117
11	Sosna J, Sella T	2008	Meta anal ysis	-	70.2-71.5			131
12	Leslie A, Virjee JP	2002	70	Follow up after 5 years			93	132
13	Rockey DC, Koch J	2004	100	Colonoscopy	27-33		97- 100	133
14	Culpan DG, Mitchell AJ	2002	239	Follow up after 3 years		90.6		134
15	Connolly DJ, Traill ZC	2002	880	Follow up after 2 years		90.2	99. 5	135
16	Smith GA, O'Dwyer PJ	2001	138 9	Follow up after 1-2 years	21.7	83		26
17	Gillespie JS, Kelly BE	2001	160	Follow up after 2 years		96.5		136
18	Rex DK, Rahmani EY	1997	219 3	Medical records			81.8 - 85.2	27
19	Jaramillo E, Slezak P	1992	288	Colonoscopy	98	100		137
20	Jensen J, Kewenter J	1990	853	Follow up after 1-2 years	72			39
21	Ahovuo J, Linden H	1990	57	Colonoscopy	81			138
22	Reiertsen O, Bakka A	1988	303	Known cases			85	28
23	Myllylä V, Päivänsalo M	1984	112	Known cases and medidal records		82-83		139
24	Thoeni RF, Petras A.	1982	53	Known cases			88	29
25	Fork FT.	1981	250	Known cases			90	96.5 30
26	Jensen J, Kewenter J	1986	458	FS	77			41

Table 5. Range of sensitivity/specificity of screening methods obtained from systematic review of related articles

Type	Colonoscopy	Sigmoidoscopy	Barium enema	FOBT	CT Colonography
Sen ¹ for large polyps	74-100	82-100	21.7-98	4-41	33-100
Sen ¹ for cancer	80-100	67.5-94.2	68-100	5.6-100	82-100
Overall sen*	76-97	77.8-93.8	41-93	14.3-82	50-100
Spc ² for large polyps**	-	-	-	-	67-100
Spc ² for cancer**	-	-	-	-	97-100
Overall sSpc ²	99.6-100	83.9***	57-100	59.4-98	45-100

1. Sen: Sensitivity

2. Spc: Specificity

*Overall sensitivity does not mean the sum of other sensitivities. This term was exactly reported in some articles (refer to previous tables).

**For all methods except CT colonography specificity for large polyps, cancer and overall specificity are pooled together because of inadequacy of evidences and is explained as overall specificity.

***There was only one article that reported this value.

Table 6. Average sensitivity/specificity of screening methods \pm standard deviation obtained from systematic review of related articles

Type	Colonoscopy	Sigmoidoscopy	Barium enema	FOBT	CT colonography
Sen ¹ for large polyps	92.5 \pm 6.2	93.6 \pm 6.9	57.6 \pm 20.8	18.5 \pm 11.8	79.9 \pm 15.9
Sen ¹ for cancer	94.7 \pm 4.6	82.0 \pm 9.3	82.3 \pm 8.7	45.7 \pm 26.5	95.7 \pm 5.9
Overall sen*	84.9 \pm 8.5	86.3 \pm 8.1	81.75 \pm 16.8	37.0 \pm 19.4	80.1 \pm 15.6
Spc ² for large polyps**	-	-	-	-	90.1 \pm 8.3
Spc ¹ for cancer**	-	-	-	-	98.5 \pm 1.3
Overall Spc ²	99.8 \pm 0.2	83.9***	92.4 \pm 14.7	87.6 \pm 11.4	85.0 \pm 14.0

1. Sen: Sensitivity

2. Spc: Specificity

*Overall sensitivity does not mean the sum of other sensitivities. This term was exactly reported in some articles (refer to previous tables).

**For all methods except CT colonography specificity for large polyps, cancer and overall specificity are pooled together because of inadequacy of evidences and is explained as overall specificity.

***There was only one article that reported this value.

Conclusion

In overall all of the tests except stool exams have acceptable sensitivity and specificity for detecting colorectal cancer. Based on available evidences, colonoscopy has the best sensitivity and specificity among the selected screening methods and fecal occult blood test has the lowest sensitivity. Doing an analysis of cost effectiveness among these methods is encouraged.

Acknowledgement

This study was performed in Isfahan University of Medical Sciences and with kind assistance of Pursina Hakim Research Center (Gastrointestinal and Hepatic Research Center).

Conflict of Interest

The authors declare that they have no conflict of interest in this article.

Authors' Contribution

EMH designed the study and contributed to analyzing the data. DM contributed to designing the study, analyzing the data, writing the paper and literature review. AZ carried out the study, reviewed the literature, analyzed the data and wrote the paper. All authors read and approved the final manuscript.

References

1. Fauci AS. Harrison's principles of internal medicine. 17th ed. New York: McGraw-Hill Medical ; London : McGraw-Hill [distributor]; 2008.
2. Irvine EJ, Hunt RH. Evidence-based gastroenterology. Hamilton, Ont. ; London: BC Decker; 2001.
3. Feldman M, Friedman LS, Sleisenger MH. Sleisenger & Fordtran's gastrointestinal and liver disease : pathophysiology, diagnosis, management. 7th ed. / [edited by] Mark Feldman, Lawrence S. Friedman, Marvin H. Sleisenger. ed. Philadelphia ; London: Saunders; 2002.
4. Fletcher RH, Fletcher SW. Clinical epidemiology : the essentials. 4th ed. Baltimore, Md.: Lippincott Williams & Wilkins; 2005.
5. National cancer control programmes. Policies and managerial guide lines. 2nd Edition, Geneva: World health organization, 2002.
6. Dominitz JA, Eisen GM, Baron TH. Complications of colonoscopy. *Gastrointest Endosc* 2003;57(4):441-445.
7. Baxter NN, Goldwasser MA, Paszat LF, Saskin R, Urbach DR, Rabeneck L. Association of Colonoscopy and Death From Colorectal Cancer. *Annals of Internal Medicine* 2009;150(1):1-8.
8. Bielen DJLE, Bosmans HTC, Wever LLID. Clinical validation of high-resolution fast spin-echo MR colonography after colon distention with air. *Journal of Magnetic Resonance Imaging* 2005;22(3):400-5.
9. Menardo G. Sensitivity of diagnostic examinations for colorectal polyps. *Techniques in Coloproctology* 2004;8(0):s273-s5.
10. Yee J, Kumar NN, Godara S. Extracolonic Abnormalities Discovered Incidentally at CT Colonography in a Male Population. *Radiology* 2005;236(2):519-26.
11. Imperiale TF, Ransohoff DF, Itzkowitz SH, Turnbull BA, Ross ME. Fecal DNA versus fecal occult blood for colorectal-cancer screening in an average-risk population. *N Engl J Med* 2004;351(26):2704-14.
12. Heresbach D, Barrioz T, Lapalus MG, Coumaros D, Bauret P, Potier P, et al. Miss rate for colorectal neoplastic polyps: a prospective multicenter study of back-to-back video colonoscopies. *Endoscopy* 2008;40(4):284-90.
13. Hosokawa O, Hattori M, Douden K, Hayashi H, Ohta K, Kaizaki Y. Difference in accuracy between gastroscopy and colonoscopy for detection of cancer. *Hepatogastroenterology* 2007;54(74):442-4.
14. Bressler B, Paszat LF, Chen Z, Rothwell DM, Vinden C, Rabeneck L. Rates of new or missed colorectal cancers after colonoscopy and their risk factors: a population-based analysis. *Gastroenterology* 2007;132(1):96-102.
15. Rockey DC, Paulson E, Niedzwiecki D, Davis W, Bosworth HB, Sanders L, et al. Analysis of air contrast barium enema, computed tomographic colonography, and colonoscopy: prospective comparison. *Lancet*. 2005 Jan 22-28;365(9456):305-11.
16. Hoppe H, Netzer P, Spreng A, Quattropiani C, Mattich J, Dinkel HP. Prospective comparison of contrast enhanced CT colonography and conventional colonoscopy for detection of colorectal neoplasms in a single institutional study using second-look colonoscopy with discrepant results. *Am J Gastroenterol* 2004;99(10):1924-35.
17. Rex DK, Cutler CS, Lemmel GT, Rahmani EY, Clark DW, Helper DJ, et al. Colonoscopic miss rates of adenomas determined by back-to-back colonoscopies. *Gastroenterology* 1997;112(1):24-8.
18. Warneke J, Petrelli N, Herrera L, Nava H. Accuracy of colonoscopy for the detection of colorectal polyps. *Dis Colon Rectum* 1992;35(10):981-5.
19. Gelfand DW, Chen MY, Ott DJ. Benign colorectal neoplasms undetected by colonoscopy. *Gastrointest Radiol*. 1992 Fall;17(4):344-6.
20. Byrd RL, Boggs HW, Jr., Slagle GW, Cole PA. Reliability of colonoscopy. *Dis Colon Rectum* 1989;32(12):1023-1025.
21. Kalra L, Hamlyn AN. Comparative evaluation of investigations for colorectal carcinoma in symptomatic patients. *Postgrad Med J*. 1988 Sep;64(755):666-8.
22. Walleiser S, Griffiths A, Lord SJ, Howard K, Solomon MJ, Gebiski V. What is the value of computered tomography colonography in patients screening positive for fecal occult blood? A systematic review and economic

evaluation. *Clin Gastroenterol Hepatol* 2007;5(12):1439-1446; quiz 1368.

23. Cotton PB, Durkalski VL, Pineau BC, Palesch YY, Mauldin PD, Hoffman B, et al. Computed tomographic colonography (virtual colonoscopy): a multicenter comparison with standard colonoscopy for detection of colorectal neoplasia. *JAMA*. 2004 Apr 14;291(14):1713-9.

24. Iannaccone R, Catalano C, Mangiapane F, Murakami T, Lamazza A, Fiori E, et al. Colorectal polyps: detection with low-dose multi-detector row helical CT colonography versus two sequential colonoscopies. *Radiology* 2005;237(3):927-37.

25. Pickhardt PJ, Choi JR, Hwang I, Butler JA, Puckett ML, Hildebrandt HA, et al. Computed tomographic virtual colonoscopy to screen for colorectal neoplasia in asymptomatic adults. *N Engl J Med* 2003;349(23):2191-200.

26. Smith GA, O'Dwyer PJ. Sensitivity of double contrast barium enema and colonoscopy for the detection of colorectal neoplasms. *Surg Endosc* 2001;15(7):649-52.

27. Rex DK, Rahmani EY, Haseman JH, Lemmel GT, Kaster S, Buckley JS. Relative sensitivity of colonoscopy and barium enema for detection of colorectal cancer in clinical practice. *Gastroenterology*. 1997 Jan;112(1):17-23.

28. Reiertsen O, Bakka A, Tronnes S, Gauperaa T. Routine double contrast barium enema and fiberoptic colonoscopy in the diagnosis of colorectal carcinoma. *Acta Chir Scand* 1988;154(1):53-5.

29. Thoeni RF, Petras A. Double-contrast barium-enema examination and endoscopy in the detection of polypoid lesions in the cecum and ascending colon. *Radiology* 1982;144(2):257-260.

30. Fork FT. Double contrast enema and colonoscopy in polyp detection. *Gut* 1981;22(11):971-7.

31. Soon MS, Kozarek RA, AyubK, Soon A, Lin TY, Lin OS. Screening colonoscopy in Chinese and Western patients: a comparative study. *Am J Gastroenterol* 2005;100(12):2749-55.

32. Graser A, Stieber P, Nagel D, Schäfer C, Horst D, Becker CR, et al. Comparison of CT colonography, colonoscopy, sigmoidoscopy and faecal occult blood tests for the detection of advanced adenoma in an average risk population. *Gut* 2009;58(2):241-8.

33. Sung JJ, Chan FK, Leung WK, Wu JC, Lau JY, Ching J, et al. Screening for colorectal cancer in Chinese: Comparison of fecal occult blood test, flexible sigmoidoscopy, and colonoscopy. *Gastroenterology* 2003;124(3):608-14.

34. Schoenfeld P, Lipscomb S, Crook J, Dominguez J, Butler J, Holmes L, et al. Accuracy of polyp detection by gastroenterologists and nurse endoscopists during flexible sigmoidoscopy: a randomized trial. *Gastroenterology*. 1999 Aug;117(2):312-8.

35. Saito Y, Slezak P, Rubio C. The diagnostic value of combining flexible sigmoidoscopy and double-contrast barium enema as a one-stage procedure. *Gastrointest Radiol* 1989; 14(4):357-9.

36. Dubow RA, Katon RM, Benner KG. Short (35-cm) versus long (60-cm) flexible sigmoidoscopy: A comparison of findings and tolerance in asymptomatic patients screened for colorectal neoplasia. *Gastrointestinal Endoscopy* 1985;31(5):305-8.

37. Farrands PA, Vellacott KD, Amar SS, Balfour TW, Hardcastle JD. Flexible fiberoptic sigmoidoscopy and double-contrast barium-enema examination in the identification of adenomas and carcinoma of the colon. *Dis Colon Rectum* 1983; 26(11):725-7.

38. Hardcastle JD, Farrands PA, Balfour TW, Chamberlain J, Amar SS, SheldonMG. Controlled trial of faecal occult blood testing in the detection of colorectal cancer. *Lancet* 1983;2(8340):1-4.

39. Jensen J, Kewenter J, Asztely M, Lycke G, Wojciechowski J. Double contrast barium enema and flexible rectosigmoidoscopy: a reliable diagnostic combination for detection of colorectal neoplasm. *Br J Surg* 1990;77(3):270-2.

40. Rozen P, Ron E, Fireman Z, Hallak A, Grossman A, Baratz M, et al. The relative value of fecal occult blood tests and flexible sigmoidoscopy in screening for large bowel neoplasia. *Cancer*. 1987 Nov 15;60(10):2553-8.

41. Jensen J, Kewenter J, Haglund E, Lycke G, Svensson C, Ahrén C. Diagnostic accuracy of double-contrast enema and rectosigmoidoscopy in connection with faecal occult blood testing for the detection of rectosigmoid neoplasms. *Br J Surg*. 1986 Dec;73(12):961-4.

42. Cheng TI, Wong JM, Hong CF, Cheng SH, Cheng TJ, Shieh MJ, et al. Colorectal cancer screening in asymptomatic adults: comparison of colonoscopy, sigmoidoscopy and fecal occult blood tests. *J Formos Med Assoc* 2002;101(10):685-90.

43. Brevinge H, Lindholm E, Buntzen S, Kewenter J. Screening for colorectal neoplasia with faecal occult blood testing compared with flexible sigmoidoscopy directly in a 55-56 years' old population. *Int J Colorectal Dis* 1997;12(5):291-5.

44. Bang KM, Tillett S, Hoar SK, Blair A, McDougall V. Sensitivity of fecal hemoccult testing and flexible sigmoidoscopy for colorectal cancer screening. *J Occup Med* 1986; 28(8):709-13.

45. Demers RY, Stawick LE, Demers P. Relative sensitivity of the fecal occult blood test and flexible sigmoidoscopy in detecting polyps. *Prev Med* 1985;14(1):55-62.

46. Ahlquist DA, Sargent DJ, Loprinzi CL, Levin TR, Rex DK, Ahnen DJ, et al. Stool DNA and occult blood testing for screen detection of colorectal neoplasia. *Ann Intern Med* 2008; 149(7):441-50, W481.

47. Collins JF, Lieberman DA, Durbin TE, Weiss DG. Accuracy of screening for fecal occult blood on a single stool sample obtained by digital rectal examination: a comparison with recommended sampling practice. *Ann Intern Med* 2005;142(2):81-5.

48. Kristinsson J, Nygaard K, Aadland E, Barstad S, Sauar J, Hofstad B, et al. Screening of first degree relatives of patients operated for colorectal cancer: evaluation of fecal calprotectin vs. hemoccult II. *Digestion* 2001;64(2):104-10.

49. Tibble J, Sigthorsson G, Foster R, Sherwood R, Fagerhol M, Bjarnason I. Faecal calprotectin and faecal occult blood tests in the diagnosis of colorectal carcinoma and adenoma. *Gut* 2001;49(3):402-8.

50. Olynyk JK, Platell CF, Collett JA. Fecal occult blood and flexible sigmoidoscopy screening for colorectal cancer: modeling the impact on colonoscopy requirements and cancer detection rates. *J Gastroenterol Hepatol* 2001;16(4):389-92.

51. Launoy G, Smith TC, Duffy SW, Bouvier V. Colorectal cancer mass-screening: estimation of faecal occult blood test sensitivity, taking into account cancer mean sojourn time. *Int J Cancer* 1997;73(2):220-4.

52. Nakama H, Kamijo N, Fujimori K, Horiuchi A, Fattah AS, Zhang B. Clinical diagnostic accuracy of faecal occult blood test for anal diseases. *Int J Qual Health Care* 1997;9(2):139-41.

53. Hope RL, Chu G, Hope AH, Newcombe RG, Gillespie PE, Williams SJ. Comparison of three faecal occult blood tests in the detection of colorectal neoplasia. *Gut* 1996;39(5):722-5.

54. Allison JE, Tekawa IS, Ransom LJ, Adrain AL. A comparison of fecal occult-blood tests for colorectal-cancer screening. *N Engl J Med* 1996;334(3):155-9.

55. Niv Y, Sperber AD. Sensitivity, specificity, and predictive value of fecal occult blood testing (Hemoccult II) for colorectal neoplasia in symptomatic patients: a prospective study with total colonoscopy. *Am J Gastroenterol* 1995;90(11):1974-7.

56. Jeanson A, Jamart J, Maisin JM, Vanheuverzwyn R, Gohy P, Debongnie JC, et al. Assessment of the new immunological test Hemoblot for detecting occult blood in faeces. *Eur J Cancer Prev* 1994;3(5):407-12.

57. Jahn H, Joergensen OD, Kronborg O, Fenger C. Can Hemoccult-II replace colonoscopy in surveillance after radical surgery for colorectal cancer and after polypectomy? *Dis Colon Rectum* 1992;35(3):253-6.

58. Ahlquist DA, Wieand HS, Moertel CG, McGill DB, Loprinzi CL, O'Connell MJ, et al. Accuracy of fecal occult blood screening for colorectal neoplasia. A prospective study using Hemoccult and HemoQuant tests. *JAMA* 1993;269(10):1262-7.

59. Murakami R, Otani T, Nakanishi K, Fudemoto Y, Ishikawa H, Hiyama T, et al. Diagnostic validity of fecal occult blood tests for detecting gastroenterological cancers. *Jpn J Cancer Res* 1992;83(2):141-5.

60. Favennec L, Kapel N, Meillet D, Chochillon C, Gobert JG. Detection of occult blood in stools: comparison of three guaiac tests and a latex agglutination test. *Ann Biol Clin (Paris)* 1992;50(5):311-13.

61. Bertario L, Spinelli P, Gennari L, Sala P, Pizzetti P, Severini A, et al. Sensitivity of Hemoccult test for large bowel cancer in high-risk subjects. *Dig Dis Sci* 1988;33(5):609-613.

62. Letsou G, Ballantyne GH, Zdon MJ, Zucker KA, Modlin IM. Screening for colorectal neoplasms. A comparison of the fecal occult blood test and endoscopic examination. *Dis Colon Rectum* 1987;30(11):839-43.

63. Poleski MH, Gordon PH. Sensitivity of the Hemoccult II slide test in detecting colonic neoplasms. *Can J Surg* 1986;29(2):99-101.

64. Adamsen S, Kronborg O, Hage E, Fenger C. Reproducibility and diagnostic value of Hemoccult-II test. A colonoscopic evaluation in asymptomatic patients. *Scand J Gastroenterol* 1985;20(9):1073-7.

65. Farrands PA, O'Regan D, Taylor I. An assessment of occult blood testing to determine which patients with large bowel symptoms require urgent investigation. *Br J Surg* 1985;72(10):835-7.

66. Saito H, Tsuchida S, Nakaji S, Kakizaki R, Aisawa T, Munakata A, et al. An immunologic test for fecal occult blood by counter immunoelectrophoresis. Higher sensitivity and higher positive reactions in colorectal cancer than single radial immunodiffusion and hemoccult test. *Cancer* 1985;56(7):1549-52.

67. Adlercreutz H, Partanen P, Virkola P, Liewendahl K, Turunen MJ. Five guaiac-based tests for occult blood in faeces compared invitro and in vivo. *Scand J Clin Lab Invest* 1984;44(6):519-28.

68. Crowley ML, Freeman LD, Mottet MD, Strong RM, Sweeney BF, Brower RA, et al. Sensitivity of guaiac-impregnated cards for the detection of colorectal neoplasia. *J Clin Gastroenterol* 1983;5(2):127-30.

69. Rozen P, Levi Z, Hazazi R, Waked A, Vilkin A, Maoz E, et al. Quantitative colonoscopic evaluation of relative efficiencies of an immunochemical faecal occult blood test and a sensitive guaiac test for detecting significant colorectal neoplasms. *Aliment Pharmacol Ther* 2009;29(4):450-7.

70. St John DJ, Young GP, Alexeyeff MA, Deacon MC, Cuthbertson AM, Macrae FA, et al. Evaluation of new occult blood tests for detection of colorectal neoplasia. *Gastroenterology* 1993;104(6):1661-8.

71. Zou H, Taylor WR, Harrington JJ, Hussain FT, Cao X, Loprinzi CL, et al. High detection rates of colorectal neoplasia by stool DNA testing with a novel digital melt curve assay. *Gastroenterology* 2009;136(2):459-70.

72. Ribet A, Escourrou J, Frexinos J, Delpu J. Screening for colorectal tumors - results of two years experience. *Cancer Detect Prev* 1980;3(2):449-61.

73. Johnson CD, Toledano AY, Herman BA, Dachman AH, McFarland EG, Barish MA, et al. Computerized tomographic colonography: performance evaluation in a retrospective multicenter setting. *Gastroenterology* 2003;125(3):688-95.

74. Chen SC, Lu DS, Hecht JR, Kadell BM. CT colonography: value of scanning in both the supine and prone positions. *AJR Am J Roentgenol* 1999;172(3):595-9.

75. Suzuki K, Yoshida H, Nappi J, Armato SG, 3rd, Dachman AH. Mixture of expert 3D massive-training ANNs for reduction of multiple types of false positives in CAD for detection of polyps in CT colonography. *Med Phys* 2008;35(2):694-703.

76. White TJ, Avery GR, Kennan N, Syed AM, Hartley JE, Monson JR. Virtual colonoscopy vs conventional colonoscopy in patients at high risk of colorectal cancer--a prospective trial of 150 patients. *Colorectal Dis* 2009;11(2):138-45.

77. Selçuk D, Demirel K, Ozer H, Baca B, Hatemi I, Mihmanli I, et al. Comparison of virtual colonoscopy with conventional colonoscopy in detection of colorectal polyps. *Turk J Gastroenterol* 2006;17(4):288-93.
78. Ng CS, Doyle TC, Pinto EM, Courtney HM, Bull RK, Prevost AT, et al. Evaluation of CT in identifying colorectal carcinoma in the frail and disabled patient. *Eur Radiol* 2002;12(12):2988-97.
79. Wong BC, Wong WM, Chan JK, Lai KC, Hu WH, Chan CK, et al. Virtual colonoscopy for the detection of colorectal polyps and cancers in a Chinese population. *J Gastroenterol Hepatol* 2002;17(12):1323-7.
80. Laghi A, Iannaccone R, Carbone I, Catalano C, Panebianco V, Di Giulio E, et al. Computed tomographic colonography (virtual colonoscopy): blinded prospective comparison with conventional colonoscopy for the detection of colorectal neoplasia. *Endoscopy*. 2002 Jun;34(6):441-6.
81. Spinzi G, Belloni G, Martegani A, Sangiovanni A, Del Favero C, Minoli G. Computed tomographic colonography and conventional colonoscopy for colon diseases: a prospective, blinded study. *Am J Gastroenterol* 2001;96(2):394-400.
82. Mendelson RM, Foster NM, Edwards JT, Wood CJ, Rosenberg MS, Forbes GM. Virtual colonoscopy compared with conventional colonoscopy: a developing technology. *Med J Aust* 2000;173(9):472-5.
83. Miao YM, Amin Z, Healy J, Burn P, Murugan N, Westaby D, et al. A prospective single centre study comparing computed tomography pneumocolon against colonoscopy in the detection of colorectal neoplasms. *Gut* 2000;47(6):832-7.
84. Kay CL, Kulling D, Hawes RH, Young JW, Cotton PB. Virtual endoscopy--comparison with colonoscopy in the detection of space-occupying lesions of the colon. *Endoscopy* 2000;32(3):226-32.
85. Fenlon HM, Nunes DP, Schroy PC 3rd, Barish MA, Clarke PD, Ferrucci JT. A comparison of virtual and conventional colonoscopy for the detection of colorectal polyps. *N Engl J Med*. 1999 Nov 11;341(20):1496-503.
86. Beaulieu CF, Jeffrey RB, Jr., Karadi C, Paik DS, Napel S. Display modes for CT colonography. Part II. Blinded comparison of axial CT and virtual endoscopic and panoramic endoscopic volume-rendered studies. *Radiology* 1999;212(1):203-12.
87. Johnson CD, Chen MH, Toledano AY, Heiken JP, Dachman A, Kuo MD, et al. Accuracy of CT colonography for detection of large adenomas and cancers. *N Engl J Med* 2008;359(12):1207-17.
88. Kim YS, Kim N, Kim SH, Park MJ, Lim SH, Yim JY, et al. The efficacy of intravenous contrast-enhanced 16-row multidetector CT colonography for detecting patients with colorectal polyps in an asymptomatic population in Korea. *J Clin Gastroenterol* 2008;42(7):791-8.
89. Summers RM, Handwerker LR, Pickhardt PJ, Van Uitert RL, Deshpande KK, Yeshwant S, et al. Performance of a previously validated CT colonography computer-aided detection system in a new patient population. *AJR Am J Roentgenol* 2008;191(1):168-74.
90. Juchems MS, Ernst A, Johnson P, Virmani S, Brambs HJ, Aschoff AJ. Electronic colon-cleansing for CT colonography: diagnostic performance. *Abdom Imaging*. 2008 Mar 15. [Epub ahead of print]
91. Jensch S, de Vries AH, Peringa J, Bipat S, Dekker E, Baak LC, et al. CT colonography with limited bowel preparation: performance characteristics in an increased-risk population. *Radiology* 2008;247(1):122-32.
92. Roberts-Thomson IC, Tucker GR, Hewett PJ, Cheung P, Sebben RA, Khoo EE, et al. Single-center study comparing computed tomography colonography with conventional colonoscopy. *World J Gastroenterol* 2008;14(3):469-3.
93. Yun JY, Ro HJ, Park JB, Choi JB, Chung JE, Kim YJ, et al. Diagnostic performance of CT colonography for the detection of colorectal polyps. *Korean J Radiol* 2007;8(6):484-91.
94. Arnesen RB, von Benzon E, Adamsen S, Svendsen LB, Raaschou HO, Hansen OH. Diagnostic performance of computed tomography colonography and colonoscopy: a prospective and validated analysis of 231 paired examinations. *Acta Radiol* 2007;48(8):831-7.
95. Baker ME, Bogoni L, Obuchowski NA, Dass C, Kendzierski RM, Remer EM, et al. Computer-aided detection of colorectal polyps: can it improve sensitivity of less-experienced readers? Preliminary findings. *Radiology* 2007;245(1):140-9.
96. Chaparro Sánchez M, del Campo Val L, Maté Jiménez J, Cantero Perona J, Barbosa A Olivares D, et al. Computed tomography colonography compared with conventional colonoscopy for the detection of colorectal polyps. *Gastroenterol Hepatol* 2007;30(7):375-80.
97. Sallam BM, Pilch-Kowalczyk A, Gruszczynska K, Baron J, Pugliese F. Diagnostic performance of CT colonography in a population with high prevalence of large bowel disease. *Med Sci Monit* 2007;13 Suppl 1:105-10.
98. Purkayastha S, Athanasiou T, Tekkis PP, Constantinides V, Teare J, Darzi AW. Magnetic resonance colonography vs computed tomography colonography for the diagnosis of colorectal cancer: an indirect comparison. *Colorectal Dis* 2007;9(2):100-11.
99. Kalra N, Suri S, Bhasin DK, Sinha SK, Saravanan N, Kour T, et al. Comparison of multidetector computed tomographic colonography and conventional colonoscopy for detection of colorectal polyps and cancer. *Indian J Gastroenterol* 2006;25(5):229-32.
100. Reuterskiöld MH, Lasso A, Svensson E, Kilander A, Stotzer PO, Hellström M. Diagnostic performance of computed tomography colonography in symptomatic patients and in patients with increased risk for colorectal disease. *Acta Radiol* 2006;47(9):888-98.
101. Dehmeshki J, Halligan S, Taylor SA, Roddie ME, McQuillan J, Honeyfield L, et al. Computer assisted detection software for CT colonography: effect of sphericity filter on performance characteristics for patients with and without fecal tagging. *Eur Radiol* 2007;17(3):662-8.

102. Duff SE, Murray D, Rate AJ, Richards DM, Kumar NA. Computed tomographic colonography (CTC) performance: one-year clinical follow-up. *Clin Radiol*. 2006 Nov;61(11):932-6.
103. Wessling J, Fischbach R, Borchert A, Kugel H, Allkemper T, Osada N et al. Detection of colorectal polyps: comparison of multi-detector row CT and MR colonography in a colon phantom. *Radiology* 2006;241(1):125-31.
104. Juchems MS, Fleiter TR, Pauls S, Schmidt SA, Brambs HJ, Aschoff AJ. CT colonography: comparison of a colon dissection display versus 3D endoluminal view for the detection of polyps. *Eur Radiol* 2006;16(1):68-72.
105. Summers RM, Yao J, Pickhardt PJ, Franaszek M, Bitter I, Brickman D, et al. Computed tomographic virtual colonoscopy computer-aided polyp detection in a screening population. *Gastroenterology* 2005;129(6):1832-44.
106. You YT, Chang Chien CR, Wang JY, Ng KK, Chen JS, Tang R, et al. Evaluation of contrast-enhanced computed tomographic colonography in detection of local recurrent colorectal cancer. *World J Gastroenterol* 2006;12(1):123-6.
107. Abdel Razek AA, Abu Zeid MM, Bilal M, Abdel Wahab NM. Virtual CT colonoscopy versus conventional colonoscopy: a prospective study. *Hepatogastroenterology* 2005;52(66):1698-702.
108. Halligan S, Altman DG, Taylor SA, Mallett S, Deeks JJ, Bartram CI, et al. CT colonography in the detection of colorectal polyps and cancer: systematic review, meta-analysis, and proposed minimum data set for study level reporting. *Radiology* 2005;237(3):893-904.
109. Wessling J, Domagk D, Lugering N, Schierhorn S, Heindel W, Domschke W, et al. Virtual colonography: identification and differentiation of colorectal lesions using multi-detector computed tomography. *Scand J Gastroenterol* 2005;40(4):468-76.
110. Anupindi S, Perumpillichira J, Jaramillo D, Zalis ME, Israel EJ. Low-dose CT colonography in children: initial experience, technical feasibility, and utility. *Pediatr Radiol* 2005;35(5):518-24.
111. Mulhall BP, Veerappan GR, Jackson JL. Meta-analysis: computed tomographic colonography. *Ann Intern Med* 2005;142(8):635-50.
112. Arnesen RB, Adamsen S, Svendsen LB, Raaschou HO, von Benzon E, Hansen OH. Missed lesions and false-positive findings on computed-tomographic colonography: a controlled prospective analysis. *Endoscopy* 2005;37(10):937-44.
113. Park SH, Ha HK, Kim MJ, Kim KW, Kim AY, Yang DH, et al. False-negative results at multi-detector row CT colonography: multivariate analysis of causes for missed lesions. *Radiology* 2005;235(2):495-502.
114. Iannaccone R, Laghi A, Catalano C, Mangiapane F, Lamazza A, Schillaci A, et al. Computed tomographic colonography without cathartic preparation for the detection of colorectal polyps. *Gastroenterology* 2004;127(5):1300-11.
115. Pickhardt PJ, Choi JR, Hwang I, Schindler WR. Nonadenomatous polyps at CT colonography: prevalence, size distribution, and detection rates. *Radiology* 2004;232(3):784-90.
116. Van Gelder RE, Nio CY, Florie J, Bartelsman JF, Snel P, De Jager SW et al. Computed tomographic colonography compared with colonoscopy in patients at increased risk for colorectal cancer. *Gastroenterology* 2004;127(1):41-8.
117. Johnson CD, MacCarty RL, Welch TJ, Wilson LA, Harmsen WS, Ilstrup DM et al. Comparison of the relative sensitivity of CT colonography and double-contrast barium enema for screen detection of colorectal polyps. *Clin Gastroenterol Hepatol* 2004;2(4):314-21.
118. Macari M, Bini EJ, Jacobs SL, Naik S, Lui YW, Milano A et al. Colorectal polyps and cancers in asymptomatic average-risk patients: evaluation with CT colonography. *Radiology* 2004; 230(3):629-36.
119. Iannaccone R, Laghi A, Catalano C, Brink JA, Mangiapane F, Trenna S et al. Detection of colorectal lesions: lower-dose multi-detector row helical CT colonography compared with conventional colonoscopy. *Radiology*. 2003 Dec;229(3):775-81.
120. Sosna J, Morrin MM, Kruskal JB, Lavin PT, Rosen MP, Raptopoulos V. CT colonography of colorectal polyps: a metaanalysis. *AJR Am J Roentgenol* 2003;181(6):1593-8.
121. Munikrishnan V, Gillams AR, Lees WR, Vaizey CJ, Boulos PB. Prospective study comparing multislice CT colonography with colonoscopy in the detection of colorectal cancer and polyps. *Dis Colon Rectum* 2003;46(10):1384-90.
122. Johnson CD, Harmsen WS, Wilson LA, MacCarty RL, Welch TJ, Ilstrup DM et al. Prospective blinded evaluation of computed tomographic colonography for screen detection of colorectal polyps. *Gastroenterology* 2003;125(2):311-9.
123. Gluecker T, Dorta G, Keller W, Jornod P, Meuli R, Schnyder P. Performance of multidetector computed tomography colonography compared with conventional colonoscopy. *Gut* 2002;51(2):207-11.
124. Yee J, Akerkar GA, Hung RK, Steinauer-Gebauer AM, Wall SD, McQuaid KR. Colorectal neoplasia: performance characteristics of CT colonography for detection in 300 patients. *Radiology* 2001;219(3):685-92.
125. Britton I, Dover S, Vallance R. Immediate CT pneumocolon for failed colonoscopy; comparison with routine pneumocolon. *Clin Radiol* 2001;56(2):89-93.
126. McFarland EG, Brink JA, Pilgram TK, Heiken JP, Balfe DM, Hirselj DA et al. Spiral CT colonography: reader agreement and diagnostic performance with two- and three-dimensional image-display techniques. *Radiology* 2001;218(2):375-83.
127. Civelli EM, Gallino G, Mariani L, Cozzi G, Biganzoli E, Salvetti M et al. Double-contrast barium enema and computerised tomography in the pre-operative evaluation of rectal carcinoma: are they still useful diagnostic procedures? *Tumori* 2000;86(5):389-92.

128. Thorson AG, Christensen MA, Davis SJ. The role of colonoscopy in the assessment of patients with colorectal cancer. *Dis Colon Rectum* 1986;29(5):306-11.
129. McPherson A, Payne JE. Importance of total colonoscopy in the diagnosis of colonic disorders. *Med J Aust* 1983;1(4):170-2.
130. Wolff WI, Shinya H, Geffen A, Ozoktay S, DeBeer R. Comparison of colonoscopy and the contrast enema in five hundred patients with colorectal disease. *Am J Surg* 1975;129(2):181-6.
131. Sosna J, Sella T, Sy O, Lavin PT, Eliahou R, Fraifeld S et al. Critical analysis of the performance of double-contrast barium enema for detecting colorectal polyps $>$ or $=$ 6 mm in the era of CT colonography. *AJR Am J Roentgenol* 2008;190(2):374-85.
132. Leslie A, Virjee JP. Detection of colorectal carcinoma on double contrast barium enema when double reporting is routinely performed: an audit of current practice. *Clin Radiol* 2002;57(3):184-7.
133. Rockey DC, Koch J, Yee J, McQuaid KR, Halvorsen RA. Prospective comparison of air-contrast barium enema and colonoscopy in patients with fecal occult blood: a pilot study. *Gastrointest Endosc* 2004;60(6):953-8.
134. Culpan DG, Mitchell AJ, Hughes S, Nutman M, Chapman AH. Double contrast barium enema sensitivity: a comparison of studies by radiographers and radiologists. *Clin Radiol* 2002;57(7):604-7.
135. Connolly DJ, Traill ZC, Reid HS, Copley SJ, Nolan DJ. The double contrast barium enema: a retrospective single centre audit of the detection of colorectal carcinomas. *Clin Radiol* 2002;57(1):29-32.
136. Gillespie JS, Kelly BE. Double contrast barium enema and colorectal carcinoma: sensitivity and potential role in screening. *Ulster Med J* 2001;70(1):15-8.
137. Jaramillo E, Slezak P. Comparison between double-contrast barium enema and colonoscopy to investigate lower gastrointestinal bleeding. *Gastrointest Radiol* 1992;17(1):81-3.
138. Ahovuo J, Linden H, Kinnunen J, Edgren J, Kellokumpu I, Husa A. Double-contrast barium examination and endoscopy in the detection of small polyps of the large intestine. *Ann Chir Gynaecol* 1990;79(3):143-6.
139. Myllyla V, Paivansalo M, Laitinen S. Sensitivity of single and double contrast barium enema in the detection of colorectal carcinoma. *Rofo* 1984;140(4):393-7.