

# Screening for Colorectal Cancer

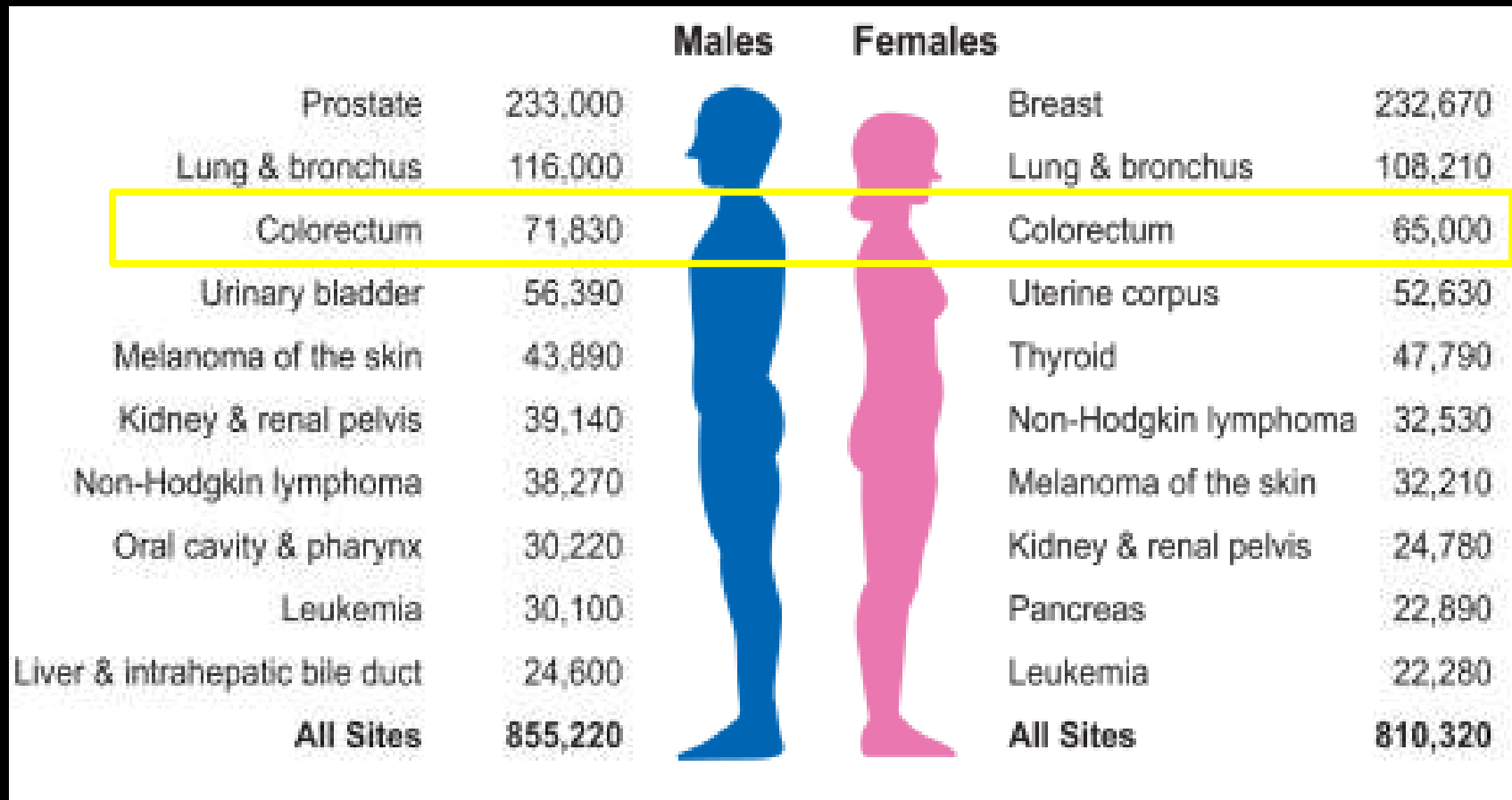
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**Bruce D. Greenwald, MD**

Professor of Medicine

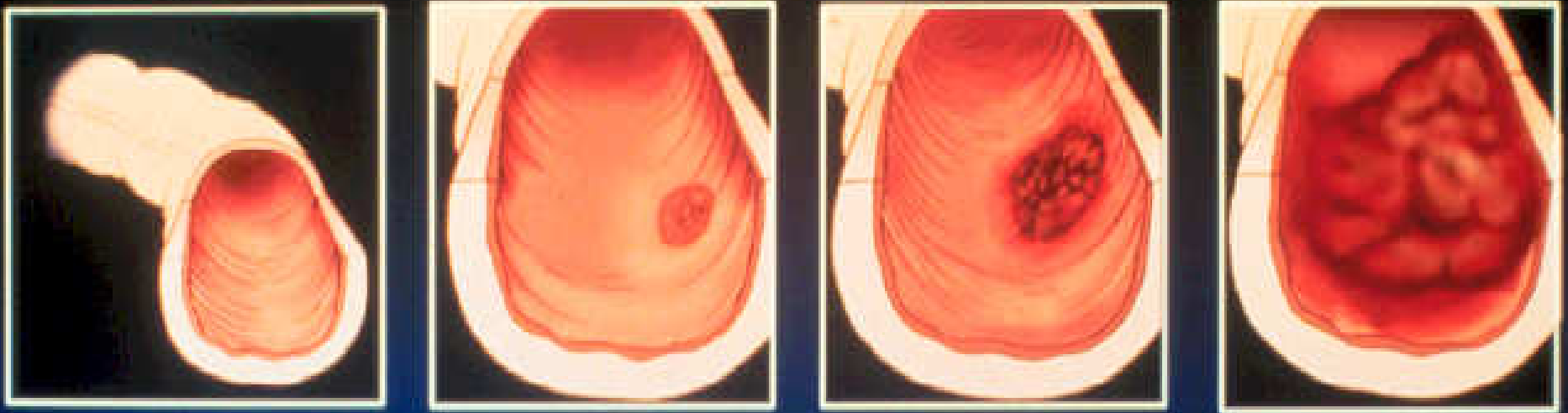
University of Maryland School of Medicine and  
Greenebaum Cancer Center

# 2014 Estimated US Cancer Cases



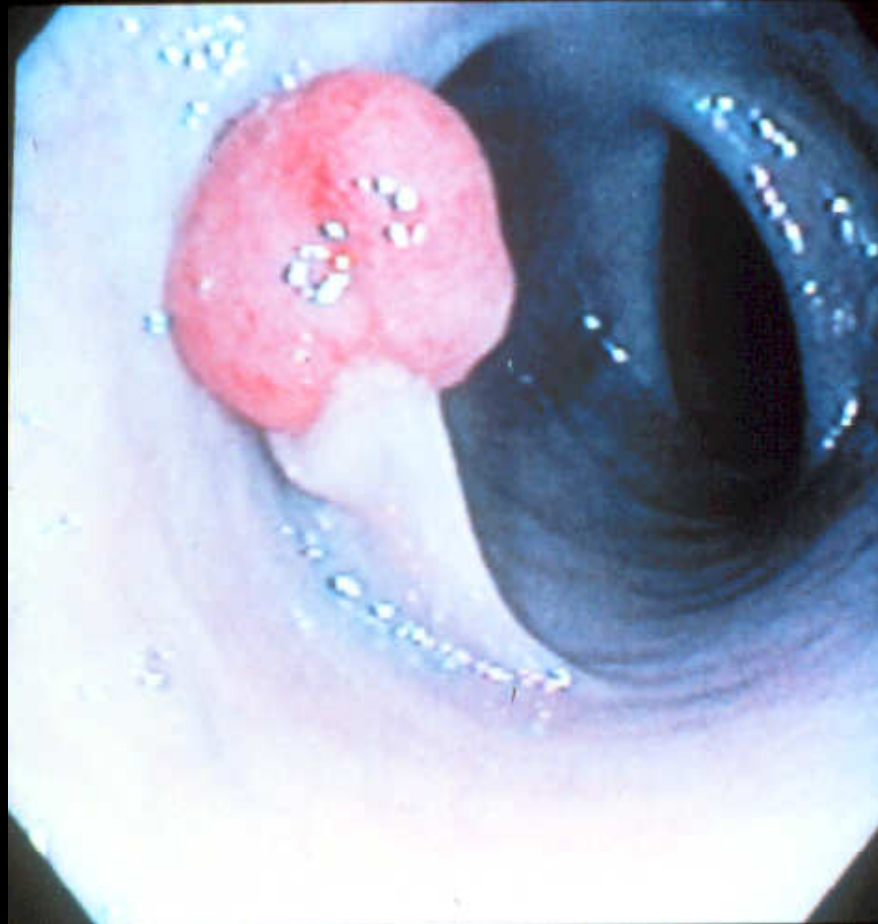
Siegel et al. CA: A cancer journal for clinicians; 2014; 64:9

# Adenoma – Carcinoma Sequence

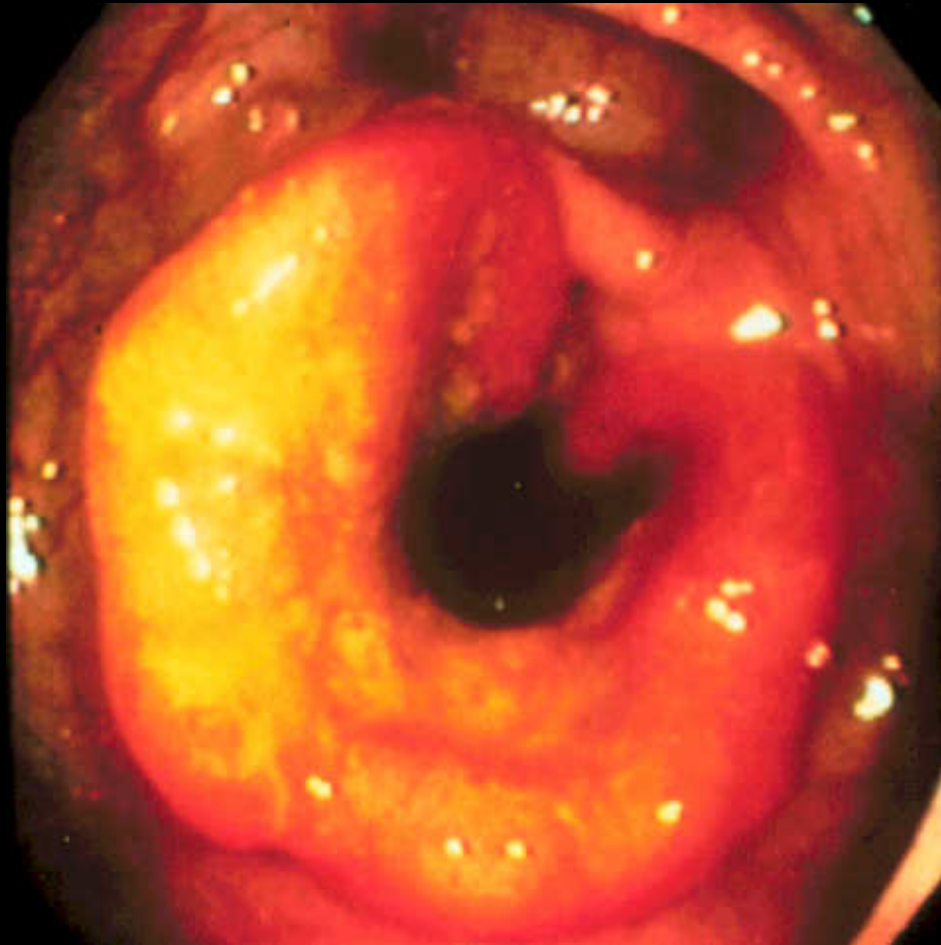


**Normal Mucosa** → **Adenoma** → **Severe Dysplasia** → **Cancer**

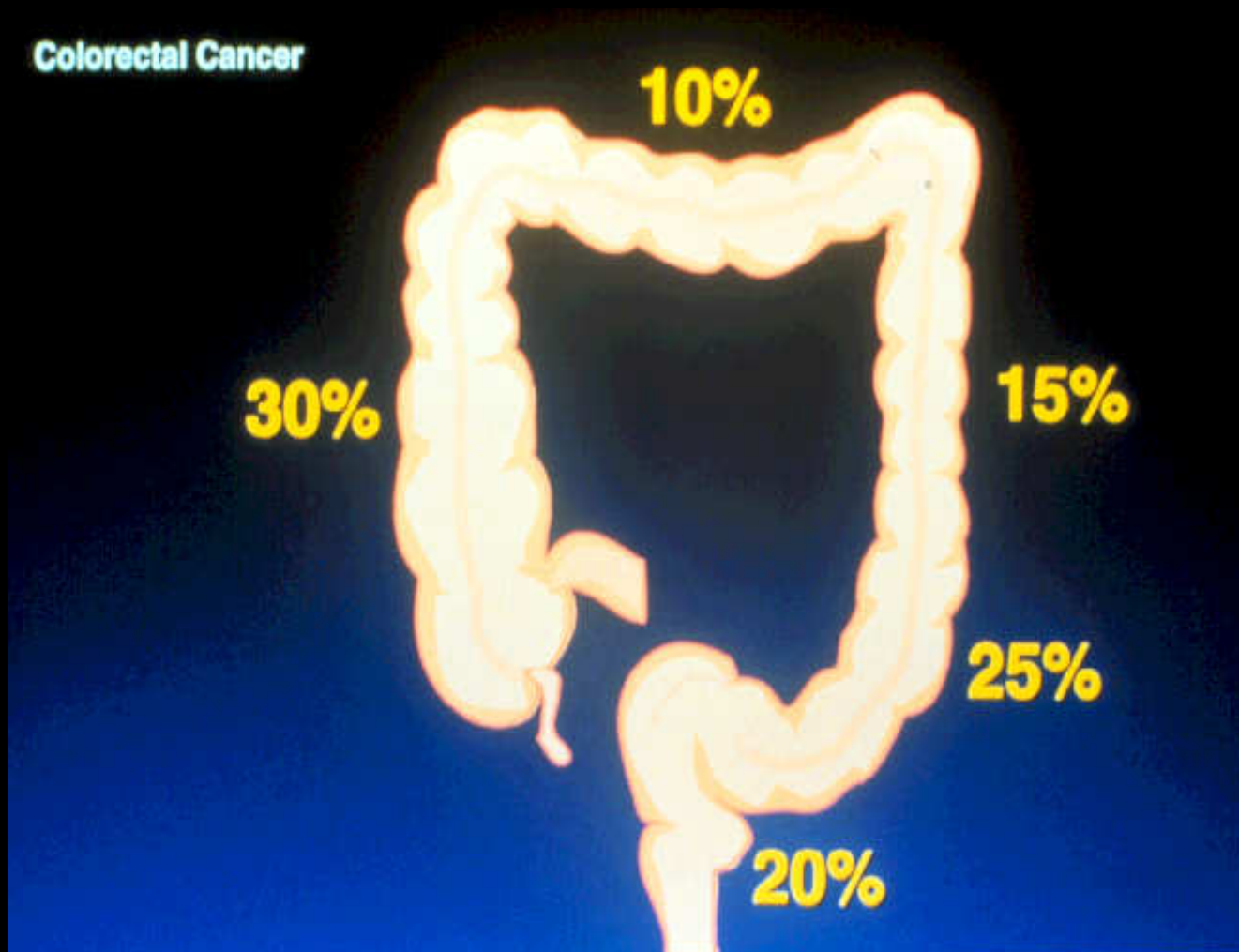
# Colon Polyp



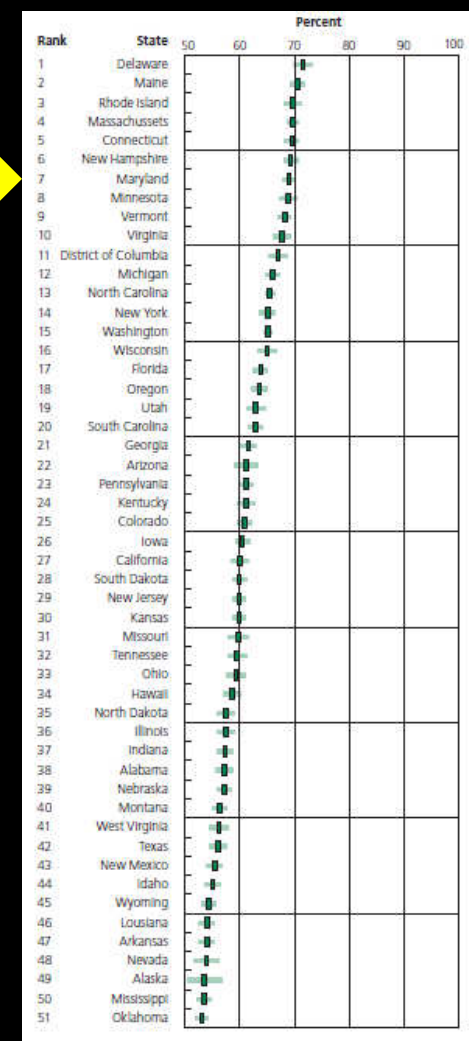
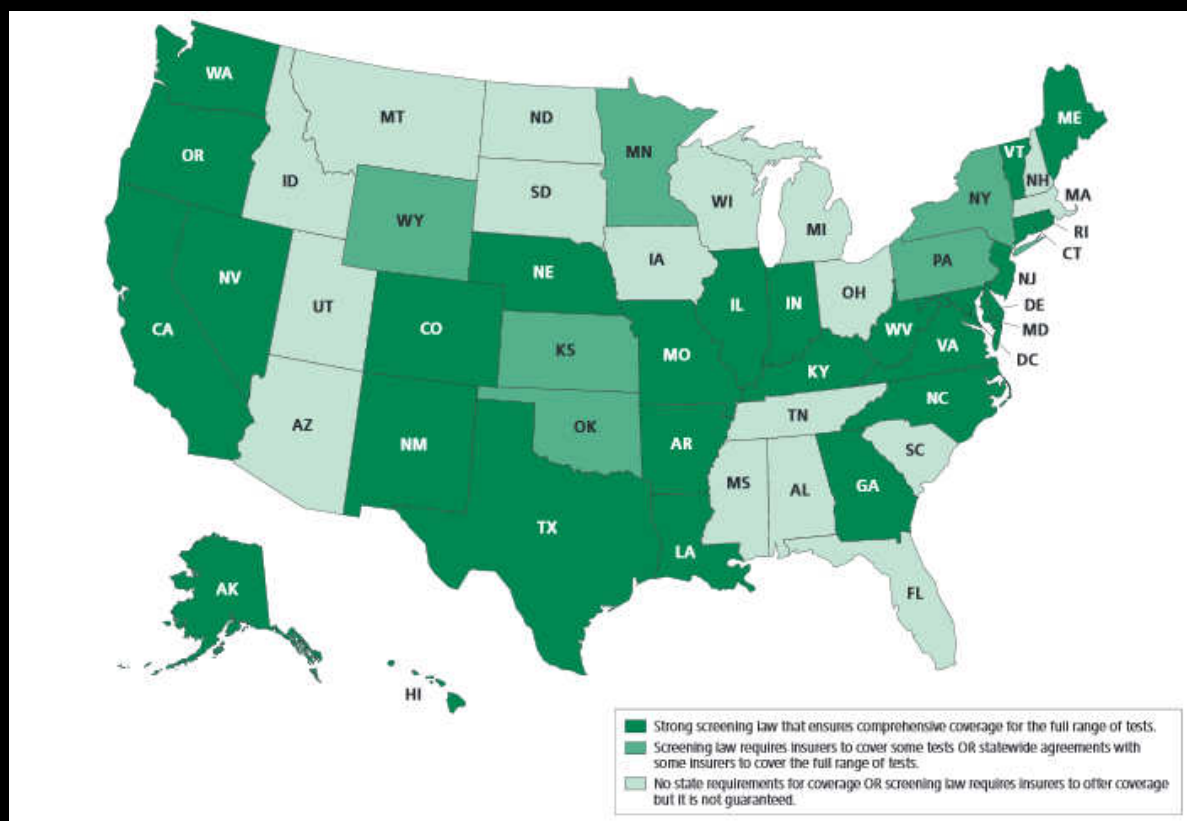
# Colon Cancer



# Distribution of Colorectal Cancer



# Maryland is pro-colon cancer screening



Colorectal cancer screening prevalence by state, 2006-2008

Colorectal cancer facts and figures 2011-2013.

<http://www.cancer.org/acs/groups/content/@epidemiologysurveillance/documents/document/acspc-028323.pdf>. Accessed 2/10/14

# Colorectal cancer screening

## First assess RISK

### **AVERAGE RISK INDIVIDUAL**

- All patients age 50 years and older, the asymptomatic general population

### **HIGH RISK**

- Personal history
- Family history



# Colorectal Cancer Screening

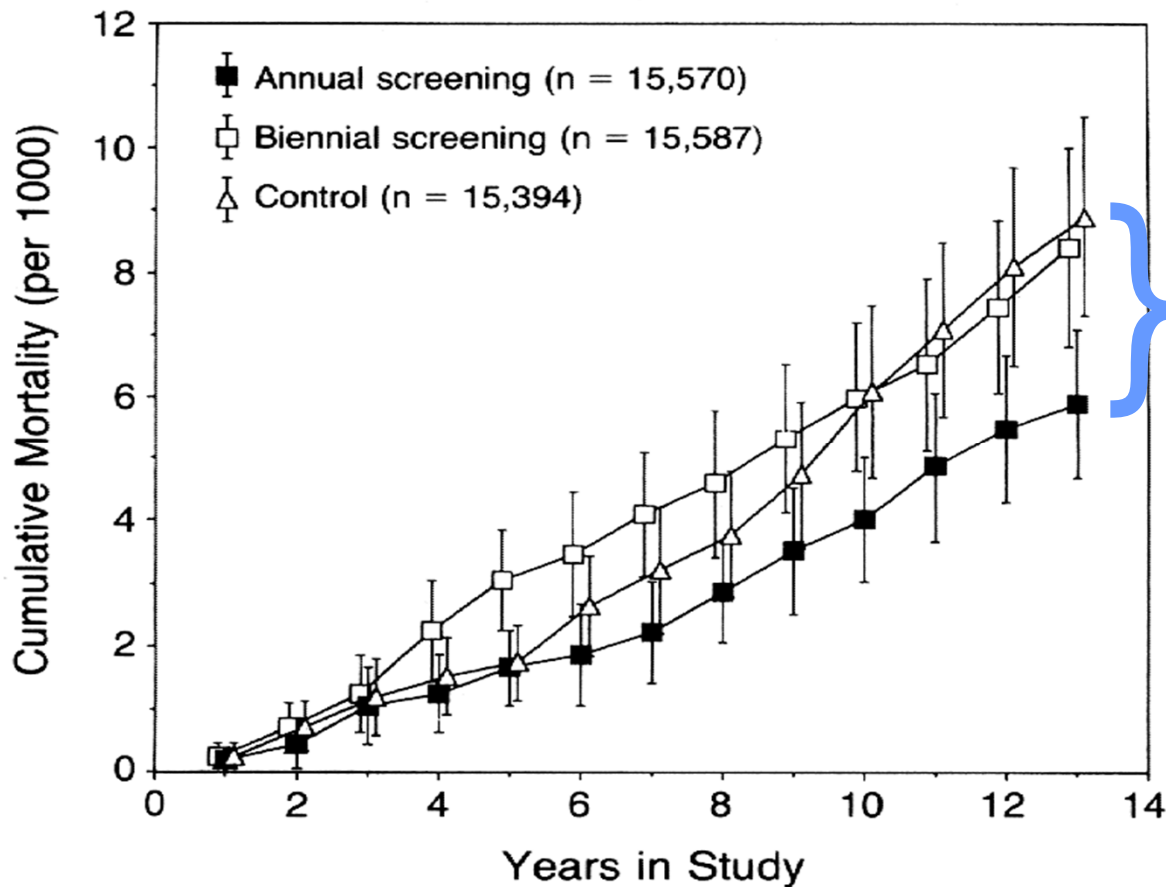
## Average risk

- Fecal occult blood testing (FOBT)
- Flexible sigmoidoscopy
- Barium enema
- Colonoscopy
- CT colography
- Stool genetic testing

## FOBT – Clinical Issues

- Guaiac-based (gFOBT) or fecal immunochemical testing (FIT – detects human globin)
- Test multiple stools
- Diet modification is necessary (gFOBT)
- OK to test when patient is on low-dose ASA or warfarin in therapeutic range
- All positives lead to full colon evaluation (colonoscopy)
- Rehydration leads to higher sensitivity and lower specificity

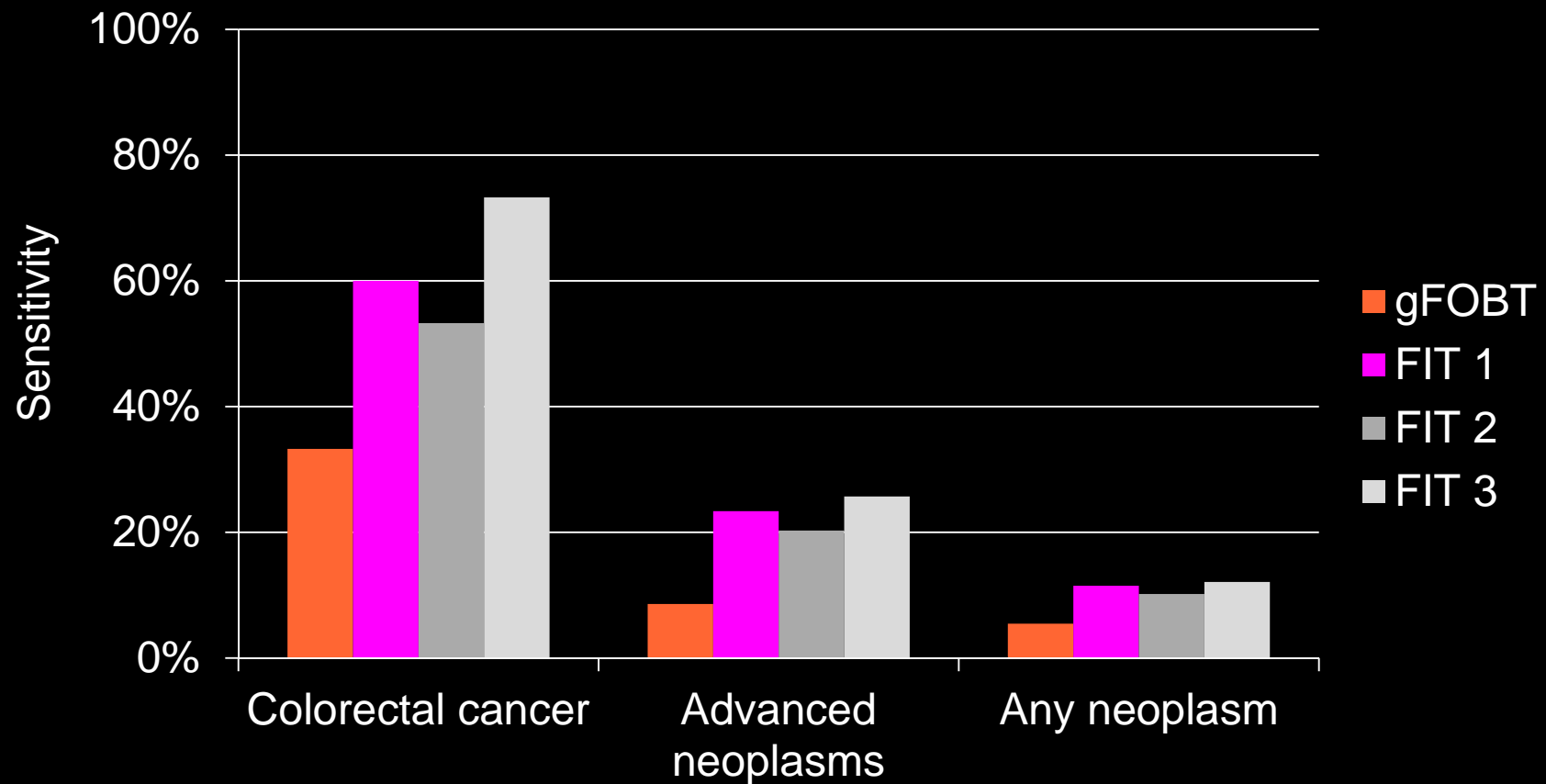
# Annual FOBT Saves Lives!



**33%  
reduction**

Mandel JS et al. N Engl J Med 1993 328:1365-71.

# FIT is more sensitive than gFOBT



# Follow-up of Positive FOBT

- Colonoscopy recommended

## HOWEVER:

- Only 52% of primary care physicians would recommend colonoscopy
- Only 29% of internal medicine residents would recommend colonoscopy

Winawer et al. Gastroenterol 1997;112:584

Sharma et al. Am J Gastroenterol 2000;95:1551

Sharma et al. Am J Gastroenterol 2000;95:1914

# Flexible Sigmoidoscopy

## PROS:

- May be done in the office
- Inexpensive, cost-effective
- Mortality from rectal cancer reduced by 60-70% in case-control studies
- Easier bowel preparation, usually done without sedation

## CONS:

- Detects only one-half of adenomas
- 40% of cancers arise proximal to splenic flexure
- 75% of proximal cancers have no adenomas distal to splenic flexure
- Often limited by discomfort, poor bowel preparation

Selby et al. N Engl J Med 1992;326:653  
Rex et al. Gastrointest Endosc 1999;99:727  
Newcomb et al. J Natl Canc Inst 1992;84:1572

Stewart Aust NZ J Surg 1999;69:2  
Painter et al. Endoscopy 1999;3:269

# Flexible Sigmoidoscopy Misses 50% of Lesions

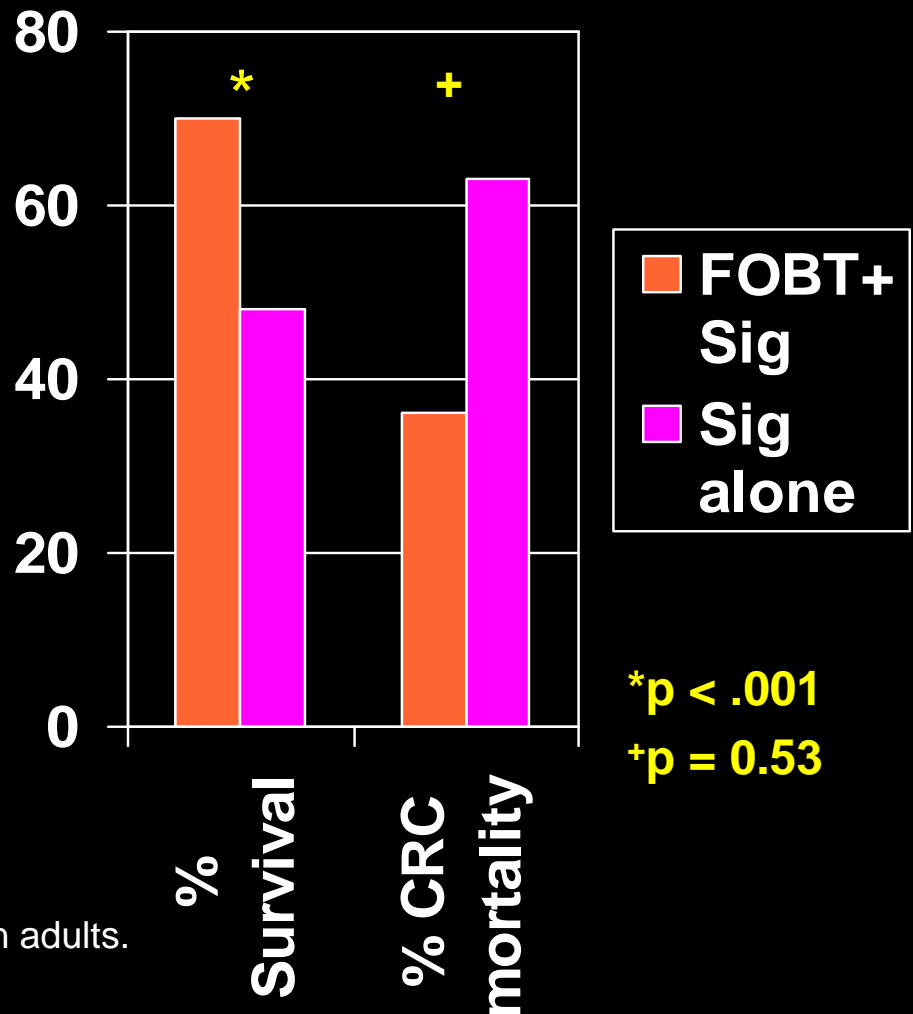
- Colonoscopy comparison studies:  
  
46-52% of patient with advanced proximal neoplasia (> 1 cm, villous, high-grade dysplasia or cancer) had no adenomas distal to the splenic flexure

Lieberman et al. N Engl J Med 2000; 343:162-8.

Imperiale et al. N Engl J Med 2000; 343:169-174.

# Combined FOBT and Sigmoidoscopy

- Case-control trial (N=21,750) w/rigid sigmoidoscopy – improved survival
- Other trials: FS + FOBT
  - Improved yield over FOBT alone
  - Adding FOBT to FS alone may not improve yield



Winawer et al. J Natl Cancer Inst 1993;85:1311  
Pignone et al. Screening for colorectal cancer in adults.  
<http://www.ahrq.gov/clinic/serfiles.htm>



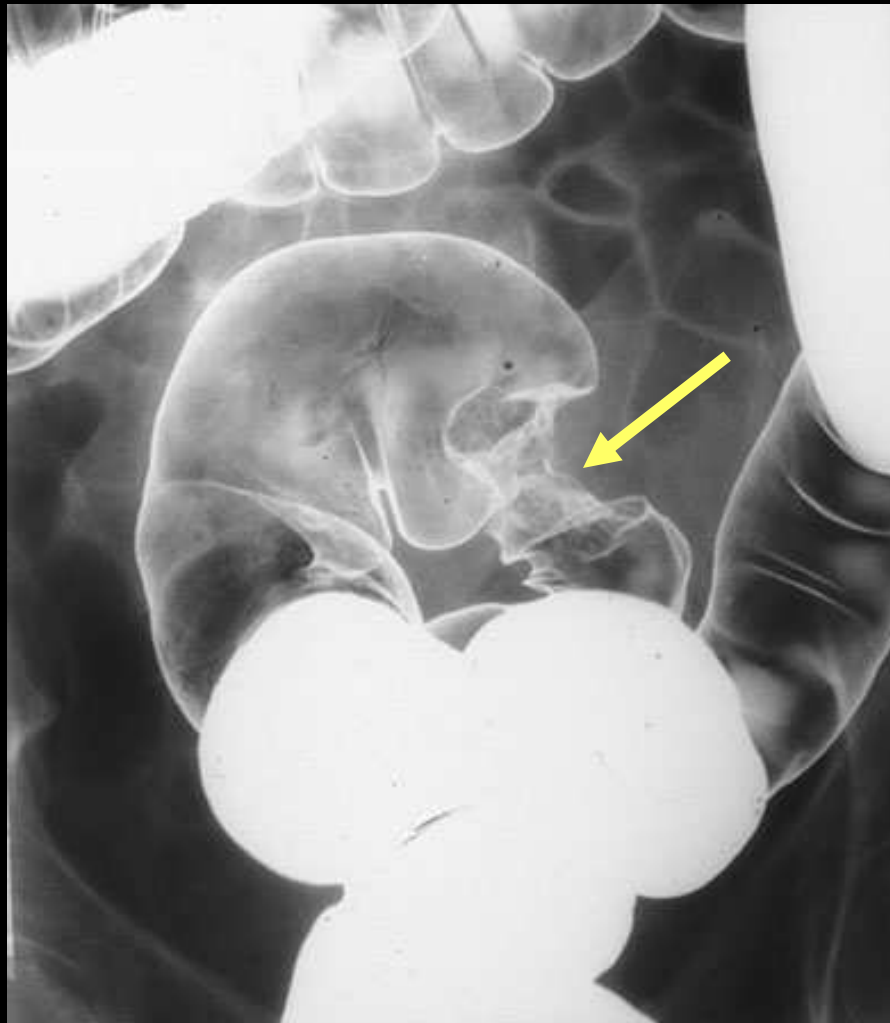
# FOBT + Flexible Sigmoidoscopy Misses 24% of Lesions

- Colonoscopy comparison studies:  
  
24.2% of patient with advanced proximal neoplasia (> 1 cm, villous, high-grade dysplasia or cancer) had negative FOBT and no adenomas distal to the splenic flexure.

Lieberman and Weiss. N Engl J Med 2001; 345:555-60.

# Colorectal Cancer Screening: Double-Contrast Barium Enema

## Colon Cancer



# Double-contrast Barium Enema

## PROS:

- Low cost, exams whole colon

## CONS:

- Never studied as a screening test
- Missed 50% of adenomas < 1 cm in National Polyp Study
- Sensitivity for cancer in patients with positive FOBT: 50-75%
- Poor specificity; best interval unknown

Winawer et al. Gastroenterol 1997;112:599

Rex. Endoscopy 1995;27:200

Lieberman et al. N Engl J Med 2000;343:163

# Colonoscopy



# Colonoscopy

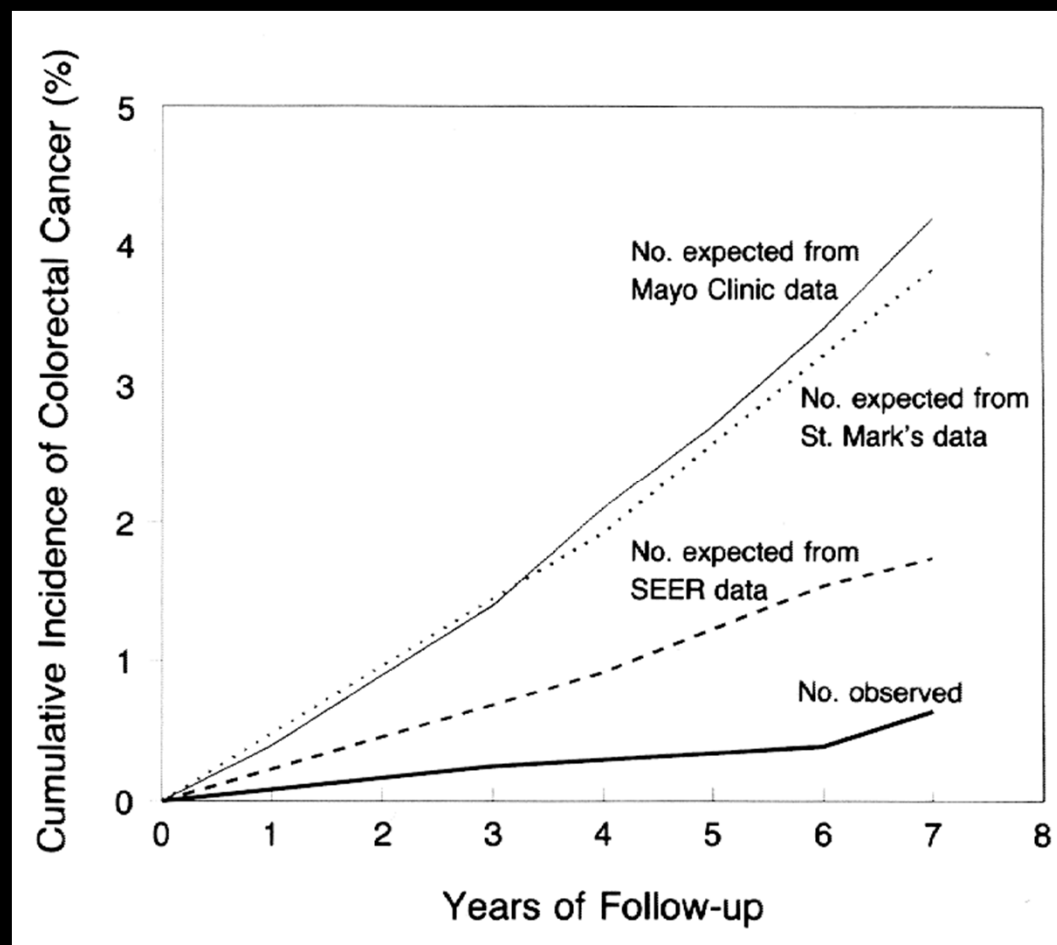
## PROS:

- Exams entire colon
- Therapeutic – polyps removed at time of procedure

## CONS:

- Invasive, risk of complications
- Requires bowel prep, missed work, escort home
- Incomplete procedures ~5%
- Missed polyps
- Randomized trials lacking

# Colonoscopic Polypectomy Reduces Colorectal Cancer Incidence



Winawer et al. N Engl J Med 1993; 329:1977-81.

# Miss Rate for Colonoscopy

	Comparison group	
	Tandem Colonoscopy	CT Colography
Adenoma $\leq$ 5 mm	27%	--
Adenoma 6 – 9 mm	13%	9%
Adenoma $\geq$ 10 mm	6%	12%

Rex et al. Gastroenterol 1997; 112:24-28.

Pickhardt et al. N Engl J Med 2003;349:2191-2200.

# Colonoscopy Complications

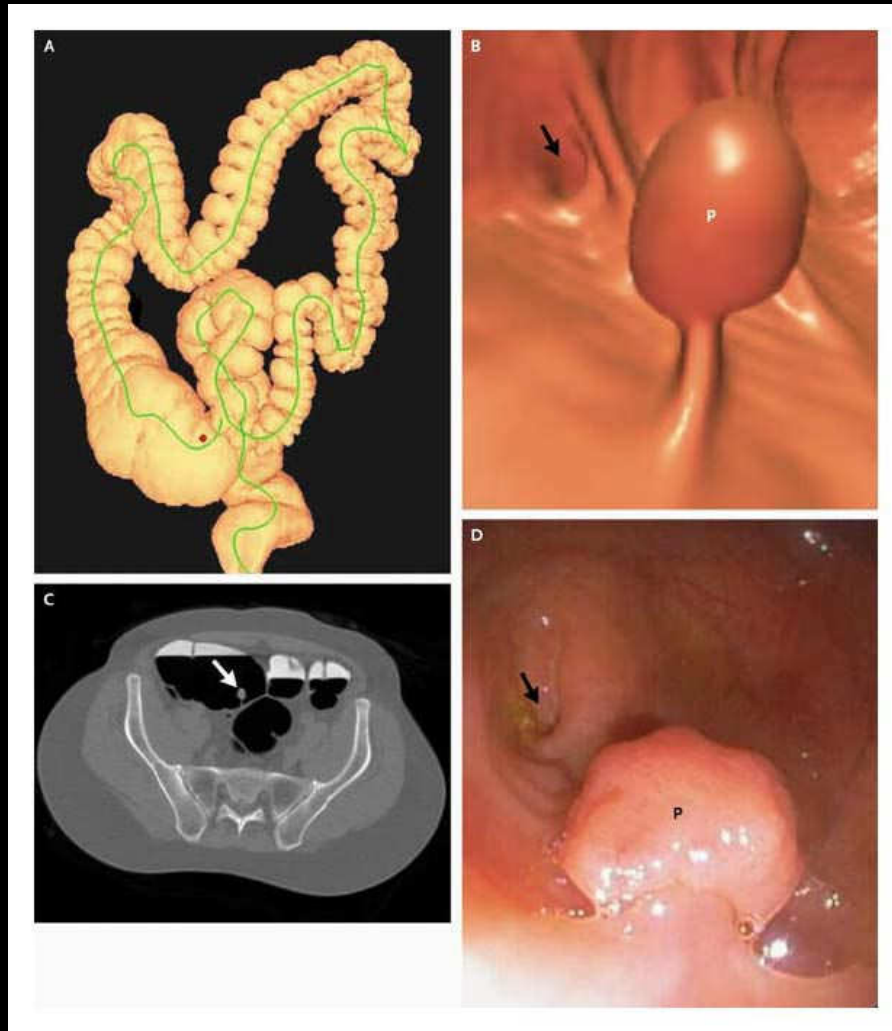
- Perforation 1-2/1000 procedures
- Bleeding 3/1000 procedures
- Mortality 1/10,000 procedures



# Cost-Effectiveness of Colorectal Cancer Screening

Screening method	Cost-effectiveness ratio (cost per life-years saved)
FOBT annually	\$ 5,691 - \$17,805
Flexible sigmoidoscopy every 5 years	\$12,477 - \$19,068
FOBT annually + flexible sigmoidoscopy every 5 yrs	\$13,792 - \$22,518
Double contrast barium enema every 5 years	\$11,168 - \$25,624
Colonoscopy every 10 yrs	\$ 9,038 - \$22,012

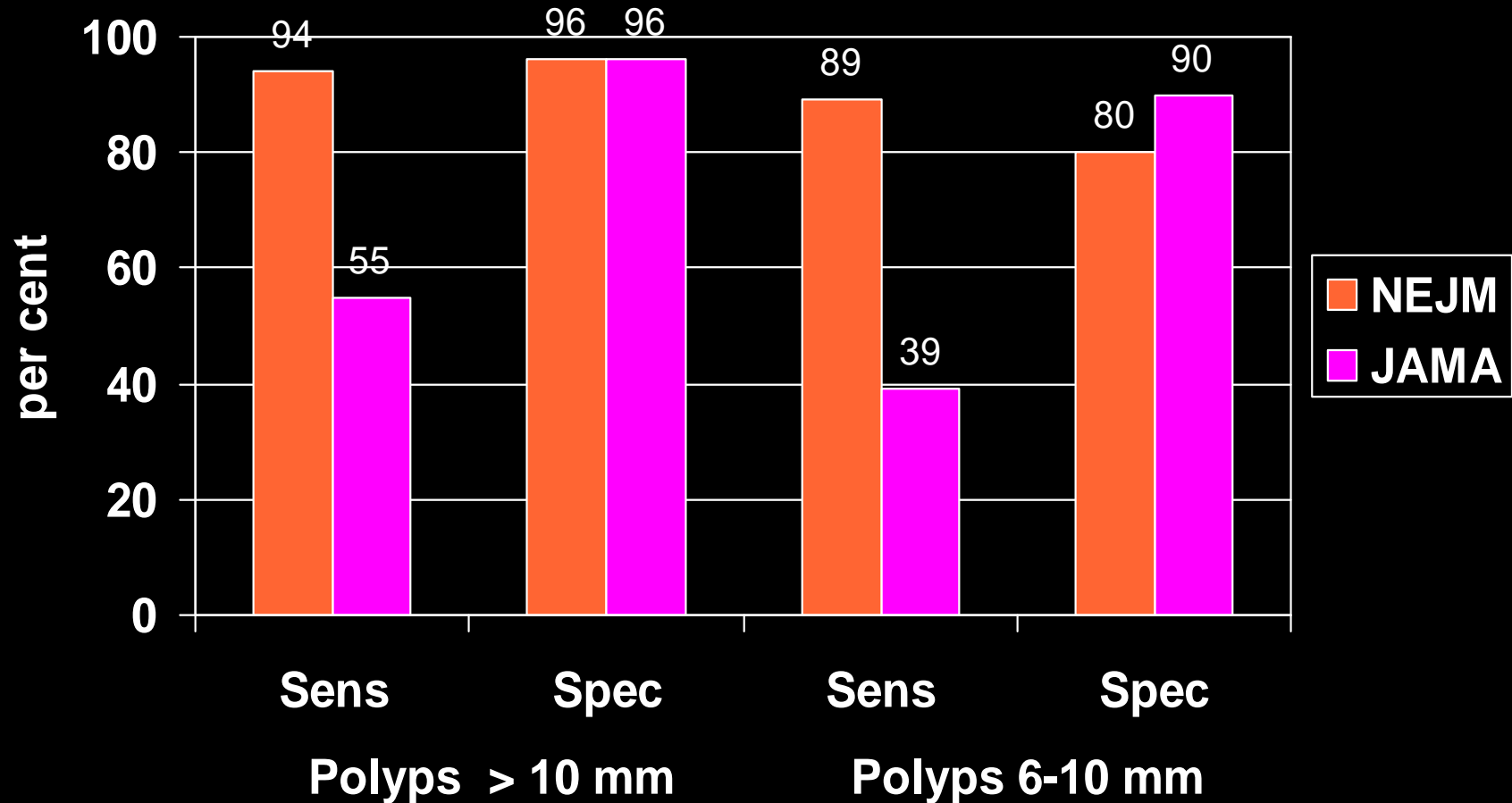
# CT Colonography



**Solitary 16-mm  
Pedunculated Cecal  
Polyp in a 55-Year-Old  
Man at Average Risk for  
Colorectal Neoplasia**

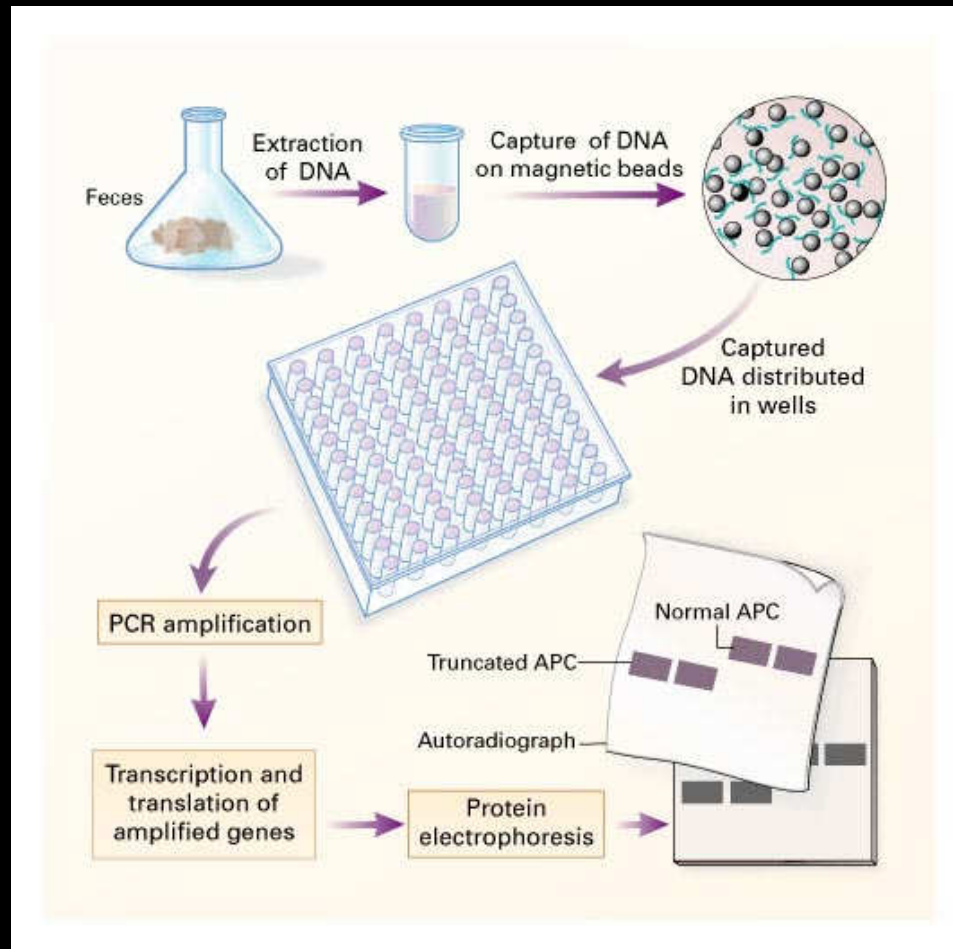
Pickhardt et al. N Engl J Med  
2003;349:2191-2200

# Virtual Colonoscopy Results are Variable



Pickhardt et al. N Engl J Med 2003;349:2191  
Cotton et al. JAMA 2004, 291:1731

# Stool DNA Testing



Schwartz. N Engl J Med 2002;346:302-4

# Stool DNA Testing

- Pros
  - No sedation or preparation necessary
  - Home-based (patient mails sample)
  - No risk
- Cons
  - Not clinically available
  - Cost and frequency of exam unknown
  - False negatives occur

# Colorectal Cancer Screening

Test	Interval
<b>Detects cancer and adenomatous polyps</b>	
Colonoscopy	Every 10 years
Flexible sigmoidoscopy	Every 5 years
CT colonography	Every 5 years
<b>Primarily detects cancer</b>	
Fecal occult blood testing with immunochemical test	Every year on 1-2 samples
Fecal occult blood testing with guaiac reagent	Every year on 3 samples

# Barriers to Screening

- Cost and lack of access to health care
- Lack of awareness of need for colorectal cancer screening
- Inadequate communication by health care providers
- Differences between patient and provider preferences for screening
- Low levels of education and income
- Personal barriers – fear and embarrassment