

Screening Dinner

Advances in Colorectal Cancer Screening: The State of the Science



Lisa A. Hall, MHA Senior Director of Prevention & Screening Colorectal Cancer Alliance

Access the Alliance's Lookbook Here:



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Welcome and Introductions

Wi-Fi: National Union Building

Password: 918fstdc

Screening Dinner Discussion Agenda

5:30-5:45pm

Registration and Buffet

5:45-6:00pm

Welcome and Introduction

- Remarks by Michael Sapienza, Chief
 Executive Officer, Colorectal Cancer Alliance
- Lisa Hall, Senior Director of Prevention and Screening, Colorectal Cancer Alliance

6:00-6:30pm

Screening Landscape

 Dr. Robert Smith, Vice President of Early Cancer Detection Science, American Cancer Society

6:30-7:00pm

Evaluating CRC Testing Options: A Practical Framework

- Dr. Djenaba Joseph, CAPT, U.S. Public Health Service, Chief, Program Service Branch, Centers for Disease Control Prevention
- Dr. Richard Wender, Professor and Chair Family Medicine and Community Health University of Pennsylvania

7:00-8:50pm

Screening Modality Expert Panel and Meeting Participant Discussion

Co-facilitated by:

- Dr. Djenaba Joseph
- Dr. Richard Wender

Panelists:

- Dr. Lance Baldo, Chief Medical Officer, Freenome
- Dr. Erica Barnell, Chief Medical Officer and Co-Founder, Geneoscopy
- Dr. Craig Eagle, Chief Medical Officer, Guardant
- Dr. Steven Itzkowitz, Professor of Medicine, Oncological Sciences and Medical Education, Ichan School of Medicine at Mount Sinai, Chair, American Cancer Society's National Colorectal Cancer Round Table
- Dr. Todd Kelley, Vice President of Medical Affairs, Polymedco
- Dr. Theodore Levin, Research Scientist, Intern Associate Director of Cancer Section, Kaiser North Division of Research
- Dr. Paul Limburg, Chief Medical Officer, Screening Exact Sciences

8:15-8:30pm

Closing and Next Steps

Remarks by Michael Sapienza, Chief Executive Officer, Colorectal Cancer Alliance Lisa Hall, Senior Director of Prevention and Screening, Colorectal Cancer Alliance

8:30-9:30pm Cocktail Hour





Michael Sapienza

Chief Executive Officer Colorectal Cancer Alliance

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What do you believe are the biggest factors driving the 20% higher incidence rate within black and brown communities? (choose your top 2 factors)



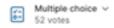
| What do you believe are the biggest factors driving the 20% higher incidence rate within black and brown communities? (choose your top 2 factors) | |
|---|-----|
| A. The screening test used | |
| | 16% |
| B. Screening test quality | |
| | 8% |
| C. On-time screening | 72% |
| D. Follow-up to positive stool/blood | |
| | 34% |
| E. Environmental factors | |
| | 46% |
| F. Other | |
| | 6% |







What do you believe are the biggest factors driving the 35% increased mortality rate within black and brown communities? (choose your top 3 factors)



What do you believe are the biggest factors driving the 35% increased mortality rate within black and brown communities? (choose your top 3 factors)

A. Stage of diagnosis

| | 81% |
|--|------|
| B. Co-morbid diseases | |
| | 31% |
| C. Access to care | 0.5% |
| | 85% |
| D. Treatment provided | 29% |
| E Ability to consistently resolve treatment | |
| E. Ability to consistently receive treatment | 50% |
| F. Other | |
| | 4% |





Robert Smith, PhD

Vice President of Early Cancer Detection Science, American Cancer Society

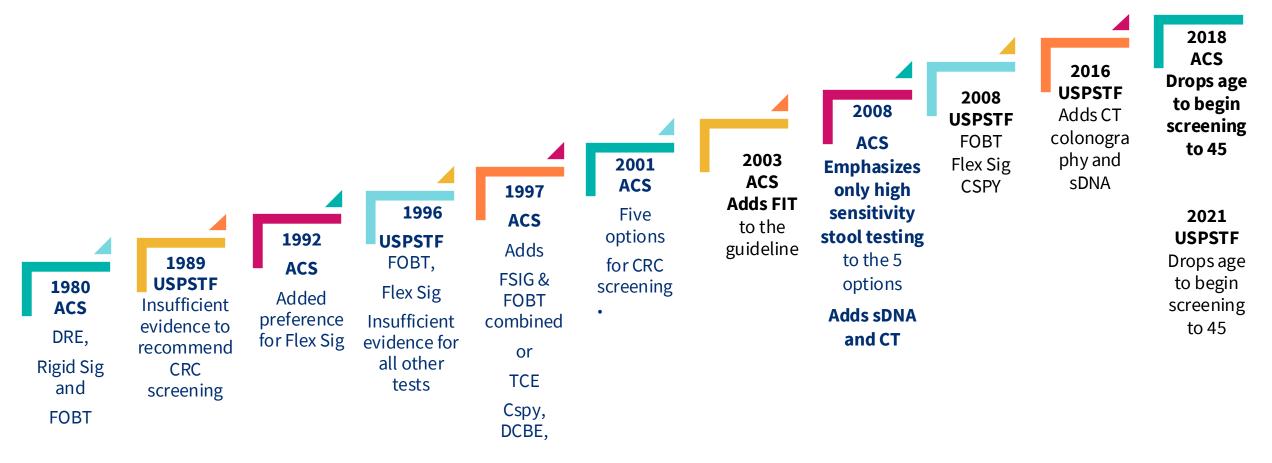
The CRC Screening Landscape

Robert A. Smith, PhD Director, Center for Early Cancer Detection Science American Cancer Society

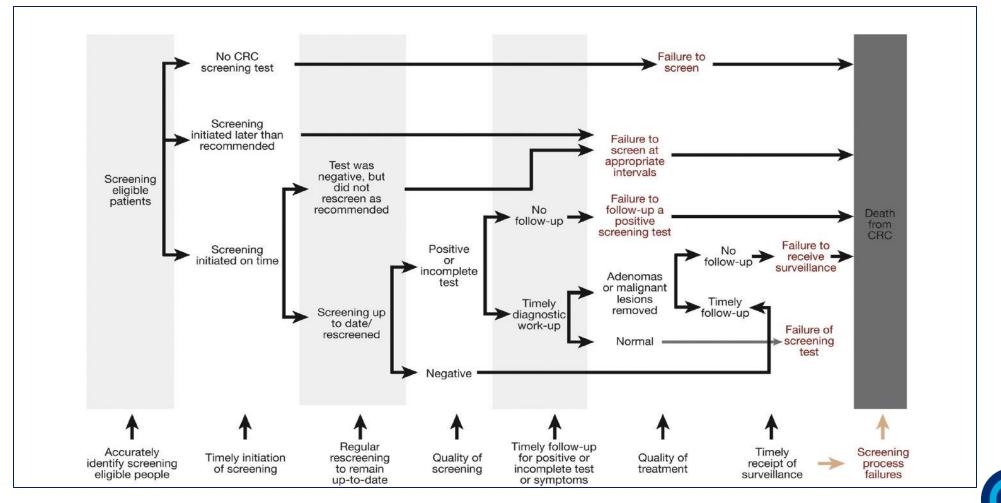
Advances in Colorectal Cancer Screening: The State of the Science Colorectal Cancer Alliance Washington, DC, May 19, 2024



Brief History of ACS & USPSTF Colorectal Cancer Screening Guidelines



CRC test exposure trajectories and failures in people who died of CRC, KPNC and KPSC 2006–2012

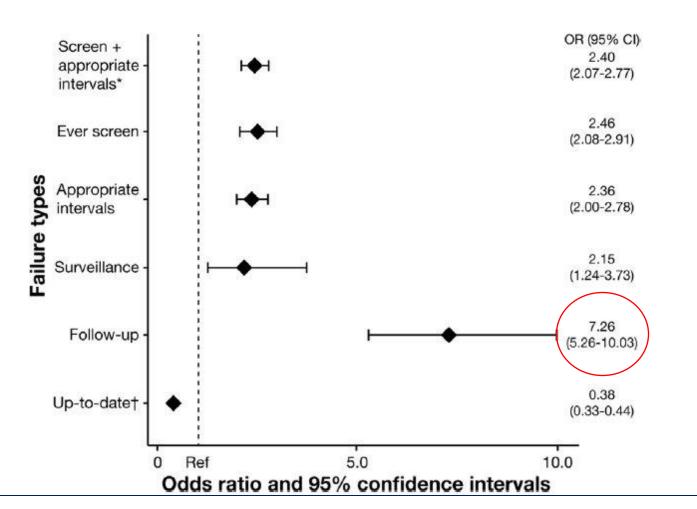


Doubeni, Gastroenterology 2018, Vol. 156, No. 1

colorectal cancer

alliance

Association between screening patterns and death from CRC, KPNC and KPSC 2006–2012



Doubeni, Gastroenterology 2018, Vol. 156, No. 1

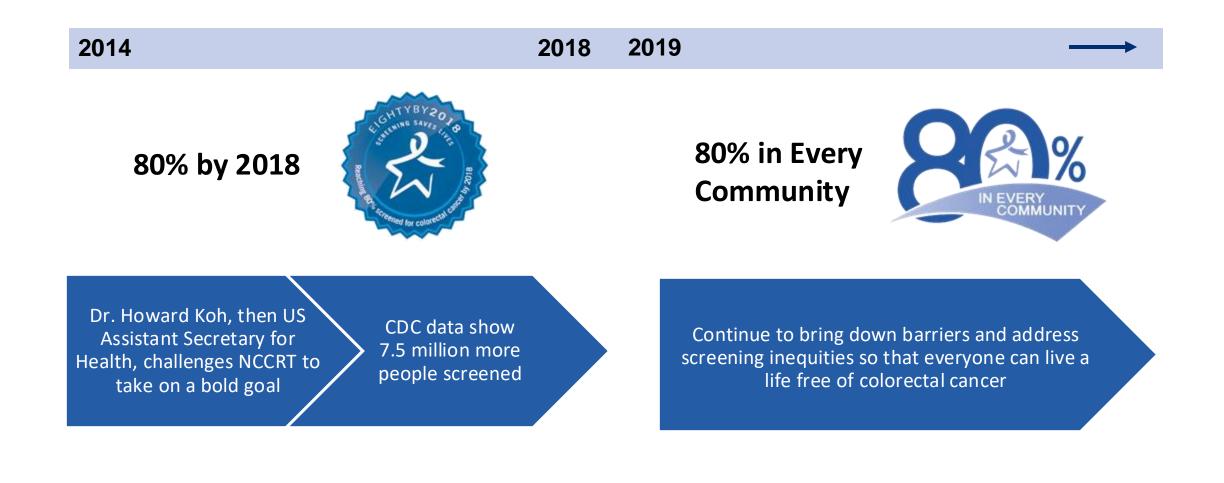
Being up to date on screening lowered the risk of dying of CRC by more than 60%.

Relative to those who were up to date, the risk of death from CRC was more than 2-fold higher in people who had failed to screen

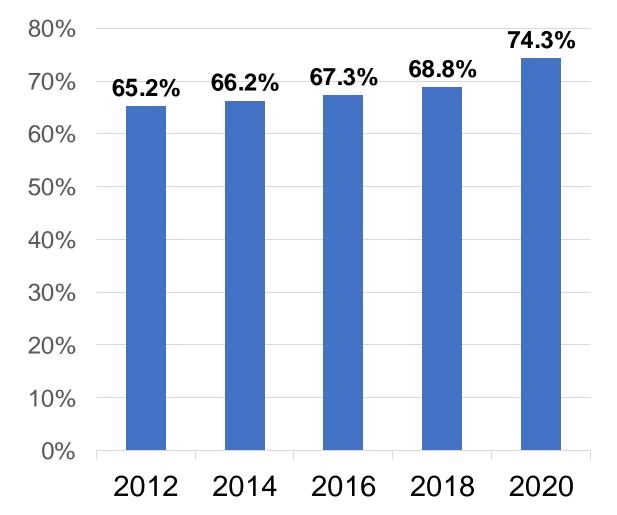
Risk of death was 7-fold higher in those with failure of followup.



The Timeline of Two 80% Campaigns

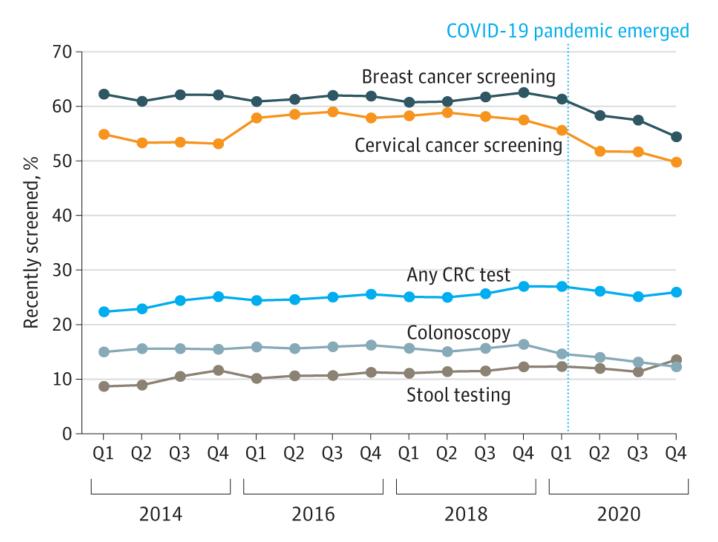


Percent of US Adults Up to Date with CRC Screening (BRFSS)



9.3 million additional people screened by 2018!

Recent CRC Screen in 2020, Ages 50-75





Fedewa SA, Star J, Bandi P, et al. Changes in Cancer Screening in the US During the COVID-19 Pandemic. *JAMA Netw Open*. 2022;5(6):e2215490. Published 2022 Jun 1. doi:10.1001/jamanetworkopen.2022.15490



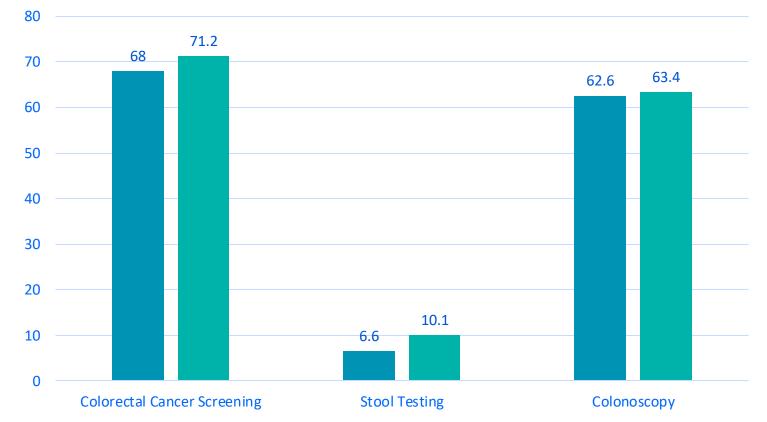
UTD CRC Screening in 2021, Ages 50-75



Colorectal cancer: 4% (aPR, 1.04; 95% Cl, 1.02-1.05)

Stool testing: 53% (aPR, 1.53; 95% CI, 1.39 to 1.68)

Colonoscopy: (aPR, 1.0; 95% Cl, 0.98 to 1.02)

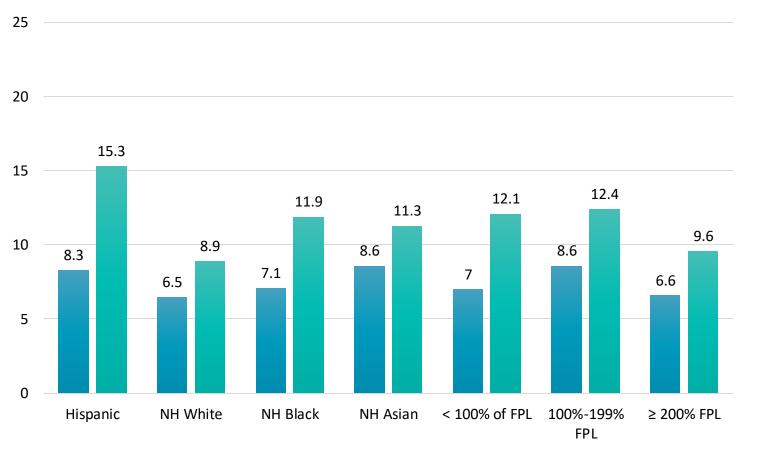


2019 2021

Star J, Bandi P, Nargis N, et al. Updated Review of Major Cancer Risk Factors and Screening Test Use in the United States, with a Focus on Changes During the COVID-19 Pandemic [published online ahead of print, 2023 May 2]. Cancer Epidemiol Biomarkers Prev. 2023;EPI-23-0114. doi:10.1158/1055-9965.EPI-23-0114



Stool Testing by Race/Income, Ages 50-75



2019 2021

Star J, Bandi P, Siegel RL, et al. Cancer Screening in the United States During the Second Year of the COVID-19

20 Pandemic

[published online ahead of print, 2023 Feb 23]. J Clin Oncol. 2023; JCO2202170. doi:10.1200/JCO.22.02170



Colorectal Cancer Screening (%), Adults 45 Years and Older, US, 2021

| | Stool test ^a | Colonoscopy ^b | Up to date ^c | |
|-------------|-------------------------|--------------------------|-------------------------|-------|
| | ≥45 years | ≥45 years | ≥45 years | 45-75 |
| Overall | 10 | 54 | 59 | 58 |
| Sex | | | | |
| Males | 9 | 54 | 58 | 56 |
| Females | 10 | 55 | 60 | 60 |
| Age (years) | | | | |
| 45-49 | 3 | 18 | 20 | _ |
| 50-54 | 9 | 43 | 50 | - |
| 55-64 | 11 | 65 | 70 | |
| 65-75 | _ | - | - | 83 |
| 65-74 | 15 | 74 | 80 | - |
| ≥75 | 10 | 67 | 70 | _ |



Colorectal Cancer Screening (%), Adults 45 Years and Older, US, 2021

| | Stool test ^a | Colonoscopy ^b | Up to date ^c | |
|-----------------------|-------------------------|--------------------------|-------------------------|-------|
| | ≥45 years | ≥45 years | ≥45 years | 45-75 |
| Overall | 10 | 54 | 59 | 58 |
| Race/ethnicity | | | | |
| Hispanic | 14 | 46 | 52 | 51 |
| White only | 9 | 57 | 61 | 60 |
| Black only | 11 | 57 | 61 | 59 |
| Asian only | 10 | 45 | 50 | 48 |
| AIAN only or multiple | 10 | 48 | 52 | 52 |
| Sexual orientation | | | | |
| Gay/lesbian | 12 | 57 | 64 | 61 |
| Straight | 10 | 55 | 59 | 58 |
| Bisexual | _ | 48 | 51 | 57 |



Colorectal Cancer Screening (%), Adults 45 Years and Older, US, 2021

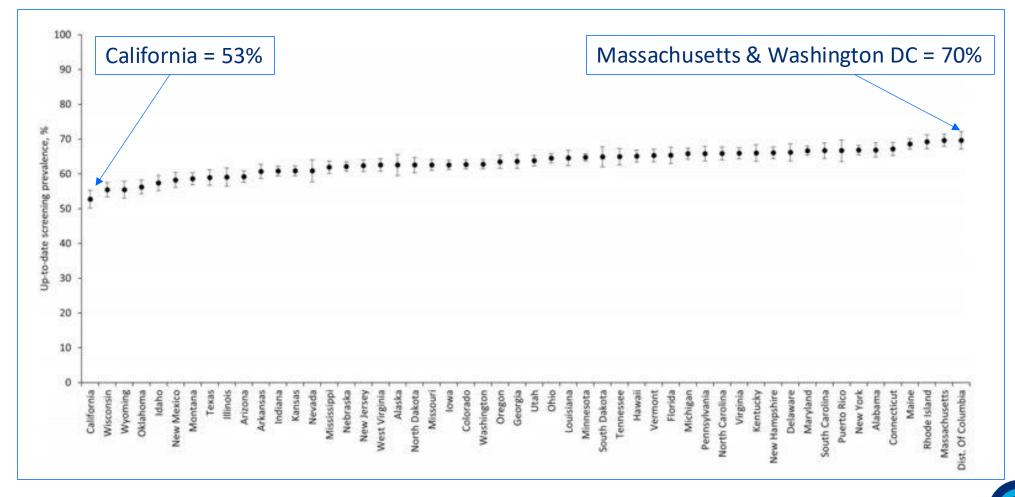
| | Stool test ^a | Colonoscopy ^b | Up to date ^c | |
|------------------------------------|-------------------------|--------------------------|-------------------------|-------|
| | ≥45 years | ≥45 years | ≥45 years | 45-75 |
| Overall | 10 | 54 | 59 | 58 |
| Immigration status | | | | |
| Born in United States/US Territory | 9 | 57 | 61 | 60 |
| In United States <10 years | 9 | 25 | 29 | 30 |
| In United States ≥10 years | 12 | 48 | 53 | 52 |
| Education | | | | |
| Less than high school | 11 | 43 | 48 | 47 |
| High school diploma | 9 | 51 | 55 | 54 |
| Some college | 11 | 56 | 61 | 59 |
| College graduate | 9 | 60 | 64 | 63 |



Colorectal Cancer Screening (%), Adults 45 Years and Older, US, 2021

| | Stool test ^a ≥45 years | Colonoscopy ^b ≥45 years | Up to date ^c | |
|-------------------------------|--------------------------------------|---------------------------------------|-------------------------|-------|
| | | | ≥45 years | 45-75 |
| verall | 10 | 54 | 59 | 58 |
| Income level | | | | |
| <100% FPL | 11 | 42 | 47 | 46 |
| 100% to <200% FPL | 12 | 47 | 52 | 51 |
| ≥200% FPL | 9 | 58 | 62 | 61 |
| Insurance status | | | | |
| Uninsured | 4 | 18 | 21 | 22 |
| Private | 9 | 59 | 63 | 64 |
| Medicaid/Public/dual eligible | 11 | 48 | 52 | 53 |
| Medicare (ages ≥65 years) | 15 | 69 | 75 | 82 |
| Other | 15 | 68 | 73 | 74 |

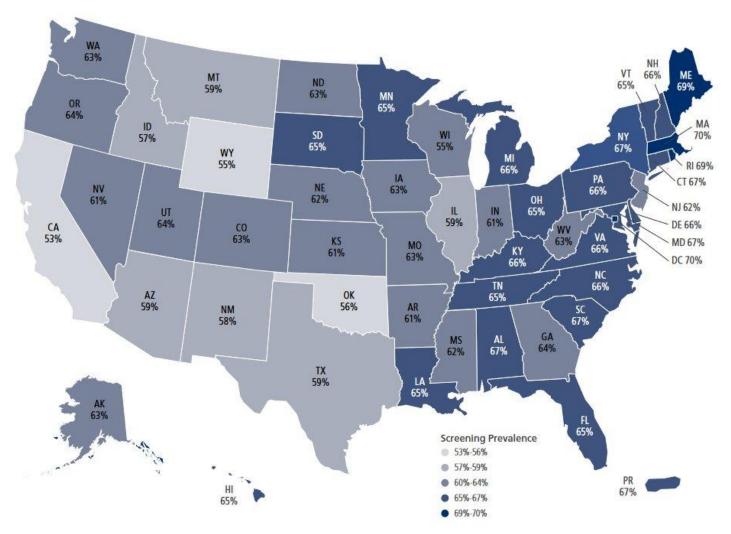
Reported Up-to-date colorectal cancer screening (%), adults aged 45 years and older by state, 2020, United States.



Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnostic examinations. Source: Behavioral Risk Factors Surveillance System, 2020. From Siegel, et al. *CA Cancer J Clin*. 2023;73:233–254.



Reported Up-to-date colorectal cancer screening (%), adults aged 45 years and older by state, 2020, United States.



*Blood/DNA stool test, sigmoidoscopy, or colonoscopy in the past 1/3, 5, or 10 years, respectively. Note: Estimates are age adjusted to the 2000 US standard population and do not distinguish between examinations for screening and diagnosis.

*Blood/DNA stool test, sigmoidoscopy, or colonoscopy in the past 1/3, 5, or 10 years, respectively. Note: Estimates are age a djusted to the 2000 US standard population and diagnosis. Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnosis. Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnosis. Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnosis. Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnosis. Prevalence is age-adjusted to the 2000 US standard population and does not distinguish between screening and diagnosis. diagnostic examinations. Source: Behavioral Risk Factors Surveillance System, 2020. From ACS Colorectal Cancer Facts and Figures, 2023-2025



JAMA Internal Medicine | Original Investigation

National Breast, Cervical, and Colorectal Cancer Screening Use in Federally Qualified Health Centers

Trisha L. Amboree, PhD; Jane R. Montealegre, PhD; Susan L. Parker, MPH; Ashvita Garg, PhD; Haluk Damgacioglu, PhD; Kathleen M. Schmeler, MD; Elizabeth Y. Chiao, MD; Elizabeth G. Hill, PhD; Kalyani Sonawane, PhD; Ashish A. Deshmukh, PhD; Prajakta Adsul, MBBS, MPH, PhD

- Cross-sectional analysis of cancer screening from January 1 through December 31, 2020. Data from 1,364 FQHCs were obtained from the Health Center Program Uniform Data System (UDS).
- Participants were 16,696,692 US adults served by FQHCs who were eligible for breast (age, 50-74 years), cervical (age, 21-64 years), and colorectal (age, 50-75 years) cancer screening.
- Nationally, screening use in FQHCs was 45.4% for breast cancer, 51.0% for cervical cancer, and 40.2% for colorectal cancer.
- The contribution of the underscreened population served by FQHCs to the national underscreened general population was 14.7% for colorectal cancer (roughly 1 in 6).

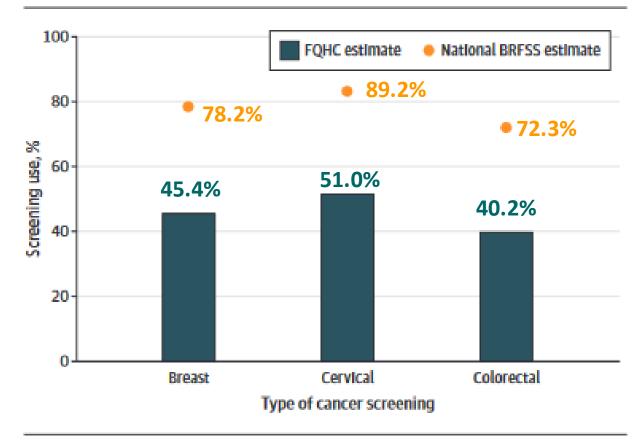


JAMA Internal Medicine Published online April 29, 2024

JAMA Internal Medicine | Original Investigation National Breast, Cervical, and Colorectal Cancer Screening Use in Federally Qualified Health Centers

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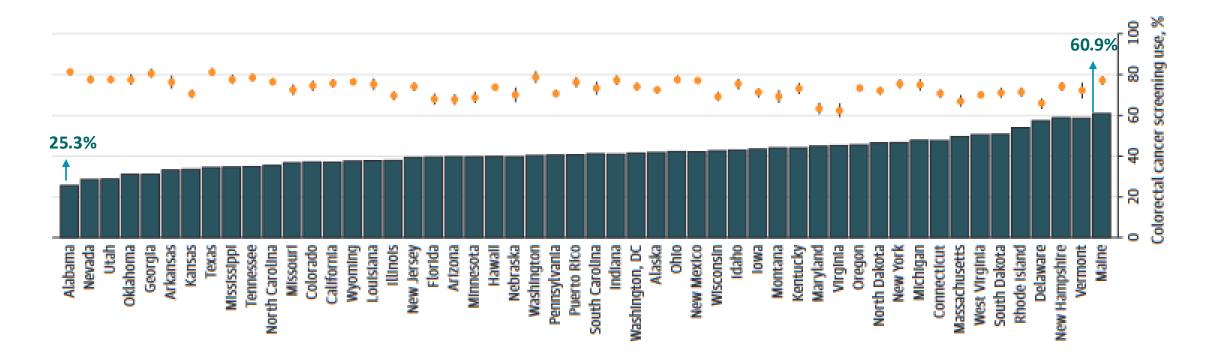
Figure 1. National Cancer Screening Use in Federally Qualified Health Centers (FQHCs) and the Behavioral Risk Factor Surveillance System (BRFSS), 2020

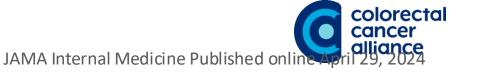




JAMA Internal Medicine Published online April 29, 2024

Figure 2. State-Level Colorectal Cancer Screening Use in Federally Qualified Health Centers (FQHCs) and the Statewide Behavioral Risk Factor Surveillance System, 2020







Analyzing Factors Influencing Non-Adherence to USPSTF Colorectal Cancer Screening Guidelines Using 2022 BRFSS Data

- Data: 2022 Behavioral Risk Factor Surveillance System (BRFSS)
- Results: Among national participants, 29.2% reported non-adherence with USPSTF guidelines for CRC screening.
- Significant factors associated with non-adherence included:
 - Lacking health insurance (OR: 2.65, 95% CI: 2.40 2.93)
 - Not completing high school (OR: 1.56, 95% CI: 1.42 1.72)
 - Identifying as non-white or being Hispanic or Latino (OR: 1.14, 95% CI: 1.10 1.19)
 - Having an income below \$35,000 (OR: 1.75, 95% CI: 1.67 1.83)
 - Longer time since last primary care appointment (5 years since last appointment versus within the last year (OR:13.55, 95% CI: 12.24 – 15.01)



CCC Cancer Causes & Control

Prevalence and predictors of colorectal cancer screening in the United States: evidence from the HINTS database 2018 to 2020

- Methods: Survey analysis of 7,505 adults aged 45–75 years who completed the 2018 to 2020 HINTS survey.
- Results: 76% of eligible adults had received screening for CRC.
- Factors associated with recent CRC screening included:
 - Increasing age
 - Having some college experience, a college degree or higher [OR 1.69; 95% CI (1.24, 2.29)]
 - Health insurance coverage [OR 4.48; 95% CI (2.96, 6.76)]
 - Primary care provider access [OR 2.48; 95% CI (1.91, 3.22)]
 - Presence of a comorbid illness [OR 1.39; 95% CI (1.12, 1.73)]
 - People who currently smoke were less likely to undergo CRC screening [OR 0.59; 95% CI (0.40, 0.87)].
 - Among adults aged 50–64 years, being of Hispanic origin [OR 0.60; 95% CI (0.3% colorecta 0.92)] was associated with a lower likelihood of CRC screening.

Barriers to CRC Screening in Rural Areas

Perceived barriers by rural residents included:

- Individual-level structural barriers
 - High cost of screening procedures, or lack of insurance coverage, and lack of time
- Screening procedure related barriers
 - Embarrassment or discomfort, fear of the test, fear of finding cancer, and fear of burdening the family

• Individual-level perception/knowledge barriers

• Lack of knowledge on screening test options and guidelines, lack of perceived need when there are no symptoms, and misperception that CRC is a male disease and CRC screening is more important for men

Provider level barriers

• Lack of provider recommendation, distrust of providers or health care system, inadequate supply of specialists (sometimes even primary care physicians, and lack of reminder system.

• Rural specific attitudes

 Lack of prevention attitude, lack of privacy, and distance/transportation to screening facilities.



Understanding misperceptions of colorectal cancer risk and cancer screening based on the health belief model: a systematic review of qualitative literature

Noor Azreen Masdor¹ · Azmawati Mohammed Nawi¹ · Rozita Hod¹

- Background: Risk perception, knowledge, and awareness are important predictors of colorectal cancer (CRC) behavior and screening. Misperception appears to be the cause of negative CRC behaviors and refusal to be screened.
- Methods: Systematic review of qualitative studies (n=15) to identify common CRC misperceptions in the general population. Health Belief Model (HBM) domains were identified.
- There were two major HBM domains for misperception of CRC risk:
 - High perceived severity (CRC is fatal and incurable) and
 - Low perceived susceptibility (older man's disease, asymptomatic, sexual behavior).
- A high perceived barrier was that screening was considered embarrassing, painful, and inappropriate, leading to screening refusal
- A high perceived severity resulted in the belief that CRC is fatal and that screening is ineffective.
- Identifying the most common misperceptions enables the strategic planning of HBMbased interventions, and more engaging risk communication could improve CRC prevention and control

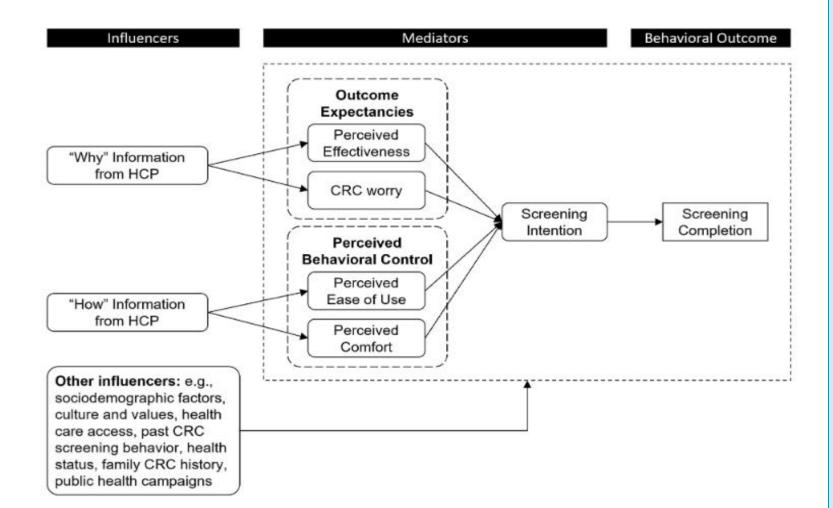


colorectal cancer Provider communication contributes to colorectal cancer screening intention through improving screening outcome expectancies and perceived behavioral control

SOCIAL SCIENCE

6

Xuan Zhu^{a,*}, Linda Squiers^b, Leah Helmueller^c, Gabriel Madson^b, Brian G. Southwell^b, Shama Alam^c, Lila J. Finney Rutten^d



The model posits that different types of information providers communicate influence patients' intention to complete screening through two sets of pathways.

1) WHY: Influences screening intention through changing patients' outcome expectancies about screening.

2) HOW: Information about preparation, procedures, and instructions about test, and how an individual would perform the behavior (changing patients' perceived behavioral control, i.e., the individual's evaluation of the feasibility of completing CRG orectal screening)



Provider communication contributes to colorectal cancer screening intention through improving screening outcome expectancies and perceived behavioral control

- Xuan Zhu^{a,*}, Linda Squiers^b, Leah Helmueller^c, Gabriel Madson^b, Brian G. Southwell^b, Shama Alam^c, Lila J. Finney Rutten^d
- Provider communication about the "Why" and "How" of mt-sDNA screening was positively associated with mt-sDNA test completion. Mt-DNA test completion mediated "Why" and "How" communication's impact on mt-sDNA screening intention.
- The authors were able to measure patients' reports of receiving specific information related to mt-sDNA screening from providers beyond the simple presence or absence of a provider recommendation, thus enabling the examination of the differential roles of communication *contents* in shaping patient decision-making.
- CRC screening interventions could consider implementing provider-patient communication strategies focusing on improving patient understanding of the rationale for CRC screening and the effectiveness of available screening options as well as addressing barriers and enhancing patients' self-efficacy in completing their preferred screening option





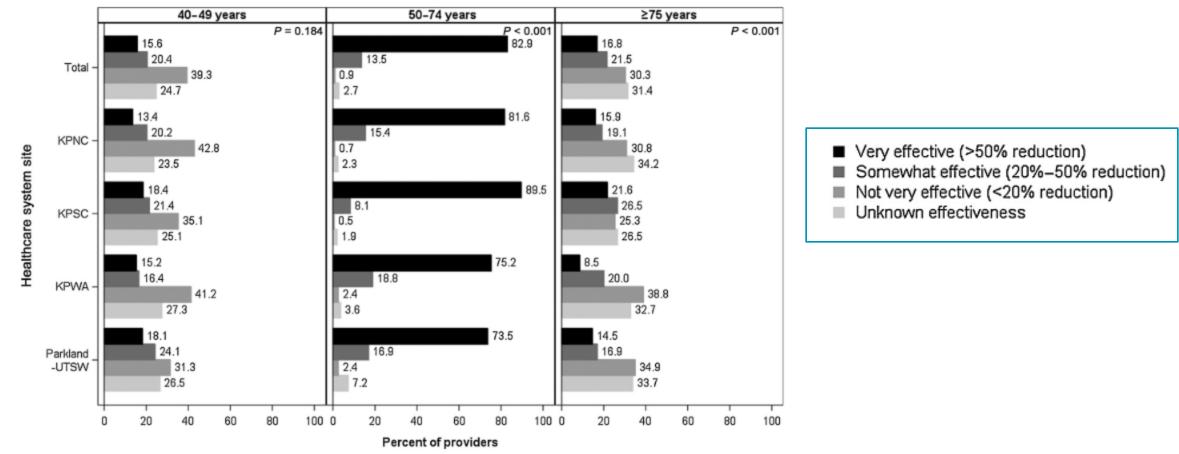
Primary Care Provider Beliefs and Recommendations About Colorectal Cancer Screening in Four Healthcare Systems

Nirupa R. Ghai¹, Christopher D. Jensen², Sophie A. Merchant², Joanne E. Schottinger³, Jeffrey K. Lee², Jessica Chubak⁴, Aruna Kamineni⁴, Ethan A. Halm^{5,6}, Celette Sugg Skinner⁶, Jennifer S. Haas⁷, Beverly B. Green⁴, Nancy T. Cannizzaro⁸, Jennifer L. Schneider², and Douglas A. Corley²

- Providers (n =1,281) within four healthcare systems completed a survey in 2017–2018 regarding their perceptions of CRC test effectiveness and recommended intervals for colonoscopy and fecal immunochemical testing (FIT) for patients ages 40–49, 50–74, and ≥75 years.
- For patients ages 40–49 and ≥75, more than one-third of providers believed the tests were somewhat or very effective, although >80% did not routinely recommend screening by either test for these age groups.



Provider beliefs about the effectiveness of colonoscopy in reducing colorectal cancer-related mortality, by patient age and healthcare system site



 x^2 P-values at the top right corner of each panel column compare provider responses across sites



Cancer Prev Res; 13(11) November 2020

Summary (1)

- The prevalence of reported up-to-date screening with any recommended test among individuals aged 50 years and older increased from 38% in 2000 to 66% in 2018 according to data from the NHIS.
- According to the NHIS, 59% of individuals aged 45 years and older were up to date on CRC screening in 2021, ranging from 50% of Asian individuals to 61% of White and Black individuals.
- Characteristics associated with low screening prevalence include:
 - Residence in the United States for <10 years (29%)
 - Being uninsured (21%)
 - Having less than a high school education (48%)
- Screening also varies widely by state, with prevalence in 2020 ranging from 53% in California to 70% in the District of Columbia and Massachusetts
- A recent study based on BRFSS data found that, in contrast to Pandemic related declines from 2018 to 2020 for breast and cervical cancer screening, CRC screening remained steady overall because a 16% decline in colonoscopy was counterbalanced by a 7% increase in stool testing.

Summary (2)

- Observational studies suggest that colonoscopy reduces CRC incidence by about 40% and mortality by about 60%.
- Increasing screening rates to 80% would result in tens of thousands fewer CRC cases and deaths
- Although overall CRC mortality continues to decline, this progress is tempered by a rapidly changing landscape of disease that foreshadows less favorable trends ahead.
 - The CRC burden is shifting to younger individuals as cohorts with elevated risk born in the last half of the 20th century age; one in five new cases now occur in individuals in their early 50s or younger.
 - There is an overall shift to later stage disease, with more individuals now diagnosed at an advanced stage than in the mid-1990s before widespread screening.
 - Finally, there is a shift from right-sided to left-sided tumors, despite higher efficacy for preventing the latter through screening, likely reflecting changes in underlying disease risk of unknown etiology.



Summary (3)

- Although a substantial proportion of CRC deaths can be prevented through screening, 4 in 10 Americans aged 45 years and older are not up to date nationally, including one half of Californians.
- Screening is especially low among younger individuals and those without health insurance.
- Striking disparities by race and geography persist, with mortality rates in Alaska Natives almost three times higher than those in non-Hispanic White individuals
- Reducing CRC inequalities and furthering progress could be achieved by incentivizing healthier lifestyles, incentivizing organized screening, strengthening the role of the referring clinician, and ensuring equitable access to high-quality health care (screening and follow-up) for all individuals, especially those in rural and other low-resource areas.



Wisdom is knowing what to do. Virtue is doing it. --David Starr Jordon

Thank you







What do you view as the biggest barriers to patients in getting the 'unscreened' screened? (choose your top 3 barriers)

| What do you view as the biggest barriers to patients in getting the 'unscreened' screened? (choose your top 3 barriers) | |
|---|------|
| A. Lack of awareness | |
| | 67% |
| B. Fear | 54% |
| | 3476 |
| C. Distrust | 30% |
| D. Time (away from work or other responsibilities) | |
| | 48% |
| E. Transportation | |
| | 4% |
| F. Concern of prep (colonoscopy) | 30% |
| G. Dealing with stool (home test) | 50% |
| C. Dealing with stor (nome test) | 11% |
| H. Distrust of the test results | |
| | 0% |
| I. Cost | |
| | 26% |
| J. Other | 2% |
| | 2.76 |





Djenaba Joseph, MD, MPH CAPT, U.S. Public Health Service, Chief, Program Service Branch, Centers for Disease Control Prevention



Richard Wender, MD

Professor and Chair Family Medicine and Community Health University of Pennsylvania Evaluating Colorectal Cancer Screening Options: A Practical Framework

 $\bullet \bullet \bullet$

Richard C. Wender MD Djenaba Joseph MD MPH

Background assumptions

- Having a choice of tests is an asset that will increase screening rates.
- Every screening strategy relies on completion of colonoscopy to be effective.
- Every test has strengths and weaknesses.
- Consideration of cost is complicated.
 - Cost to whom?
 - Cost to individual vs. to the health care system as a whole.

Factors Influencing Choice of Test

Performance

 Sensitivity Specificity

Test characteristics

- Interval
- Accessibility
- Acceptability patient, provider
 Adherence initial test, repeat testing, follow-up colonoscopy

Contextual Factors

Stakeholder - Organization, provider, patient

Performance





Sensitivity

Sensitivity for cancer by stage

Sensitivity for earlier stages is given high priority

Sensitivity for advanced polyps
Performance of other tests is generally measured by comparison to what is found at colonoscopy

Specificity

Colonoscopy for everyone is accepted as a viable strategy in the U.S.

- Roughly 55% of individuals will have completely normal colons.
- Most of the polyps that are found and removed would not progress to cancer.
- Any non-invasive option, even those with relatively low specificity, will result in a substantial reduction in number of recommended colonoscopies.

Specificity – Quantifiable Tests

Value assigned to specificity is likely to vary by context and setting.

- Higher value in settings with lower colonoscopy access.
- Factors other than specificity are likely to predominate in settings with high colonoscopy access.

 Is there a minimally acceptable specificity level regardless of context and setting?

Test Characteristics





Access

- Covered by insurance dependent on USPSTF inclusion
- Widely/easily available
- Non-invasive vs Invasive
- Home vs Facility
 - Transportation
- Patient factors
 - Income
 - Insurance status
- Cost to the individual
- Cost to the practice or health system

Acceptability - patient

All roads lead to colonoscopy

Prep and potential barriers to access.

Stool tests

 Sample vs whole stool. Clarity of instructions. Navigation available?

Blood tests

- Coverage and cost will be key to acceptability.
- Transportation and time for testing will be barriers for some people.

Patient preference

Acceptability - provider

Provider <u>perceptions</u> of following characteristics

- Accuracy
- Effectiveness
- Availability
- Acceptability to patients
- Organizational burden, including test interval
- Incentive to achieve high practice-wide screening rate

Adherence and Interval

Interval

- More frequent testing requirement is a likely barrier to achieving high practice-wide screening rates
- Maintaining frequent (e.g. annual) screening requires a population-management strategy

Adherence

- Initial test
- Repeat testing
- Follow-up colonoscopy if indicated
- Ensuring high follow-up rate is a critically important measure of program effectiveness; hard to reach 80%

Contextual Factors

Stakeholder perspective

- Patient
- Provider
- Organization
- Geography/climate
- Availability of primary care and specialty services
- Population specific (unhoused, migrant)

Applying the Framework



| Sensitivity | Cologuard | Cologuard Plus | ColoSense | | |
|---|---------------------|---------------------|--|--|--|
| Test Type | Mt-sDNA | Mt-sDNA | Mt-sRNA | | |
| CRC overall | 92% | 94% | 94.4% | | |
| Stage I-IV | 90%, 100%, 90%, 75% | 87%, 94%, 97%, 100% | I/II – 92.3% III– 100% (no IV CRCs) | | |
| APL/AA | 42% (APL) | 43% (APL) | 42.9% (TSA/TA ≥10mm) 45.9% (advanced adenoma) | | |
| High grade dysplasia | 69% | 75% | 65.2% (HGD or ≥10 adenomas) | | |
| Sessile serrated | 42% | 46% | | | |
| APL = advanced precancerous lesion = Includes advanced adenomas (high-grade dysplasia or with ≥25% villous histologic features or measuring ≥1 cm in the greatest dimension) and sessile serrated polyps measuring 1 cm or more in diameter | | | | | |

AA = Advanced Adenoma

| Sensitivity | Guardant (Shield) | Polymedco OC Auto and Light | Colonoscopy | Freenome |
|-------------------------|--|--------------------------------|-------------------------------|--------------------------|
| Test Type | Cell-free DNA blood test | FIT | Endoscopy | Blood |
| CRC overall | 83% | 73% - 90% | 18% - 100% | 79.2% |
| Stage I-IV | 65% (55% clinical), 100%, 100% , 100% | _ | _ | 57.1%, 100%, 82.4%, 100% |
| APL/AA | 13.2% | 23% - 33% (AA) | 89% - 95% (adenomas ≥10mm) | 12.5% (AA) |
| High grade dysplasia | _ | _ | _ | 29% |
| Sessile serrated | _ | _ | _ | _ |

APL = advanced precancerous lesion = Includes advanced adenomas (high-grade dysplasia or with \geq 25% villous histologic features or measuring \geq 1 cm in the greatest dimension) and sessile serrated polyps measuring 1 cm or more in diameter AA = Advanced Adenoma



| Specificity | All | Negative Colonoscopy |
|-------------------|---|----------------------|
| Cologuard | 87% | 93% |
| Cologuard Plus | 91% | 93% |
| ColoSense | 86% | 88% |
| Guardant (Shield) | 90% (advanced neoplasia) | _ |
| Polymedco | 93% - 95% | _ |
| Colonoscopy | 89% ** | _ |
| Freenome | 92% (non-advanced colorectal neoplasia) | _ |



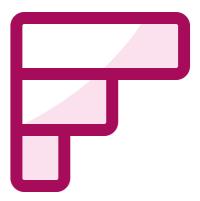
| | Interval (years) | Adherence (%) | Follow-up colo | Access | Cost |
|----------------|------------------|---------------------------|--|-----------------------------|------------------|
| Cologuard | 3 (1-3) | 51% - 71% (real-world) | 71.5% – 84.9% (real-world) | Widely available/covered | \$508 (Medicare) |
| Cologuard Plus | 3 (anticipated) | _ | _ | Not currently available | _ |
| ColoSense | 3 (anticipated) | 80% (study) | 80% 73% combined test and follow up (study) | Not currently available | Unknown |



| | Interval (years) | Adherence (%) | Follow-up colo | Access | Cost |
|-------------------|------------------|---------------|----------------|-----------------------------------|------------------------------|
| Guardant (Shield) | 1-3 (TBD) | 96% (study) | 44% (study) | Not currently available | \$895 (cash) |
| Polymedco | 1 | Varies | 58% - 83% | Widely available/covered | \$18 — \$21 |
| Colonoscopy | 10 | 55-60% | n/a | Covered, variable availability | \$2,750 (avg. cash price) |
| Freenome | 3 | 96% (study) | ?? | Not currently available | ?? |







Which of these three attributes do you view as most important? (rank in order of importance)



1.62

1.25

Which of these three attributes do you view as most important? (rank in order of importance)

- 1. B. Adherence
- 2. A. Robust data



3. C. Access



Screening Modality Expert Panel

 $\bullet \bullet \bullet$



Steven Itzkowitz, MD, FACP, FACG, AGAF

Professor of Medicine, Oncological Sciences and Medical Education, Ichan School of Medicine at Mount Sinai



Todd Kelley, MD Vice President of Medical Affairs, Polymedco



TR Levin, MD, MS Associate Director for Cancer

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Erica Barnell, MD, PhD Chief Medical Officer and Co-Founder, Geneoscopy



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Lance Baldo, MD Chief Medical Officer, Freenome



Craig Eagle, MD Chief Medical Officer, Guardant





(i) Start presenting to display the audience questions on this slide.

Closing Remarks and Next Steps

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Cocktail Reception Please join us across the hall