

# **Evaluation of New Tests Working Group: Modeling and Use of Surrogate Endpoints**

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# Early-stage proxies / surrogates for:

- Long-term effectiveness?
  - (Number needed to scope to detect 1 AN?)
- Programmatic cost-effectiveness?
  - (Cost to detect 1 AN?)

Prevalence as in	CRC	APL	NAA	Normal	Total cohort
Imperiale*	65	758	2,896	6,281	10,000

<sup>\*</sup> Imperiale *et al*, NEJM 2014; 370:1287

Test	Sens CRC	Sens APL	Sens NAA	Spec = 1- FP in normal	Interval	Test cost
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	De	Detected/Sent to colonoscopy				<u>Surrogate</u>	
n						NNS/CRC,APL	
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n	<u>48</u>	<u>182</u>	232	251	9,287	NNS/CRC,APL <b>3.1</b>	
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Cost	F	-IT + Colo T	×	FIT + Colo Dx	FIT	Cost/CRC,APL \$3,800	

<sup>\*</sup> Imperiale *et al*, NEJM 2014; 370:1287

### A simple calculator in Excel for Round 1 proxies

			Spec = 1- FP				
Sens CRC	Sens APL	Sens NAA	in normal	Interval	Test cost	Colo Dx	Colo Tx
0.74	0.24	0.08	0.96	1	\$18	\$740	\$1,083
CRC	APL	NAA	Normal	Total cohort			
65	758	2,896	6,281	10,000			
						NNS for 1	Cost for
CRC	APL	NAA	Normal	No scope		CRC/APL	CRC/AP
48	182	232	251	9,287		3.1	
\$52,958	\$200,294	\$255,080	\$190,440	\$167,167			\$3,765
	0.74 CRC 65 CRC 48	0.74  0.24    CRC  APL    65  758    CRC  APL    48  182	0.74    0.24    0.08      CRC    APL    NAA      65    758    2,896      CRC    APL    NAA      48    182    232	Sens CRC      Sens APL      Sens NAA      in normal        0.74      0.24      0.08      0.96        CRC      APL      NAA      Normal        65      758      2,896      6,281        CRC      APL      NAA      Normal        48      182      232      251	Sens CRC      Sens APL      Sens NAA      in normal      Interval        0.74      0.24      0.08      0.96      1        CRC      APL      NAA      Normal      Total cohort        65      758      2,896      6,281      10,000        CRC      APL      NAA      Normal      No scope        48      182      232      251      9,287	Sens CRC      Sens APL      Sens NAA      in normal in no	Sens CRC      Sens APL      Sens NAA      in normal      Interval      Test cost      Colo Dx        0.74      0.24      0.08      0.96      1      \$18      \$740        CRC      APL      NAA      Normal      Total cohort      Colo Dx      Normal      Normal      Noscope      NNS for 1 CRC/APL        CRC      APL      NAA      Normal      No scope      CRC/APL      3.1

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# How do proxies compare to long-term estimates from 4 models (3 CISNET models, MOSAIC)?

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- Collaboration: US, Netherlands, Israel
- Luuk van Duuren, Iris Lansdorp-Vogelaar,
  Elizabeth Half, Zohar Levi, Barbara Silverman









### PREVENTION AND EARLY DETECTION

# Effectiveness and Cost-Effectiveness of Colorectal Cancer Screening With a Blood Test That Meets the Centers for Medicare & Medicaid Services Coverage Decision



Rosita van den Puttelaar,<sup>1</sup> Pedro Nascimento de Lima,<sup>2</sup> Amy B. Knudsen,<sup>3</sup> Carolyn M. Rutter,<sup>4</sup> Karen M. Kuntz,<sup>5</sup> Lucie de Jonge,<sup>1</sup> Fernando Alarid Escudero,<sup>6</sup> David Lieberman,<sup>7</sup> Ann G. Zauber,<sup>8</sup> Anne I. Hahn,<sup>8</sup> John M. Inadomi,<sup>9</sup> and Iris Lansdorp-Vogelaar<sup>1</sup>

# Comparative Effectiveness and Cost-Effectiveness of Colorectal Cancer Screening With Blood-Based Biomarkers (Liquid Biopsy) vs Fecal Tests or Colonoscopy

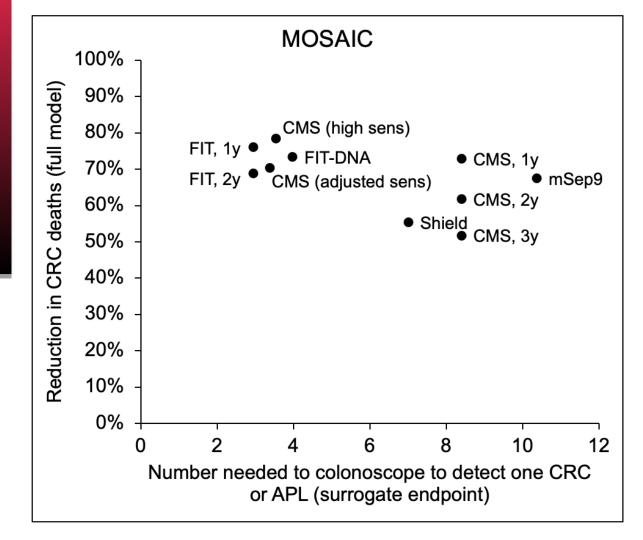


Uri Ladabaum,<sup>1,2</sup> Ajitha Mannalithara,<sup>1,2</sup> Yingjie Weng,<sup>2,3</sup> Robert E. Schoen,<sup>4</sup> Jason A. Dominitz,<sup>5,6</sup> Manisha Desai,<sup>2,3</sup> and David Lieberman<sup>7</sup>

#### NNS to detect 1 AN

VS.

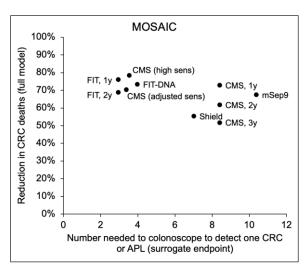
CRC death reduction

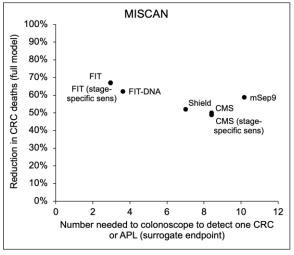


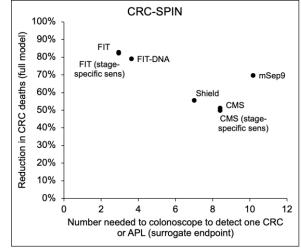
#### NNS to detect 1 AN

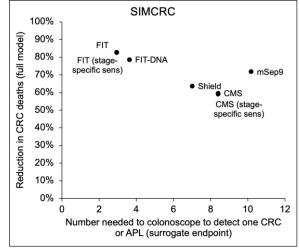
VS.

# CRC death reduction







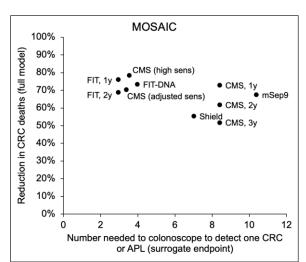


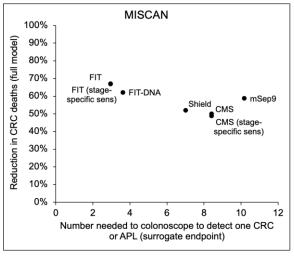
#### NNS to detect 1 AN

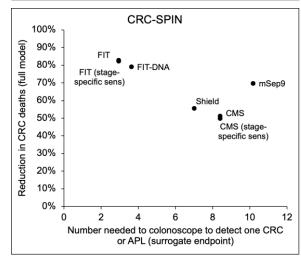
VS.

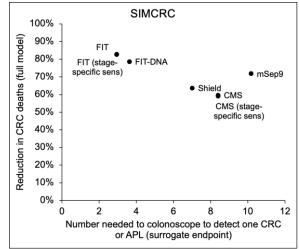
# CRC death reduction

- Fails if low specificity
- Does not reflect interval





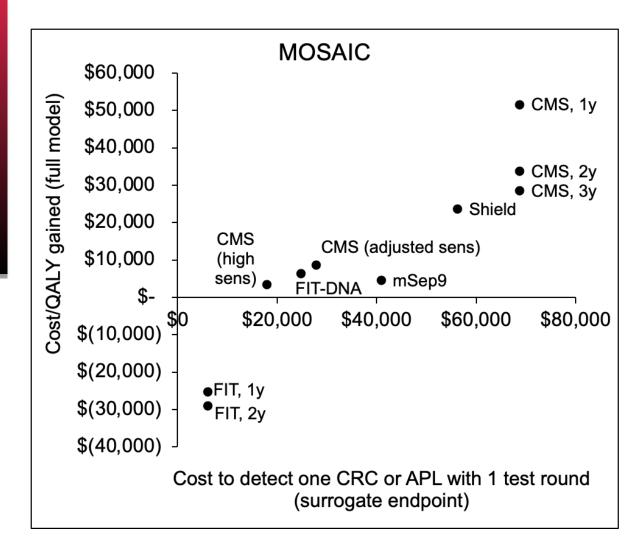




#### Cost to detect 1 AN

VS.

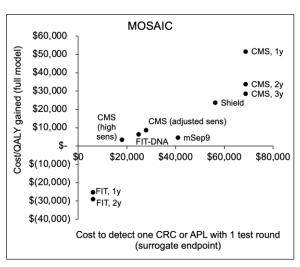
Cost/QALY gained

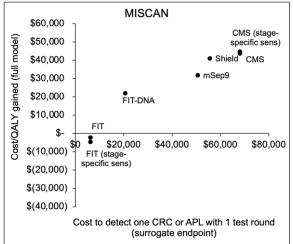


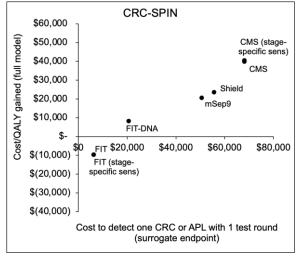
#### Cost to detect 1 AN

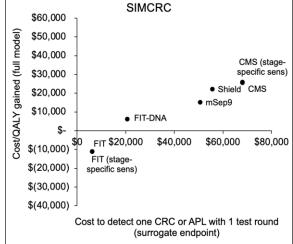
VS.

## Cost/QALY gained







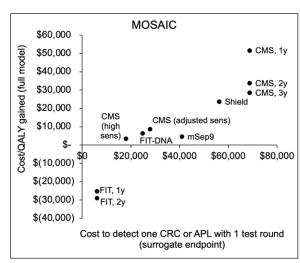


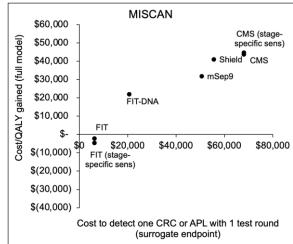
#### Cost to detect 1 AN

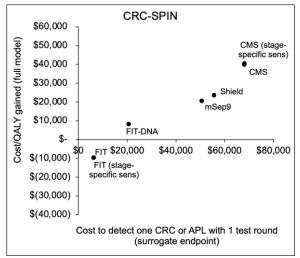
VS.

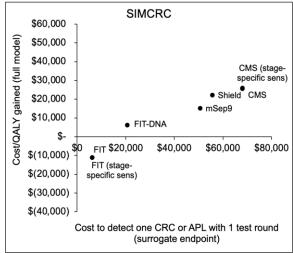
### Cost/QALY gained

 Does not reflect interval









# Discussion: Surrogate endpoints in modeling

- Number needed to colonoscope to detect one CRC/APL in round 1 of screening:
  - PROMISING proxy for effectiveness
- Cost to detect one CRC or APL in round 1 of screening:
  - PROMISING proxy for cost-effectiveness
- Further validation needed
- Incorporate in early phases of test development?