The Principles of Screening

National Academy of Medicine Workshop

Challenges in Initiating and Conducting Long-Term Health Monitoring of Population Following Nuclear and Radiological Emergencies in the United States

Washington, DC

March 12, 2019

Steven H. Woolf, MD, MPH
Professor of Family Medicine and Population Health
Virginia Commonwealth University

Definitions

- Primary prevention
- Secondary prevention
- Tertiary prevention

- Burden of suffering
- Accuracy and reliability
- Effectiveness of early detection
- Harms
- Balance of benefits and harms

Burden of Suffering

- Frequency: incidence, prevalence
- Severity: morbidity, mortality
- Clinical significance

- Burden of suffering
- Accuracy and reliability
- Effectiveness of early detection
- Harms
- Balance of benefits and harms

Accuracy and Reliability

- Accuracy
 - Sensitivity (proportion with disease who test positive)
 - Specificity (proportion without disease who test negative)
 - Positive predictive value (proportion who test positive who have the disease)
- Reliability

Positive Predictive Value and Prevalence

Prevalence = 7%

Sensitivity=100%, Specificity=98%

<u>Ultrasound</u>	<u>Cancer</u>	No Cancer	<u>Total</u>	<u>PPV</u>
Positive	7000	1860	8860	79%
Negative	0	91,140	91,140	
Total	7000	93,000	100,000	

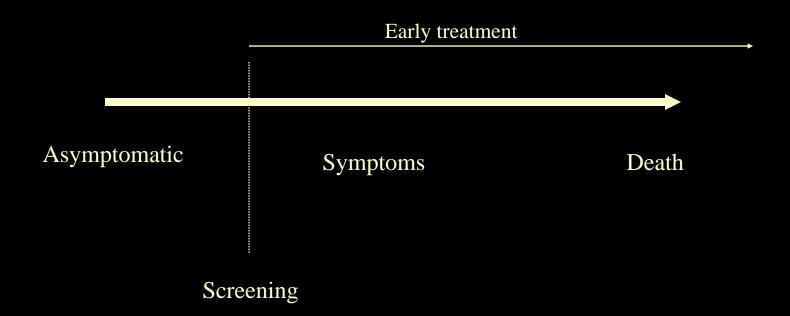
Positive Predictive Value and Prevalence

Prevalence = 0.07% Sensitivity=100%, Specificity=98%

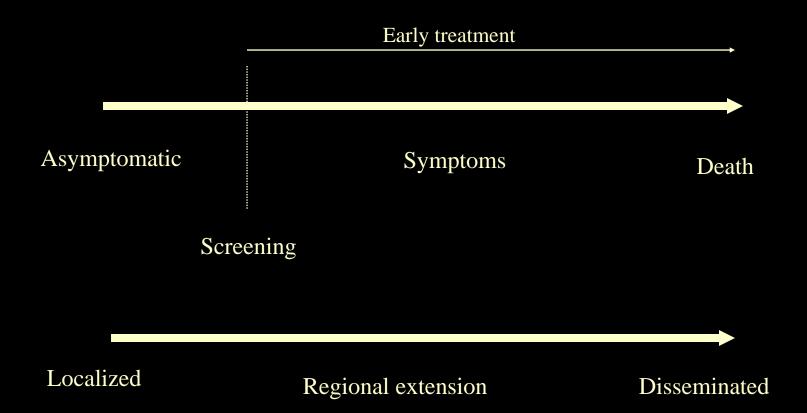
<u>Ultrasound</u>	Cancer	No Cancer	<u>Total</u>	<u>PPV</u>
Positive	70	1999	2069	3%
Negative	0	97,931	97,931	
Total	70	99,930	100,000	

- Burden of suffering
- Accuracy and reliability
- Effectiveness of early detection
- Harms
- Balance of benefits and harms

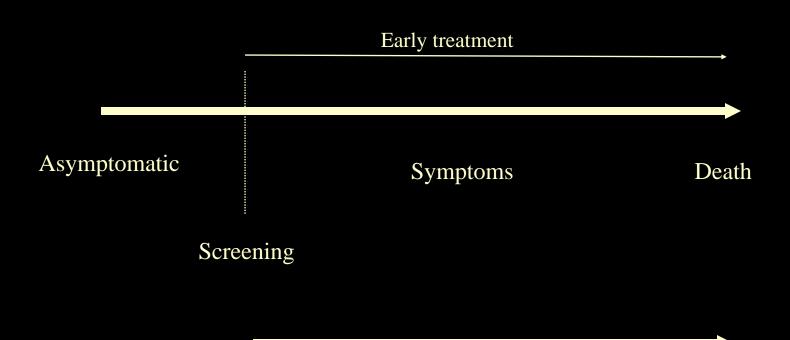
Rationale for Early Detection



Stage Shift



Lead-Time Bias



Confirming Effectiveness

- Randomized controlled trials
- Well-designed observational studies
- Relative vs. absolute benefit
- Number-needed-to-screen
- Optimal interval
- When to stop
- Selective vs universal screening

Relative versus Absolute Benefit

Example:

"drug X reduces incidence of CRF by 20%"

If baseline risk of CRF is 1:10,000 (0.010%), drug X decreases incidence to 1:12,000 (0.008%) =

Absolute reduction of 0.002%

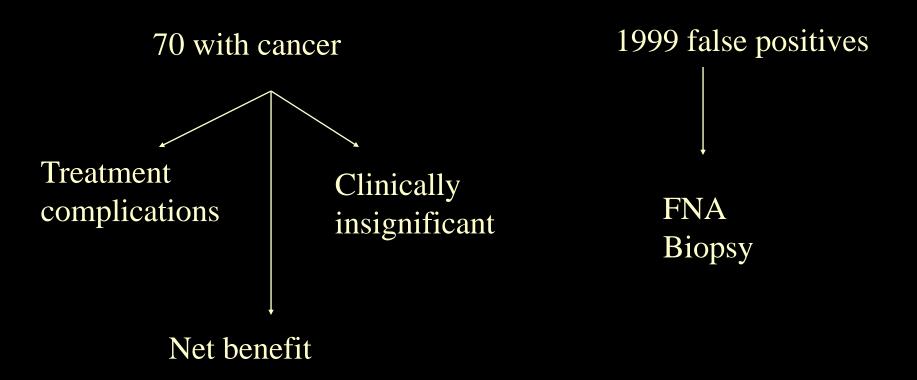
NNT = 100/0.002 = 50,000

- Burden of suffering
- Accuracy and reliability
- Effectiveness of early detection
- Harms
- Balance of benefits and harms

Harms of Screening

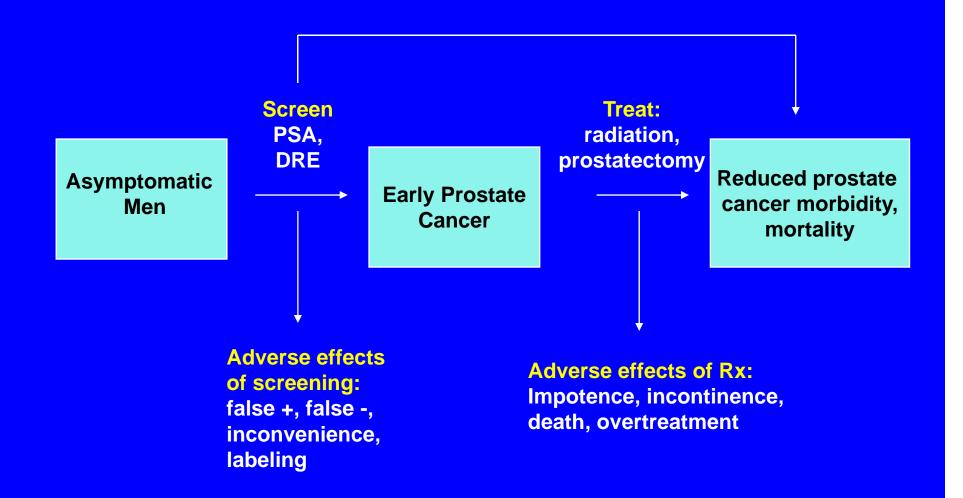
- Test procedure
- Anxiety and labeling effects
- False-positive results
- Harms of treatment

Potential Harms



- Burden of suffering
- Accuracy and reliability
- Effectiveness of early detection
- Harms
- Balance of benefits and harms

Logic for Screening



Balance of Benefits and Harms

- Objective component
- Subjective component

- Resources
- Feasibility
- Politics and public expectations
- Ethical and legal factors

Who is on the Guideline Panel?

- Topic experts and specialists vs. generalists and experts in analytic science
 - Conflicts of interest
 - Intellectual
 - Financial
 - Referal bias