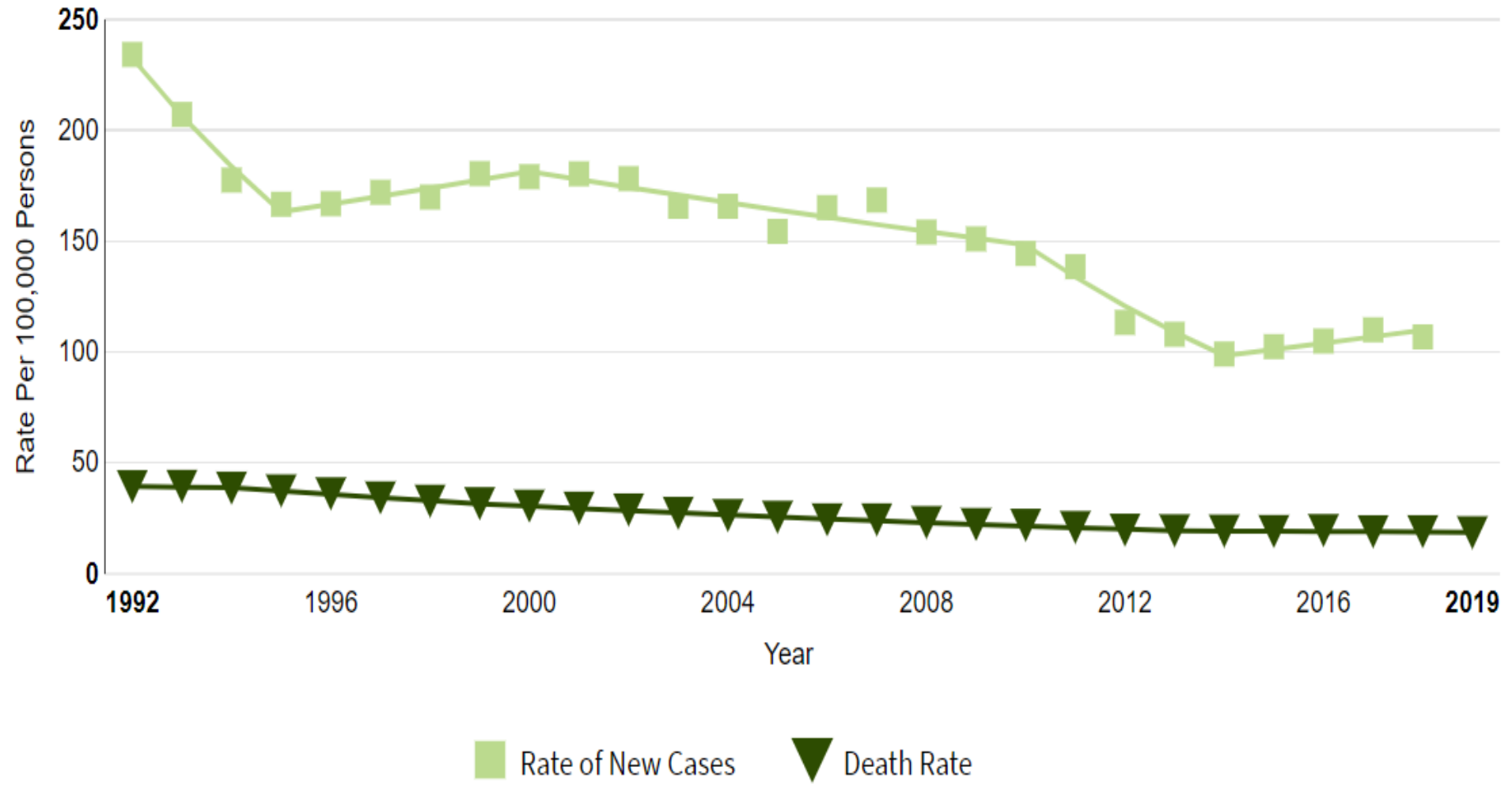


The Appropriate Use of PSA Screening and Guidelines

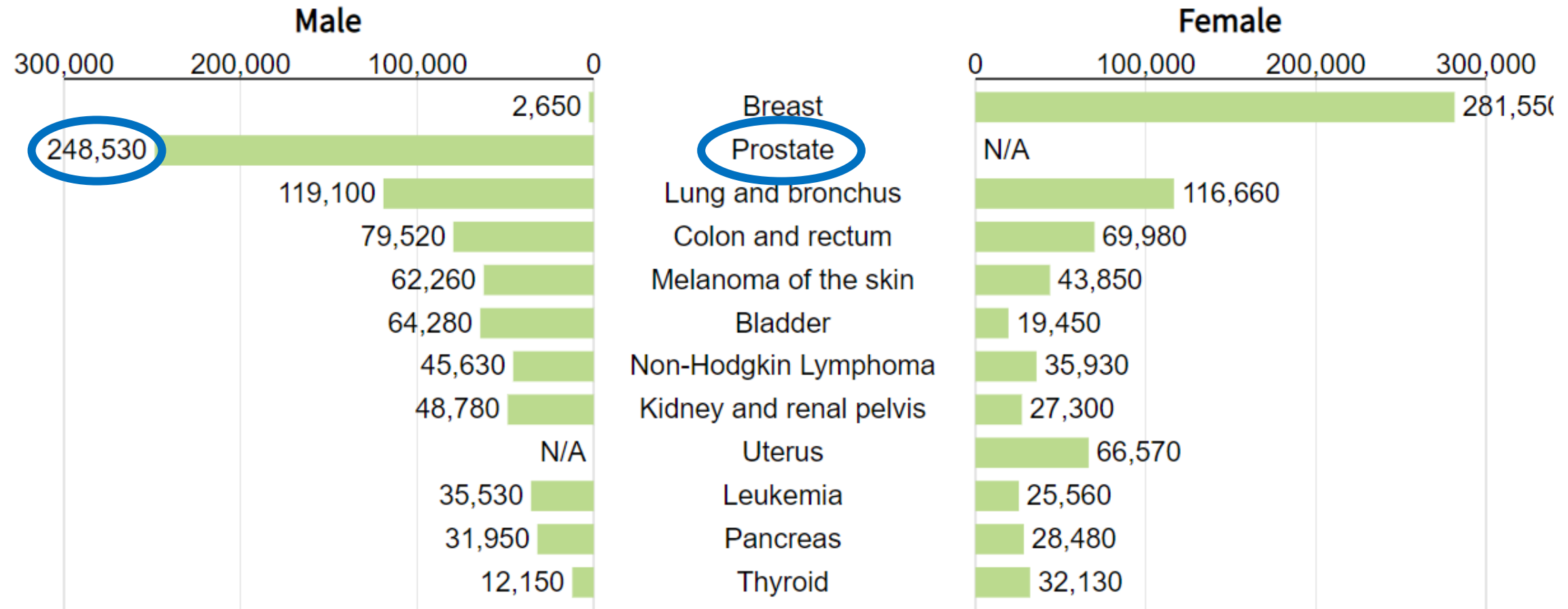
Jared Cox, *M.D.*

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SEER Data

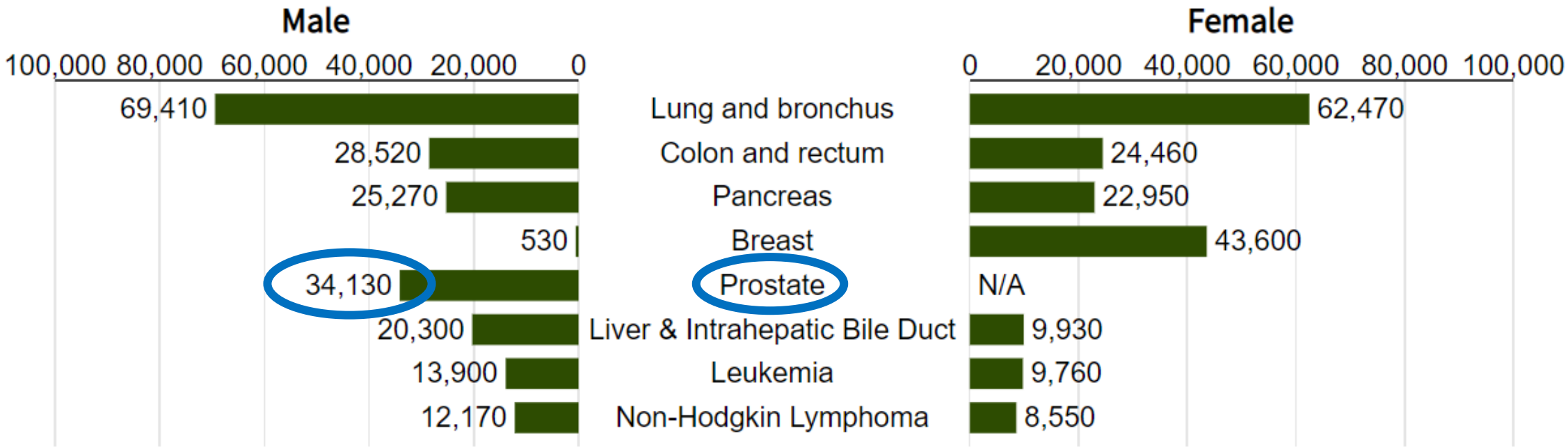


The top 12 most common cancer sites, shown below, will account for more than three quarters of all new cancer cases.



Source: Cancer Facts & Figures 2021, American Cancer Society (ACS), Atlanta, Georgia, 2021.

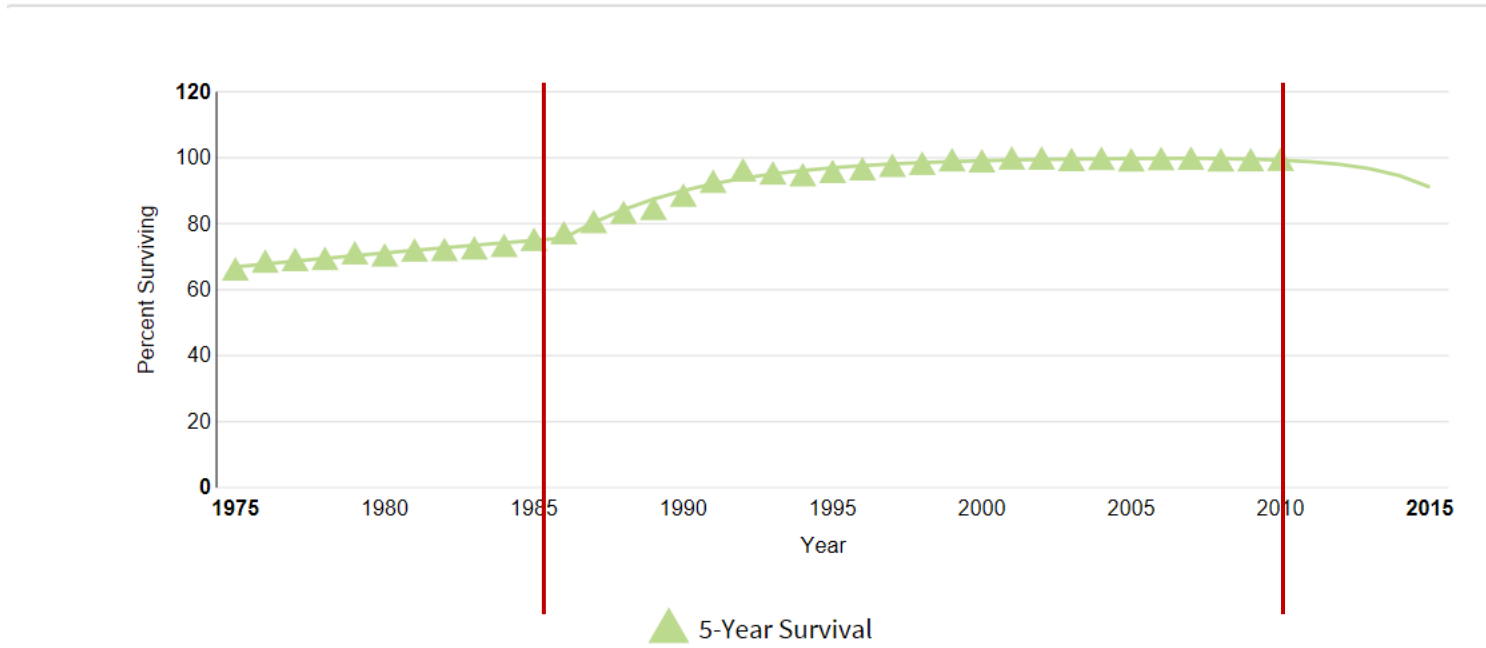
The eight deadliest cancer sites, shown below, will account for almost two-thirds of all expected cancer deaths.



Source: Cancer Facts & Figures 2021, American Cancer Society (ACS), Atlanta, Georgia, 2021.

1985			
	Data	Observed Rate	Modeled Rate
■	5-Year Survival	75.3	74.9

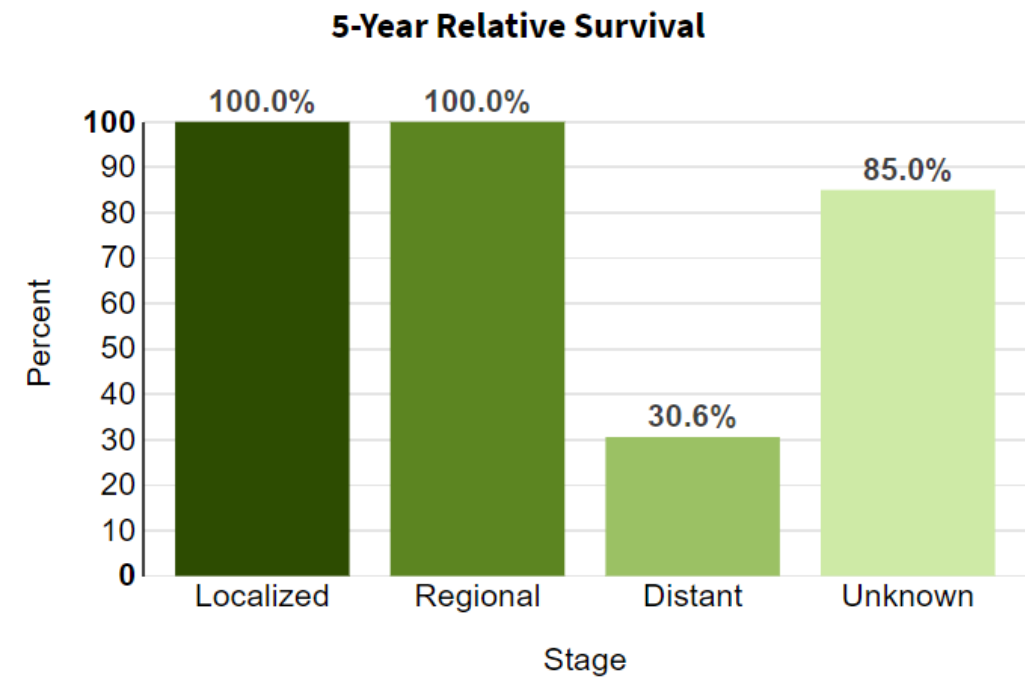
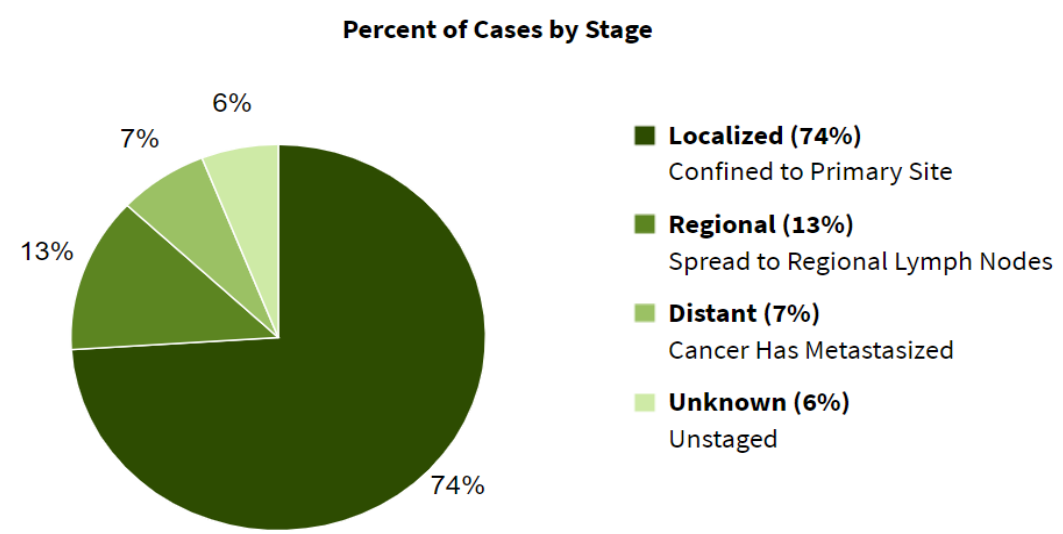
2010			
	Data	Observed Rate	Modeled Rate
■	5-Year Survival	99.6	99.3



• Cancer Trends

- Pre-PSA era
 - 75% with extraprostatic disease
 - 35% with metastatic disease
- Post-PSA era
 - 15% with extraprostatic disease
 - 4% with metastatic disease

Percent of Cases & 5-Year Relative Survival by Stage at Diagnosis: Prostate Cancer





• Before Starting

- Assessment:
 - General health
 - Medical Comorbidities
 - Life Expectancy
 - Patient Preferences
- Family history
 - 1st degree relative increases risk of prostate cancer 2.1-2.5-fold.
 - Risk higher with more affected relatives
 - Breast cancer history in family members



• Before Starting

- African Ancestry
 - 60% higher incidence
 - 38% higher mortality
 - Earlier onset
- Genetic Syndromes/Germline Mutations
 - HOX genes
 - BRCA 1/2 or other DNA repair mutations
 - Lynch Syndrome



• PSA

- Glycoprotein, protease, produced by prostate epithelial cells
- Not cancer specific
- Levels may be affected by:
 - instrumentation, infection, ejaculation, trauma, urinary retention, 5-ARIs, OTC supplements
- Different assays not necessarily directly comparable (calibration differences)
- Empiric antibiotics DO NOT have a clinical benefit in elevated PSA workup of an asymptomatic man
- Abnormal level always warrants a repeat

• DRE

- Strongly recommended in guidelines, not required
- High risk disease increased 2.7 fold with abnormal DRE
- No significant value as a stand-alone test; best used with PSA
- In an analysis of 166,104 individuals with prostate cancer diagnosed between 2004-2007 from SEER database, only 685 (0.4%) had a palpable, PSA-occult (level <2.5), Grade Group ≥ 4 disease. (Hattangadi et al, 2012)



• Practical Considerations

- Age to Initiate Screening
 - Screening studies did not include younger men
 - Baseline PSA in 40s and early 50s can give risk stratification
 - Higher baseline levels in this age group can predict death from prostate cancer
- Frequency of Testing
 - Can be determined by PSA level at early age
 - Lower frequency with lower PSA levels

• Practical Considerations

- Age to Discontinue Screening
 - >75 years old? If healthy or >10-year life expectancy
- Risk of biopsy
 - UTI, epididymitis, orchitis, prostatitis, sepsis
 - Rectal bleeding, pain, hematuria, vasovagal episode, fever, hematospermia, dysuria
 - Transperineal option?



• Other Considerations

- PSA derivatives
 - Age specific ranges – does this work?
 - PSAV (Velocity)
 - % free PSA or cPSA
 - PSAD (Density)
 - Prostate Health Index (PHI)
 - Iso-PSA



• Other Considerations

- Biomarker Testing
 - PCA-3
 - 4k score
 - Select MDx
 - Confirm MDX
 - Exo Dx IntelliScore (EPI score)
- Imaging
 - MRI
 - Other

• Guidelines

- Everyone has different recommendations for screening
 - AUA
 - NCCN
 - ASCO
 - ACS
 - USPSTF
 - EAU/ESTRO

• Screening Guidelines

- All Agree on ONE THING:

PSA-based prostate cancer screening requires an informed, shared decision-making process, and the decision should reflect the patient's understanding of the possible risks and benefits and should reflect the patient's preferences and values

- All Differ in:

- Whether or not routine screening is recommended
- What age groups and life expectancies
- What intervals

NCCN Guidelines Version 2.2018

Prostate Cancer Early Detection

BASELINE EVALUATION

- History and physical (H&P) including:
 - ▶ Family cancer history
 - ▶ Medications^a
 - ▶ History of prostate disease and screening, including prior PSA and/or isoforms, exams, and biopsies
 - ▶ Race^b
 - ▶ Family or personal history of high-risk germline mutations^c

RISK ASSESSMENT

- Start risk and benefit discussion about offering prostate screening:
- Baseline prostate-specific antigen (PSA)^d
 - Strongly consider baseline digital rectal examination (DRE)^d

Age 45–75 y

Age >75 y, in select patients (category 2B)^e

EARLY DETECTION EVALUATION

PSA <1 ng/mL,
DRE normal (if done)

PSA 1–3 ng/mL,^f
DRE normal (if done)

PSA >3 ng/mL^f
or very suspicious DRE

PSA <4 ng/mL, DRE normal
(if done), and no other
indications for biopsy

PSA ≥4 ng/mL or very
suspicious DRE

Not screened^e

Repeat testing at
2–4 year intervals^g

Repeat testing at
1–2 year intervals

[See Indications
for Biopsy \(PROSD-3\)](#)

Repeat testing in
select patients at
1–4 year intervals

[See Indications
for Biopsy \(PROSD-3\)](#)

INDICATIONS FOR BIOPSY^h

MANAGEMENT

- Repeat PSA
- DRE, if not performed during initial risk assessment
- Workup for benign disease

- Consider biomarkers that improve the specificity of screeningⁱ
- Consider multiparametric MRI^j

Transrectal ultrasound (TRUS)-guided biopsy^k

or

Follow-up in 6–12 mo with PSA/DRE^{i,l}

[See Management of Biopsy Results \(PROSD-4\)](#)

Summary

- Baseline PSA for 45-75 yo, DRE to complement
- Repeat elevated PSA to confirm
- Start at younger age (40 yo) for those with elevated risk
- PSA <1, consider testing every 2-4 years
- PSA 1-3, testing every 1-2 years
- PSA consider in those >75 yo who are healthy
- MRI if able/available
- Consider biomarkers
- **Need for biopsy – That's our job!**

● The Holy Grail for PCa Screening



- Screening/blood test with improved sensitivity and specificity
- Better image guidance for diagnosis and biopsy
- Improve on diagnostic and treatment invasiveness and risk
- DNA/RNA genomic profiling of tumors
- Goal - Delineate aggressive/dangerous tumors from indolent ones

THANK YOU

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