The Fifteen Stages

of PRESTATE CANCER

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Medical Director: Prostate Oncology Specialists, Inc. Executive Director: Prostate Cancer Research Institute Invasion of the Prostate Snatchers
The Key to Prostate Cancer



Patients Are Expected to Take Charge of the Decision Making Process

- No full-time experts in treatment selection
- Doctors have conflicts of interest
- Extremely long natural history of prostate cancer makes predictions about outcome difficult
- Treatment impacts quality-of-life more than survival

An 8-Question Staging Quiz pcri.org

Prostate Cancer Staging Guide

SKY | TEAL | AZURE | INDIGO | ROYAL

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CLICK HERE TO FIND YOUR STAGE

Click: "Take the Staging Quiz" on the PCRI home page

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Knowing Your Stage...

- Helps you speak the language of prostate cancer with your doctor
- Gives perspective on the risk of dying
- Allows you to avoid unessential information related to other stages
- Guides you to the correct treatment



The Risk of Dying

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Risk of Mortality

- · Sky
- · Teal
- Azure
- Indigo
- Royal

None

Moderate

Maximal

Mod. to Max.

Maximal

<1%

5%

10%

< 50%

> 50%

Note: These are cumulative statistics

The Timing of Mortality

American Cancer Society

		Year Diagnosed
· Five year	1%	2012
· Ten year	2%	2007
• Fifteen year	4%	2002
• More	7%	
• Never (from PC)	85%	
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The stats are pessimistic: Half of those who are diagnosed in 2017, who are destined to succumb, will benefit from the medical technology of 2032+

Other Staging Systems are Incomplete

- 1. Clinical: Digital rectal exam only
- 2. Pathologic: Surgical or biopsy findings only
- 3. TNM: 1 & 2 plus scans
- 4. <u>Risk Categories</u> (Low, Mod, High-Risk): 1 & 2 plus PSA
- 5. Stages of Blue: 1, 2, 3, 4 and whether or not there has been previous therapy

"Clinical" Stage or DRE Stage DRE = Digital Rectal Exam

	Stage	Description	Stage of Blue
	T1 :	Tumor that cannot be felt at all by DRE	Sky
THE REPORT OF THE PARTY OF THE	Т2:	Tumor confined within the prostate T2a: Tumor in < 50% of one lobe T2b: Tumor in > 50% of one lobe but not both lobes T2c: Tumor felt in both lobes	Teal Azure
	Т3:	Tumor that extends through the prostate capsule T3a: Extracapsular extension T3b: Tumor that invades seminal vesicle(s)	
	T4:	Tumor that invades rectum or bladder	•

What's Wrong with Risk Categories?

(Low-Risk, Intermediate-Risk, High-Risk)

- Doesn't include:
 - Multiparametric-MRI findings
 - Percentage biopsy-core findings
 - PET scan information
- Doesn't address relapsed and advanced Dz.
- Doesn't suggest a treatment schema
- Assigning a stage requires a professional

The Six Components of The Stages of Blue

(Risk Categories = The three factors above dotted line)

Digital rectal

Normal vs. nodule vs. mass

· Gleason Score

6 vs. 7 vs. 8-10

· PSA

<10 vs. 10-20 vs. over 20

Biopsy Cores

Few vs. mod. vs. many

Previous therapy

Yes vs. no

Imaging

Disease extent: nodes, bones

The Guts of the Five Stages of Blue

	Local Therapy	Gleason Score	Digital Rectal	PSA	MRI, CDU CT Scans	Bone Scan
5ky	No	< 6	Small nodule or no nodule	<10	No ECE	No Need
Teal	No	7	Larger nodule on one side	10-20	No ECE	Clear
Azure	No	8-10	Bilateral nodule, ECE or SV	>20	ECE, SV, Pelvic Node	Clear
Indigo	Yes	Any	Any	Rising	Pelvic Node	Clear
Royal ¹	Any	Any	Any	>100	Other Node	Positive

^{1.} Any rising PSA with a low testosterone bumps to Royal ECE = Extra-capsular Extension SV = Seminal Vesicle



The Characteristics of SKY

"Low-Risk"

- No previous therapy
- Small nodule or no nodule
- Gleason 6
- PSA under 10
- State-of-the art multiparametric-MRI (MP-MRI) showing no high-grade disease

Three Types of Sky

- Low-Sky = Two or fewer biopsy cores with cancer, no core more than 50% replaced and PSA density under 0.15 (PSA density is PSA divided by prostate size in cc)
- 2. Basic-Sky = Same as Low-Sky except can have up to 50% of core biopsies positive, but no core more than 50% replaced with cancer
- 3. High-Sky = More than 50% cores positive, or a palpable nodule (T2a) or PSA density > 0.15

Treatment for Sky

1. Low-Sky = Active Surveillance

2. Basic-Sky = Active Surveillance

3. High-Sky = Active Surveillance

The TEAL Stage "Intermediate Risk"

- No previous treatment
- Grade 7, and /or
- PSA 10 to 20 and / or
- Larger, unilateral nodule (T2b)
- MP-MRI: no unexpected high-grade disease
- Bone scan clear

Three Types of TEAL

4. Low-Teal

- Gleason 3 + 4 = 7, with < 20% grade 4
- Only 2 biopsy cores positive, each with < 50% cancer
- Only one Teal factor; all other factors are Sky

5. Basic-Teal

- Like Low-Teal but up to 50% of cores positive

6. High-Teal

- Gleason 4 + 3 = 7
- Over 50% of core biopsies positive
- Two or more **Teal** factors

Treatment for TEAL

- 4. Low-Teal = Active Surveillance
- 5. Basic-Teal = Monotherapy with Seeds or IMRT or Proton or SBRT or Surgery or hormonal therapy
- 6. High-Teal = Combo therapy with Seeds + IMRT + Hormones for 4-6 months

The Azure Stage "High-Risk"

- No previous treatment
- Gleason 8,9, 10 and / or
- PSA over 20 (and < 100) and / or</p>
- MP-MRI with ECE, SVI or Node Mets
- Bone scan clear

Three Types of Azure

- 7. Low-Azure = Small 4+4=8 tumor, PSA under 10, only a couple cores positive and favorable multiparametric-MRI findings
- Basic-Azure = Neither Low or High (most men in Azure are in the Basic subtype)
- 9. High-Azure = Gleason score of 9 or 10 and /or seminal vesicle invasion and /or pelvic lymph node metastases and / or PSA above 40

Treatment for Azure

7. Low-Azure

 Similar to High-Teal: Seeds + IMRT + hormone therapy for four to six months

8, Basic-Azure

 Seeds + IMRT to prostate and pelvic nodes + hormone therapy for 18 months

9. High-Azure

 Same as Basic-Azure with the additional consideration for adding Zytiga, Xtandi or Taxotere



The INDIGO Stage

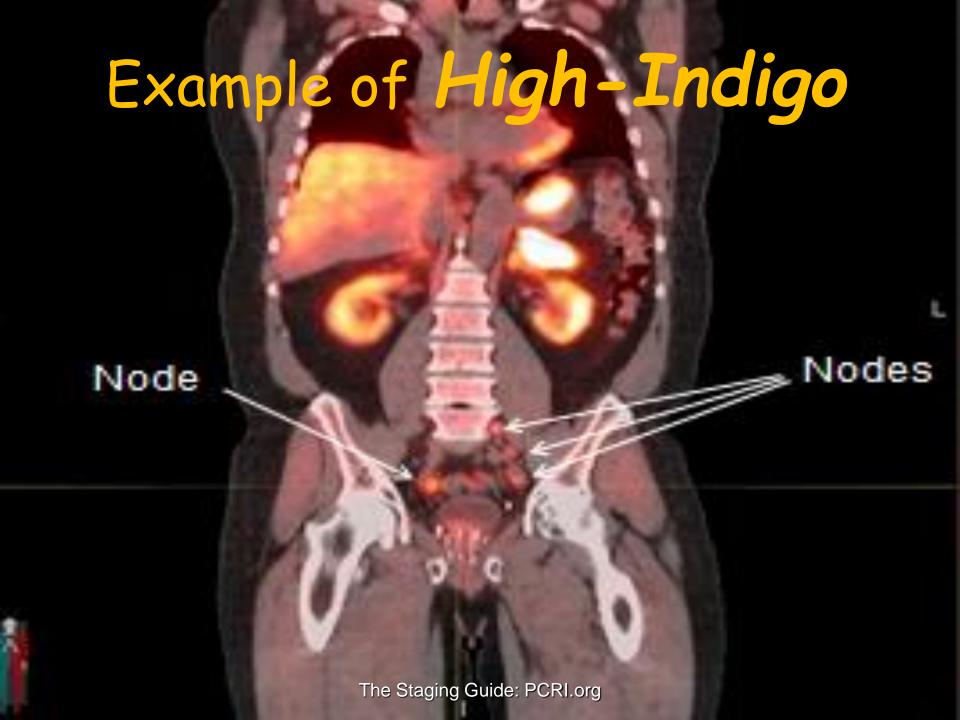
- A cancer recurrence after therapy:
 - Positive margin after surgery
 - Rising PSA (from cancer)
 - Biopsy positive prostate or nodes
 - Scan positive nodes
- No metastases beyond the pelvic nodes

Three Types of Indigo

- 10.Low-Indigo = Recurrence in the prostate or prostate fossa only
- 11 Basic-Indigo = There is a significant risk of microscopic pelvic lymph node metastases, but node mets are unproven
- 12. High-Indigo = Scans or biopsy show unequivocal pelvic node metastases

Treatment for INDIGO

- 10.Low-Indigo = IMRT to prostate fossa after previous surgery or cryotherapy after previous XRT
- 11 Basic Indigo = Same as Low-Indigo but with the addition of hormones for 6 to 12 months & pelvic node XRT.
- 12. High-Indigo = Same as Basic-Indigo with the addition of Zytiga, Xtandi and /or Taxotere



The ROYAL Stage

- Hormone resistance, which is:
 - A rising PSA
 - Testosterone < 50
- Metastases beyond pelvic nodes

Three types of ROYAL

- 13.Low-Royal = Hormone resistant, but scans show no metastases anywhere
- 14. Basic-Royal = Less than 6 metastases with at least one beyond the pelvic nodes
- 15. High-Royal = More than 5 metastases with at least one beyond the pelvic nodes

Treatment for Royal

- 13. Low-Royal = Use PET scans. Find the metastases! Once you find the cancer treat it like Basic-Royal
- 14. Basic-Royal = Provenge followed by Zytiga or Xtandi with IMRT or SBRT to all metastatic sites. (If hormone sensitive, consider early Taxotere)
- 15. High-Royal = Next slide

Treatment Principles: High-ROYAL

- Consider aggressive, early combination therapy with Provenge, Zytiga or Xtandi.
- Monitor PSA and scans closely. Goal is to achieve an undetectable PSA
- Change to Xofigo or Taxotere at the first sign of cancer progression

Don't Wait Till It's Too Late

When Provenge was administered to men before the PSA rose above 22, the average improvement in survival was <u>13 months</u> compared to only <u>4 months</u> when treatment was delayed.

Fear Cause Brain Freeze

- Prostate Cancer
- Racists
- Radiation
- · Gang members
- · Chemotherapy
- · Nazis
- · Impotence

Knowledge Combats Fear, But There Are Daunting Barriers:

- Prostate cancer is intricate and can be hard to understand
 - There is endless information on the internet
 - Much is outdated—old internet info never dies
 - Technology is evolving quickly
- Facetime with doctors comes at a premium
- Doctors are trained in only one type of therapy

What Exactly Are We Afraid of?

- Early mortality
- Loss of quality of life (pain and suffering)
- Being overwhelmed by too much information
- Mismanaging and screwing up:
 - It's painful when something bad happens
 - It's worse when you don't know what to do
 - It's even worse if the problems could have been prevented with better management

What Should We Be Afraid of:

- Fear itself: Frightened people make bad choices
- Too much respect for authority: This is a high stakes game with your longevity and quality of life at stake. Don't just do what you are told
- Laziness or intimidation toward the study process
- Impatience; treatment side effects are potentially irreversible. Usually there is plenty of time to educate yourself before embarking on therapy.

Requirements for Optimal Learning

- Set aside quality time
- Believe in yourself; you can do it
- Understand that learning is a process
- Limit your focus to only what you need
- Feed on accurate, up-to-date information
- Focus on facts, not fearful feelings

Focus Alleviates Pressure

- Identify your stage and substage by using the Staging Guide available at pcri.org
- Become very familiar with the common therapies used for your particular stage
- Consult one or more doctors <u>after</u> you have done your homework. Your are expected to voice your preferences.

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Prostate Cancer Staging Guide

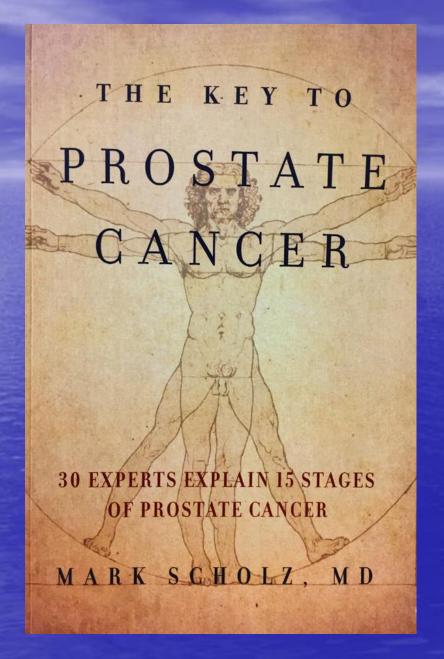
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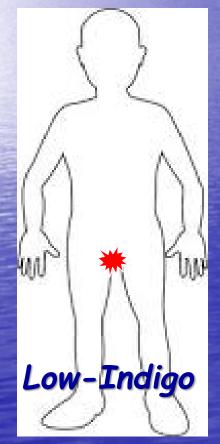
Epstein Score Clarifies Risk

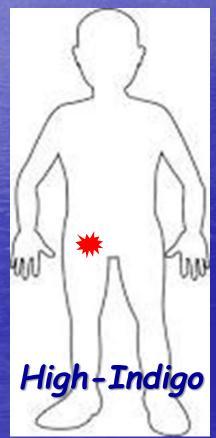
Epstein Score (1 to 5)	Implications
1 = Gleason 3 + 3 or less	Harmless
2 = Gleason 3 + 4	Barely consequential
3 = Gleason 4 + 3	Consequential
4 = Gleason 4 + 4	Worrisome
5 = Gleason 4 + 5 or more	Twice as worrisome

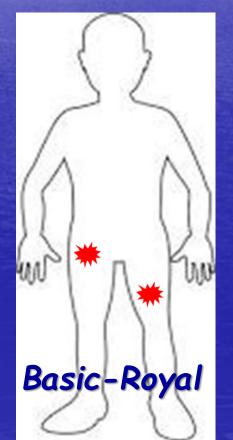
Extent of Disease Per Scans

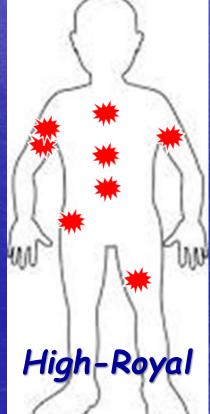
INDIGO

ROYAL



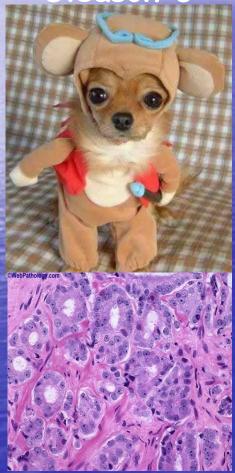




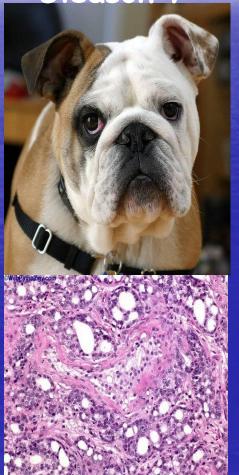


Dogs and Prostate Cancer

Epstein 1 Gleason 6



Epstein 2,3
Gleason 7



The Staging Guide: PCRI.org

Epstein 4,5 Gleason 8-10



How to Subtype Indigo

- 10.Low-Indigo = PSA <0.5 after surgery or if < 5.0 after XRT; PSA-DT > 8 months and previous stage (prior to relapse) was Basic-Teal or less
- 11 Basic-Indigo = Fails to meet criteria of either Low-Indigo or High-Indigo. Such patients have a higher likelihood of pelvic micro-mets.
- 12. High-Indigo = Unequivocal pelvic node metastases have been documented with scans or by biopsy (without metastases anywhere else in the body)

