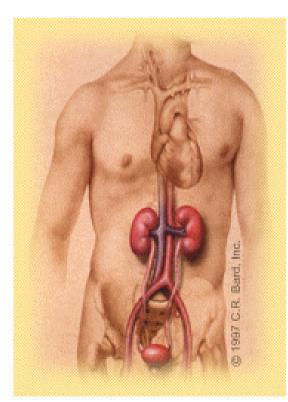
25TH ICRO DEHRADUN STAGING OF GENITOURINARY MALIGNANCIES



SPEAKER

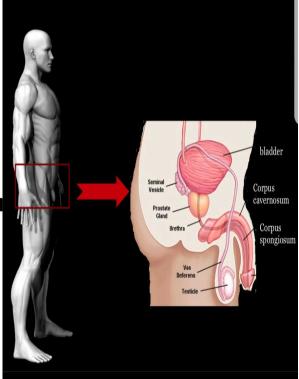
DR DEEPAK ABROL

CLINICAL ONCOLOGIST

J AND K HEALTH SERVICES

CONSULTANT ONCOLOGIST

MAHARISHI DAYANAND HOSPITAL AND MEDICAL RESEARCH CENTER AND 72 BPM HEALTH CARE PVT LTD



Introduction

- What is staging?
- Why staging is important
- Staging systems
 - TNM

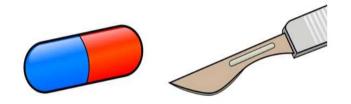
STAGING OF TUMORS

STAGE OF A TUMOR IS BASED ON

- SIZE OF PRIMARY TUMOR
- EXTENT OF INVASION INTO SURROUNDING TISSUES
- THE SPREAD TO REGIONAL LYMPH NODES
- PRESENCE OR ABSENCE OF BLOOD BORNE METASTASIS

Why is staging important?

Selecting treatment



Research

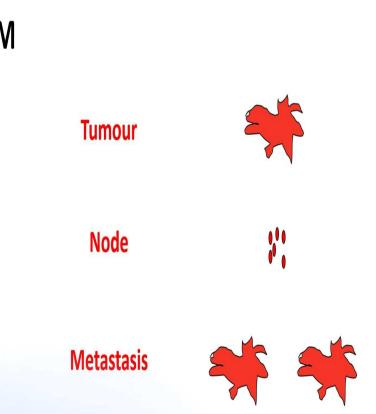


Guidelines



STAGING SYSTEM

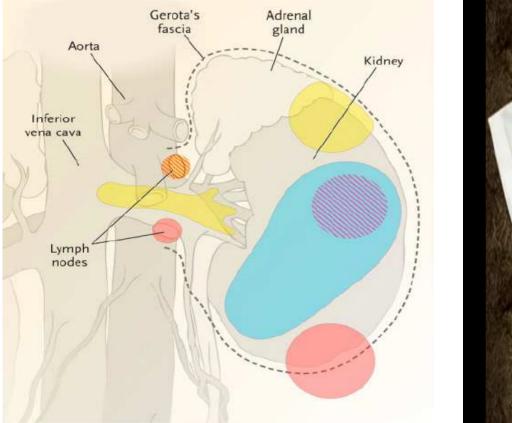
AJCC/UICC STAGING **SYSTEM** TNM **TNM CLASSIFICATION** WHERE **T** IS TUMOR **N** NODE AND **M** METASTASIS

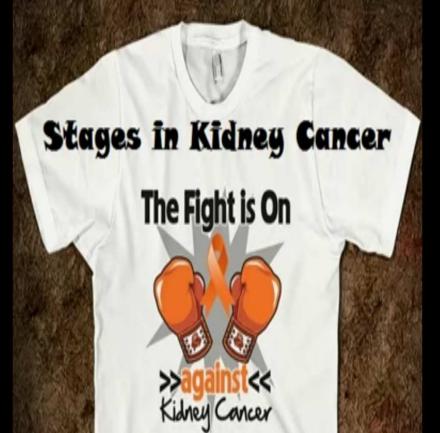


PLAN OF TODAY'S LECTURE

- Renal Cell Carcinoma
- Bladder Carcinoma
- Testicular Carcinoma
- Prostate Carcinoma
- Renal Pelvis and Ureter
- Urethra
- Penile Carcinoma

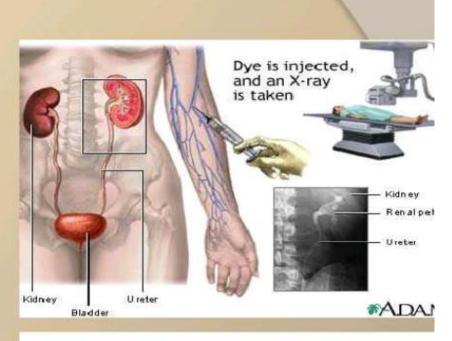
RENAL CANCER

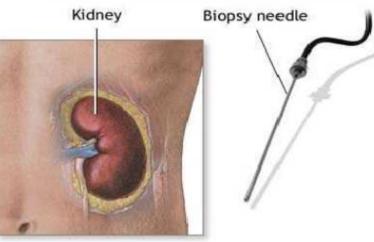




DIAGNOSIS

- Physical examination:
- Fever
- High blood pressure
- Lab tests:
- Complete blood count
- Urinanalysis
- Serum calcium
- Imaging tests:
- Ultrasound abdomen
- Abdominal CT scan
- MRI scan
- PET scan
- Renal angiography
- Intravenous pyelogram
- Chest x ray
- Bone scan
- Biopsy
 - Fine needle aspiration
 - Core needle biopsy





ADAN

American Joint Committee on Cancer (AJCC) TNM Staging System for Kidney Cancer (7th ed., 2010)

Primary Tumor (T)

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- T1 Tumor 7 cm or less in greatest dimension, limited to the kidney
- T1a Tumor 4 cm or less in greatest dimension, limited to the kidney
- T1b Tumor more than 4 cm but not more than 7 cm in greatest dimension, limited to the kidney
- T2 Tumor more than 7 cm in greatest dimension, limited to the kidney
- T2a Tumor more than 7 cm but less than or equal to 10 cm in greatest dimension, limited to the kidney
- T2b Tumor more than 10 cm, limited to the kidney
- T3 Tumor extends into major veins or perinephric tissues but not into the ipsilateral adrenal gland and not beyond Gerota's fascia
- T3a Tumor grossly extends into the renal vein or its segmental (muscle containing) branches, or tumor invades perirenal and/or renal sinus fat but not beyond Gerota's fascia
- T3b Tumor grossly extends into the vena cava below the diaphragm
- T3c Tumor grossly extends into the vena cava above the diaphragm or invades the wall of the vena cava
- T4 Tumor invades beyond Gerota's fascia (including contiguous extension into the ipsilateral adrenal gland)

Regional Lymph Nodes (N)

- NX Regional lymph nodes cannot be assessed
- N0 No regional lymph node metastasis
- N1 Metastasis in regional lymph node(s)

Distant Metastasis (M)

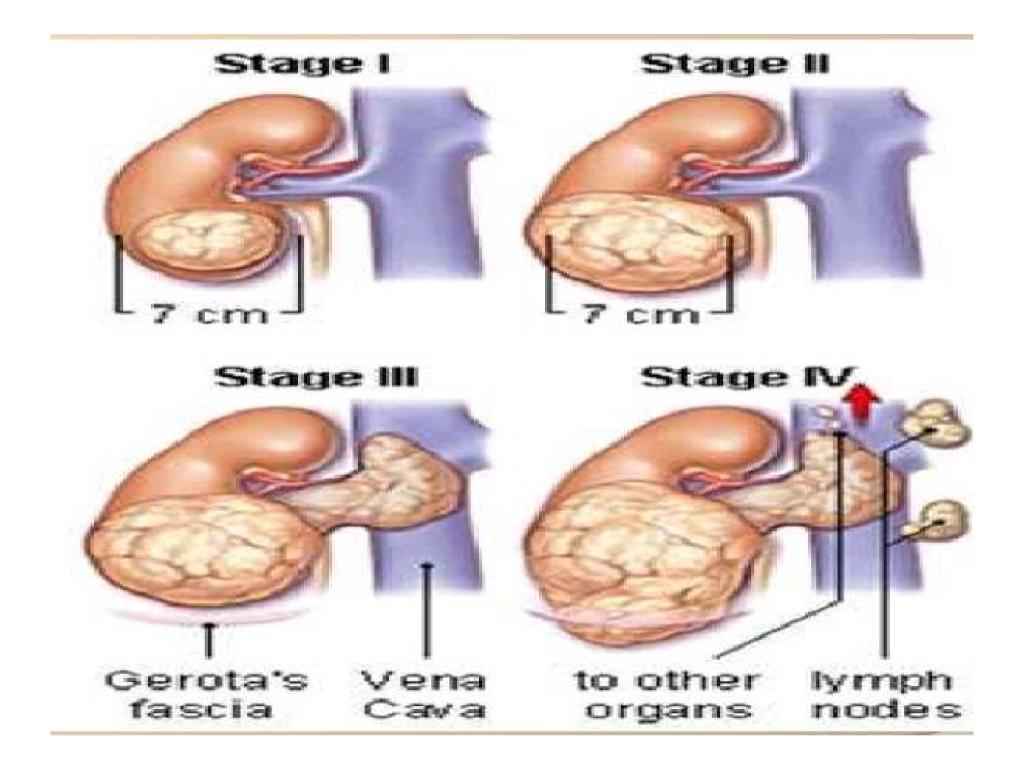
- M0 No distant metastasis
- M1 Distant metastasis

MODIFICATIONS IN AJCC 8TH EDITION

Kidney

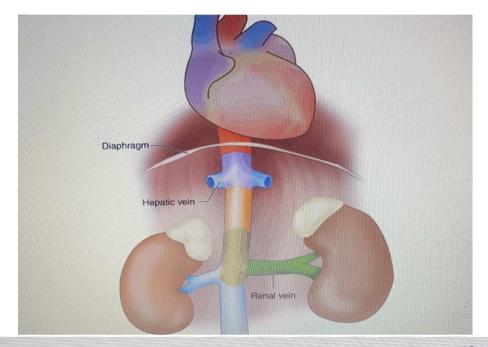
N2 is eliminated

| Anatomic Stage/Prognostic Groups | | | |
|----------------------------------|----------|----------|----|
| Stage I | T1 | N0 | M0 |
| Stage II | T2 | N0 | M0 |
| Stage III | T1 or T2 | N1 | M0 |
| | Т3 | N0 or N1 | M0 |
| Stage IV | T4 | Any N | M0 |
| | Any T | Any N | M1 |



Robson staging

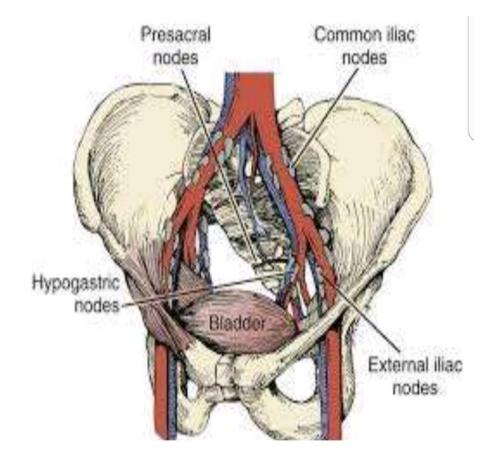
- Robson staging revolves around the relationship to Gerota's fascia, involvement of renal vein and regional nodes.
- **Stage I:** limited to kidney
- **Stage II:** involvement of perinephric fat but remains limited to Gerota's fascia
- Stage III
 - Illa: renal vein involvement
 - IIIb: nodal involvement
 - IIIc: both IIIa and IIIb
- Stage IV
 - IVa: direct invasion of adjacent organs / structures
 - IVb: distant metastases

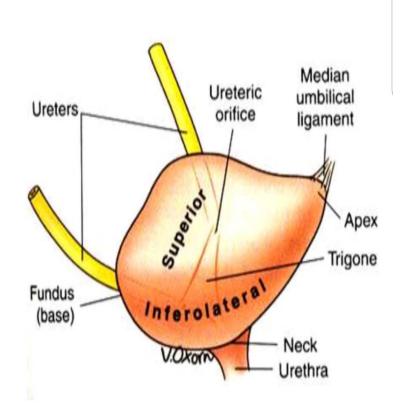


Prognostic and surgical staging systems of IVC tumor thrombus

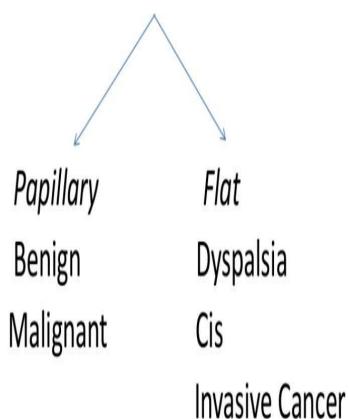
| Anatomic landmark | Staging systems | | | | |
|---|-----------------|-------|--------|--------|--------|
| | TNM | Neves | Novick | Hinman | Robson |
| RV | ТЗЬ | 0 | 1 | 1 | lla |
| IVC <2 cm above RV | | l I | 11 | | |
| IVC >2 cm above RV and below hepatic veins | | 1 | | | |
| IVC above hepatic veins and below diaphragm | | 111 | Ш | 11 | |
| IVC above diaphragm | T3c | IV | IV . | 111 | |

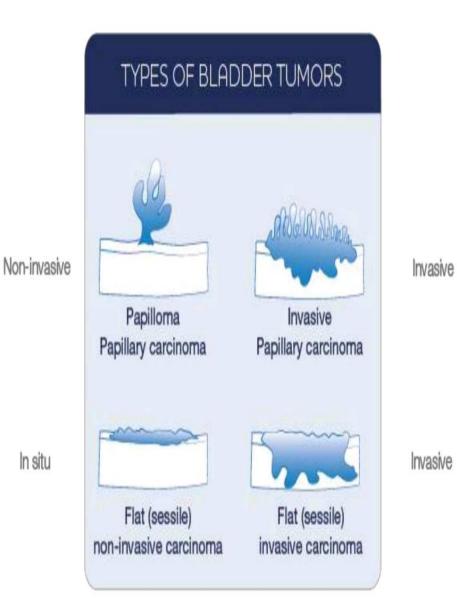
BLADDER CANCER

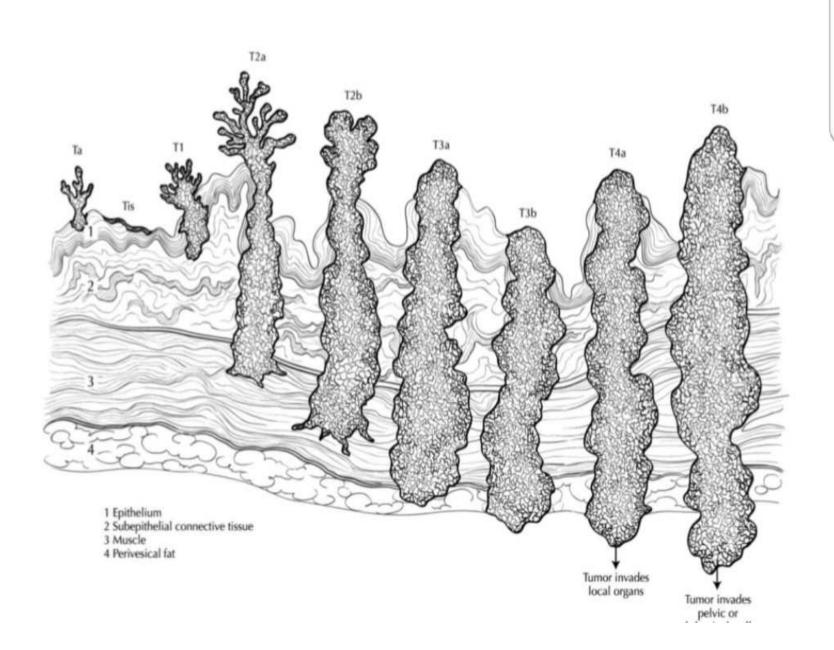




 Most Common Type is Transitional Cell Carcinoma 93%







American Joint Committee on Cancer (AJCC) TNM Staging System for Bladder Cancer (7th ed., 2010)

Primary Tumor (T)

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- Ta Noninvasive papillary carcinoma
- Tis Carcinoma in situ: "flat tumor"
- T1 Tumor invades subepithelial connective tissue
- T2 Tumor invades muscularis propria
- pT2a Tumor invades superficial muscularis propria (inner half)
- pT2b Tumor invades deep muscularis propria (outer half)
- T3 Tumor invades perivesical tissue
- pT3a Microscopically
- pT3b Macroscopically (extravesical mass)
- T4 Tumor invades any of the following: prostatic stroma, seminal vesicles, uterus, vagina, pelvic wall, abdominal wall
- T4a Tumor invades prostatic stroma, uterus, vagina
- T4b Tumor invades pelvic wall, abdominal wall

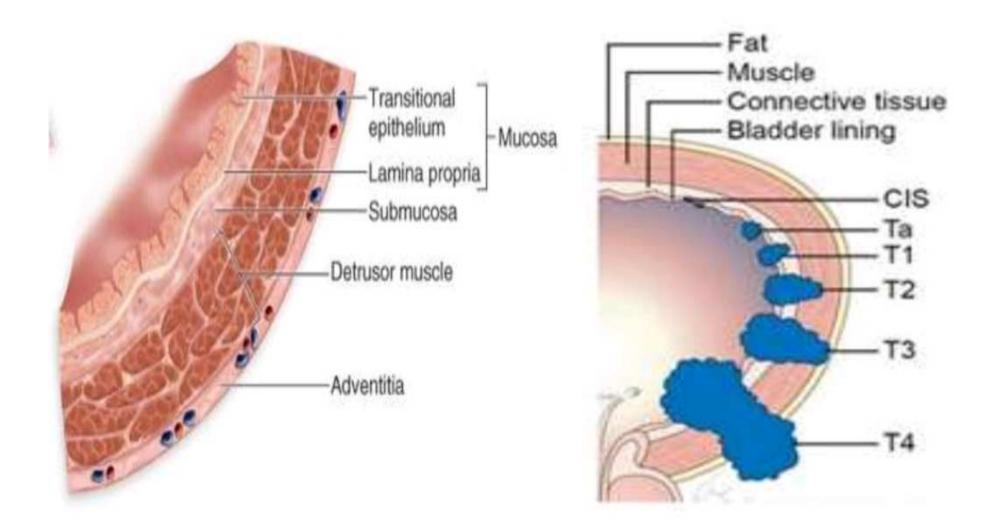
Regional Lymph Nodes (N)

Regional lymph nodes include both primary and secondary drainage regions. All other nodes above the aortic bifurcation are considered distant lymph nodes.

- NX Lymph nodes cannot be assessed
- N0 No lymph node metastasis
- N1 Single regional lymph node metastasis in the true pelvis (hypogastric, obturator, external iliac, or presacral lymph node)
- N2 Multiple regional lymph node metastasis in the true pelvis (hypogastric, obturator, external iliac, or presacral lymph node metastasis)
- N3 Lymph node metastasis to the common iliac lymph nodes

Distant Metastasis (M)

- M0 No distant metastasis
- M1 Distant metastasis



HISTOLOGIC GRADE

- LG low grade
- HG high grade (WHO/ISUP) IF NOT SPECIFIED
- **GX** can not be assessed
- G1 well differentiated
- G2 moderately differentiated
- G3 poorly differentiated
- G4 undifferentiated

| ANATOMIC ST Stage 0a | Tage/PF | N0 | M0 |
|-------------------------|---------|-------|----|
| Stage 0is | Tis | N0 | MO |
| Stage I | T1 | N0 | MO |
| Stage II | T2a | N0 | M0 |
| | T2b | N0 | M0 |
| Stage III | T3a | N0 | M0 |
| | T3b | N0 | M0 |
| | T4a | N0 | M0 |
| Stage IV | T4b | N0 | M0 |
| | Any T | N1-3 | M0 |
| | Any T | Any N | M1 |

MODIFICATIONS IN AJCC 8TH EDITION

Urinary bladder

The M category is subdivided M1a Non regional lymph nodes M1b Other distant metastasis Minor changes to **Stage**

Jewett-Marshall Staging(Clinical)

Stage A: Submucosal invasion but no muscle invasion

Stage B Bladder wall or muscle invasion

B1 Superficial

B2 Deep

Stage C Extension through serosa into perivesical fat

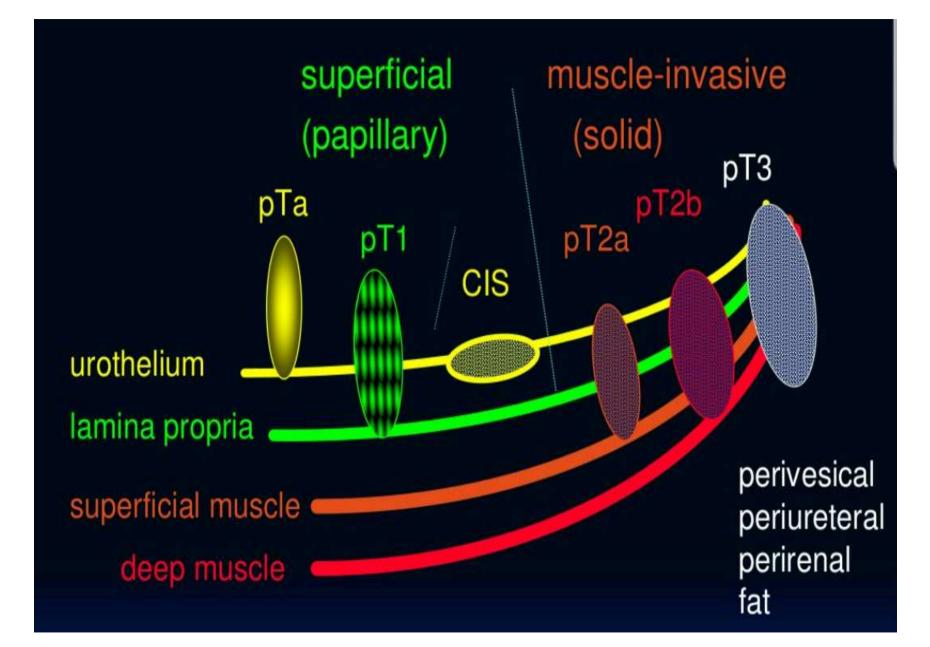
Stage D Lymph nodes and other distant metastasis

D1 regional nodes

D2 distant nodes and other distant mets

AJCC T1 TO T4 = JEWETT A TO D

N AND M PART OF **STAGE D**

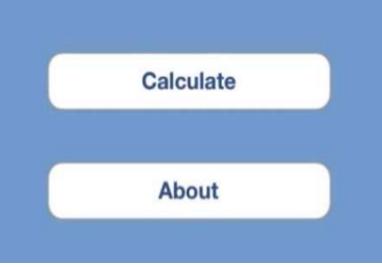


APPROXIMATE PROBABILITY OF RECURRENCE AND PROGRESSION

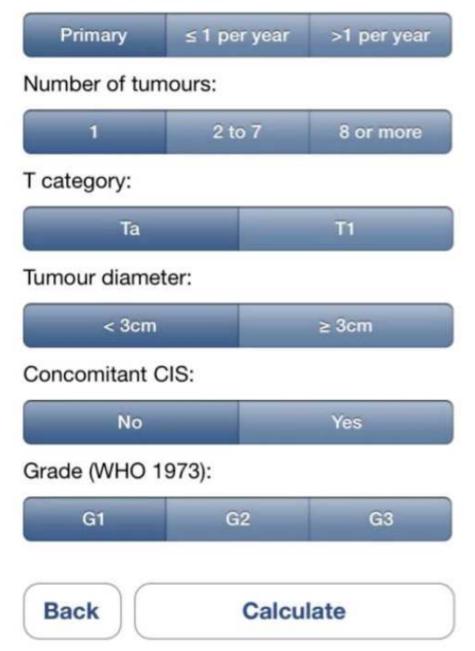
| <u>Pathology</u> | Approximate Probability of Recurrence in 5 years | Approximate Probability of Progression to Muscle Invasion |
|----------------------|---|--|
| Ta, low grade | 50% | Minimal |
| Ta, high grade | 60% | Moderate |
| T1, low grade (rare) | 50% | Moderate |
| T1, high grade | 50%-70% | Moderate-High |
| Tis | 50%-90% | High |



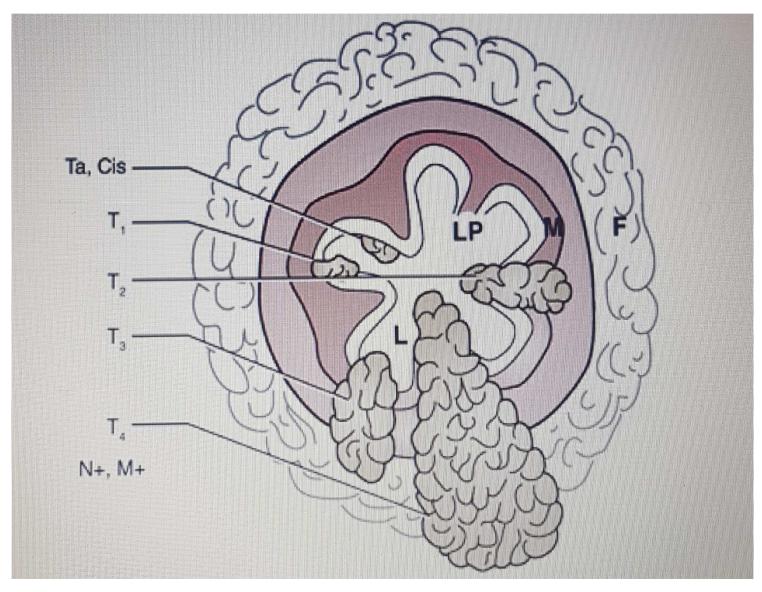
Bladder Cancer Prognosis Calculator



Prior recurrence rate:



RENAL PELVIS AND URETER



American Joint Committee on Cancer (AJCC)

TNM Staging System for Renal Pelvis and Ureter Cancer (7th ed., 2010)

Primary Tumor (T)

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- Ta Papillary noninvasive carcinoma
- Tis Carcinoma in situ
- T1 Tumor invades subepithelial connective tissue
- T2 Tumor invades the muscularis
- T3 (For renal pelvis only) Tumor invades beyond muscularis into peripelvic fat or the renal parenchyma T3. (For ureter only) Tumor invades beyond muscularis into periureteric fat
- T4 Tumor invades adjacent organs, or through the kidney into the perinephric fat.

Regional Lymph Nodes (N)*

- **NX** Regional lymph nodes cannot be assessed
- **N0** No regional lymph node metastasis
- N1 Metastasis in a single lymph node, 2 cm or less in greatest dimension
- N2 Metastasis in a single lymph node, more than 2 cm but not more than 5 cm in greatest dimension; or multiple lymph nodes, none more than 5 cm in greatest dimension
- N3 Metastasis in a lymph node, more than 5 cm in greatest dimension

* Note: Laterality does not affect the N classification.

Distant Metastasis (M)

- M0 No distant metastasis
- M1 Distant metastasis

| ANATOMIC ST Stage 0a | Tage/PF | NO NO | STIC GROUPS M0 |
|-------------------------|--|----------|----------------------|
| Stage 0is | Tis | N0 | MO |
| Stage I | T1 | N0 | MO |
| Stage II | T2 | N0 | MO |
| Stage III | Т3 | N0 | MO |
| Stage IV | T4 Any T Any T Any T Any T | N2 N3 | M0 M0 M0 M1 |

| 5 YEAR DISEASE UPPER URINARY | FREE SURVIVAL IN TRACT BY STAGE | TUMORS OF |
|---------------------------------|------------------------------------|-----------|
| STAGE | PERCENTAGE SURVIVAL | |
| рТ1 | 92 | |
| pT2 | 73 | |
| рТЗ | 41 | |
| pT4 | 0 | |

American Joint Committee on Cancer (AJCC) TNM Staging System for Urethral Carcinoma (7th ed., 2010)

Primary Tumor (T) (Male and Female)

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- Ta Noninvasive papillary, polypoid, or verrucous carcinoma
- Tis Carcinoma in situ
- T1 Tumor invades subepithelial connective tissue
- T2 Tumor invades any of the following: corpus spongiosum, prostate, periurethral muscle
- T3 Tumor invades any of the following: corpus cavernosum, beyond prostatic capsule, anterior vagina, bladder neck
- T4 Tumor invades other adjacent organs

Regional Lymph Nodes (N)

- NX Regional lymph nodes cannot be assessed
- N0 No regional lymph node metastasis
- N1 Metastasis in a single lymph node 2 cm or less in greatest dimension
- N2 Metastasis in a single node more than 2 cm in greatest dimension, or in multiple nodes

Distant Metastasis (M)

- M0 No distant metastasis
- M1 Distant metastasis

Urothelial (Transitional Cell) Carcinoma of the Prostate

- Tis pu Carcinoma in situ, involvement of the prostatic urethra
- Tis pd Carcinoma in situ, involvement of the prostatic ducts
- T1 Tumor invades urethral subepithelial connective tissue
- T2 Tumor invades any of the following: prostatic stroma, corpus spongiosum, periurethral muscle
- T3 Tumor invades any of the following: corpus cavernosum, beyond prostatic capsule, bladder neck (extraprostatic extension)
- T4 Tumor invades other adjacent organs (invasion of the bladder)

| ANATOMIC ST Stage 0a | TAGE/PF | N0 | M0 |
|-------------------------|---------|-------|----|
| Stage 0is | Tis | N0 | M0 |
| | Tis pu | N0 | M0 |
| | Tis pd | N0 | M0 |
| Stage I | T1 | N0 | MO |
| Stage II | T2 | N0 | MO |
| Stage III | T1 | N1 | M0 |
| | T2 | N1 | M0 |
| | T3 | N0 | M0 |
| | T3 | N1 | M0 |
| Stage IV | T4 | N0 | M0 |
| | T4 | N1 | M0 |
| | Any T | N2 | M0 |
| | Any T | Any N | M1 |

MODIFICATIONS IN AJCC 8TH EDITION

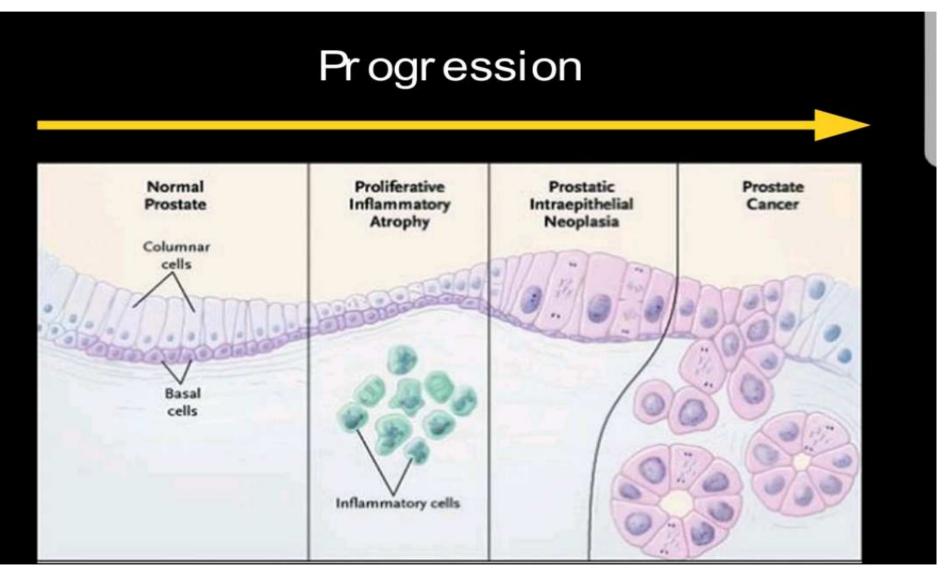
Urethra

Changes to N category

N1 Metastasis in a single lymph node

N2 Metastasis in multiple lymph nodes

PROSTATE CANCER



Staging work up

• Baseline work up

Haemogram/LFT/KFT/CXR

• Essential work up

PSA/Biopsy/Gleason scoring

• Complimentary work up

CT/MRI(becomes essential in higher clinical stage) PET CT Bone Scan(essential if bone mets suspected)

Molecular testing

TNM Staging System For Prostate Cancer Primary Tumor (T)

Clinical

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- T1 Clinically inapparent tumor neither palpable nor visible by imaging
 - T1a Tumor incidental histologic finding in 5% or less of tissue resected
 - T1b Tumor incidental histologic finding in more than 5% of tissue resected
 - T1c Tumor identified by needle biopsy (e.g., because of elevated PSA)
- T2 Tumor confined within prostate*
 - T2a Tumor involves one-half of one lobe or less
 - T2b Tumor involves more than one-half of one lobe but not both lobes
 - T2c Tumor involves both lobes
- T3 Tumor extends through the prostatic capsule**
 - T3a Extracapsular extension (unilateral or bilateral)
 - T3b Tumor invades the seminal vesicle(s)
- T4 Tumor is fixed or invades adjacent structures other than seminal vesicles: bladder, levator muscles, and/or pelvic wall.
- *Note: Tumor found in one or both lobes by needle biopsy, but not palpable or reliably visible by imaging, is classified as T1c.
- **Note: Invasion into the prostatic apex or into (but not beyond) the prostatic capsule is not classified as T3, but as T2.

Pathologic(pT)*

| pT2 Organ co | nfined |
|--------------|--------|
|--------------|--------|

- pT2a Unilateral, involving one-half of one side or less
- pT2b Unilateral, involving more than one-half of one side but not both sides
- pT2c Bilateral disease
- pT3 Extraprostatic extension
 - pT3a Extraprostatic extension or microscopic invasion of the bladder neck**
 - pT3b Seminal vesicle invasion

pT4 Invasion of bladder, rectum

*Note: There is no pathologic T1 classification.

**Note: Positive surgical margin should be indicated by an R1 descriptor (residual microscopic disease).

Regional Lymph Nodes (N)

Clinical

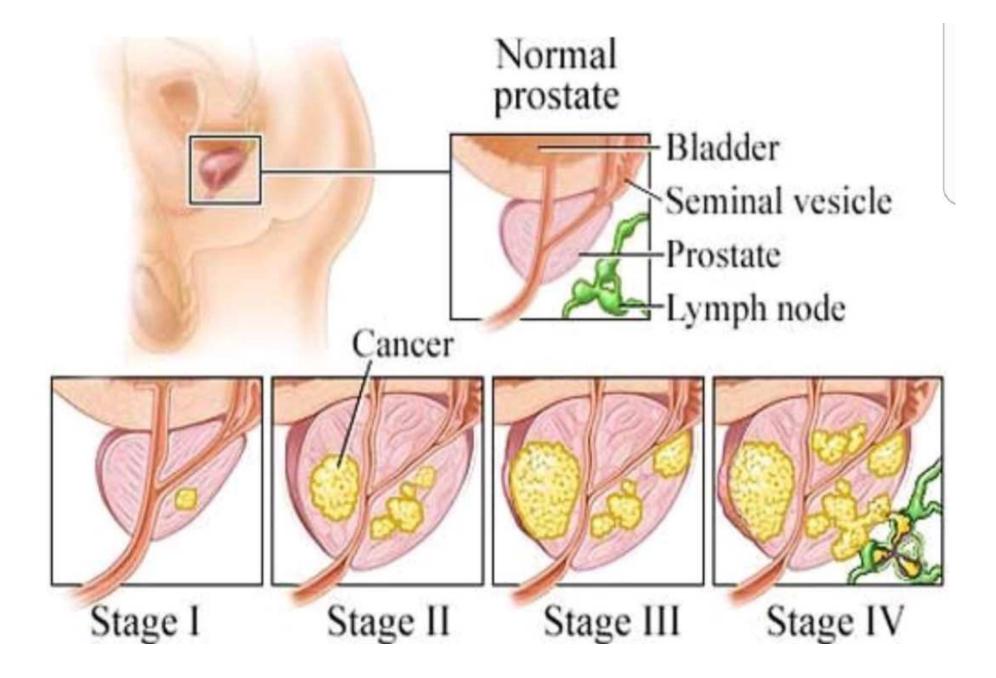
- NX Regional lymph nodes were not assessed
- N0 No regional lymph node metastasis
- N1 Metastasis in regional lymph node(s)

Pathologic

- PNX Regional nodes not sampled
- pN0 No positive regional nodes
- pN1 Metastases in regional nodes(s)

Distant Metastasis (M)*

- M0 No distant metastasis
- M1 Distant metastasis
 - M1a Non-regional lymph node(s)
 - M1b Bone(s)
 - M1c Other site(s) with or without bone disease
- *Note: When more than one site of metastasis is present, the most advanced category is used. pMIc is most advanced.



ANATOMIC STAGE/PROGNOSTIC GROUPS *

| Group | Т | N | M | PSA | Gleason |
|-------|-------|-------|----|-------------|-------------|
| 1 | T1a-c | N0 | MO | PSA <10 | Gleason ≤6 |
| | T2a | N0 | MO | PSA <10 | Gleason ≤6 |
| | T1-2a | N0 | MO | PSA X | Gleason X |
| IIA | T1a-c | N0 | MO | PSA <20 | Gleason 7 |
| | T1a-c | N0 | MO | PSA ≥10 <20 | Gleason ≤6 |
| | T2a | N0 | MO | PSA <20 | Gleason ≤7 |
| | T2b | N0 | MO | PSA <20 | Gleason ≤7 |
| | T2b | N0 | MO | PSA X | Gleason X |
| IIB | T2c | N0 | MO | Any PSA | Any Gleason |
| | T1-2 | N0 | MO | PSA ≥20 | Any Gleason |
| | T1-2 | N0 | MO | Any PSA | Gleason ≥8 |
| 111 | T3a-b | N0 | MO | Any PSA | Any Gleason |
| IV | T4 | N0 | MO | Any PSA | Any Gleason |
| | Any T | N1 | MO | Any PSA | Any Gleason |
| | Any T | Any N | M1 | Any PSA | Any Gleason |

*Note: When either PSA or Gleason is not available, grouping should be determined by T stage and/or either PSA or Gleason as available.

MODIFICATIONS IN AJCC 8TH EDITION

Prostate

T4N0M0 is now stage III

WHO Grade should be used to record tumour grade

Stage

| Stage | I | T1, T2a | NO | M0 |
|-------|----|----------------|-------|----|
| Stage | II | T2b-2c | N0 M0 | |
| Stage | | T3 <i>,</i> T4 | N0 M0 | |
| Stage | IV | Any T | N1 M0 | |
| | | Any T | Any N | M1 |

Whitmore-Jewett staging

- A: Tumor is present, but not detectable clinically; found incidentally
 - A1: tissue resembles normal cells; found in a few chips from one lobe
 - A2: more extensive involvement
- B: Tumor can be felt on physical examination but has not spread outside the prostatic capsule
 - BIN: the tumor can be felt, it does not occupy a whole lobe, and is surrounded by normal tissue
 - B1: the tumor can be felt and it does not occupy a whole lobe
 - B2: the tumor can be felt and it occupies a whole lobe or both lobes
- C: Tumor has extended through the capsule
 - C1: the tumor has extended through the capsule but does not involve the seminal vesicles
 - C2: the tumor involves the seminal vesicles
- D: Tumor has spread to other organs

Gleason Grade

• Histological grading of prostate cancer 1-5

However

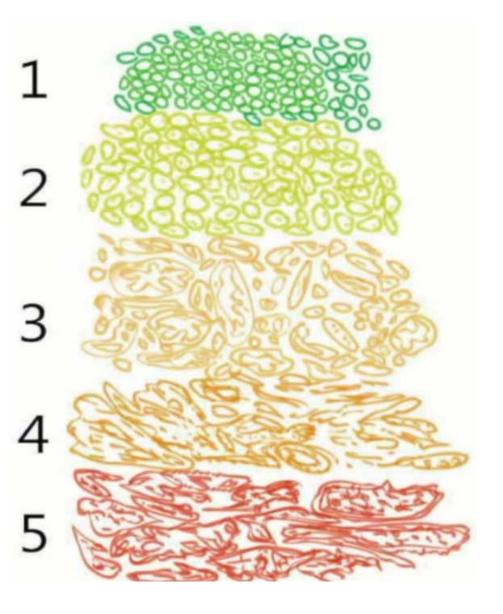
- Prostate cancer not uniform
- To aid calculations of prognosis, the sum of the *2* most prevalent islands of prostate cancer are used
- Therfore, gleason grade ranges 2-10

Gleason XGleason score cannot be processedGleason ≤6Well differentiated (slight anaplasia)Gleason 7Moderately differentiated (moderate anaplasia)Gleason 8-10Poorly differentiated/undifferentiated
(marked anaplasia)

GLEASON PATTERN

- 1. SMALL UNIFORM GLANDS
- 2. MORE STROMA BETWEEN GLANDS
- 3. DISTINCTLY INFILTERATIVE MARGINS

- 4. IRREGULAR MASSES OF NEOPLASTIC GLANDS
- 5. ONLY OCCASIONAL GLAND FORMATION



GLEASON GRADE GROUP DEFINITIONS

Gleason grade group 1: Gleason score ≤6 Only individual discrete well-formed glands

Gleason grade group 2: Gleason score 3+4=7 Predominantly well-formed glands with lesser component of poorly-formed/fused/cribriform glands

Gleason grade group 3: Gleason score 4+3=7

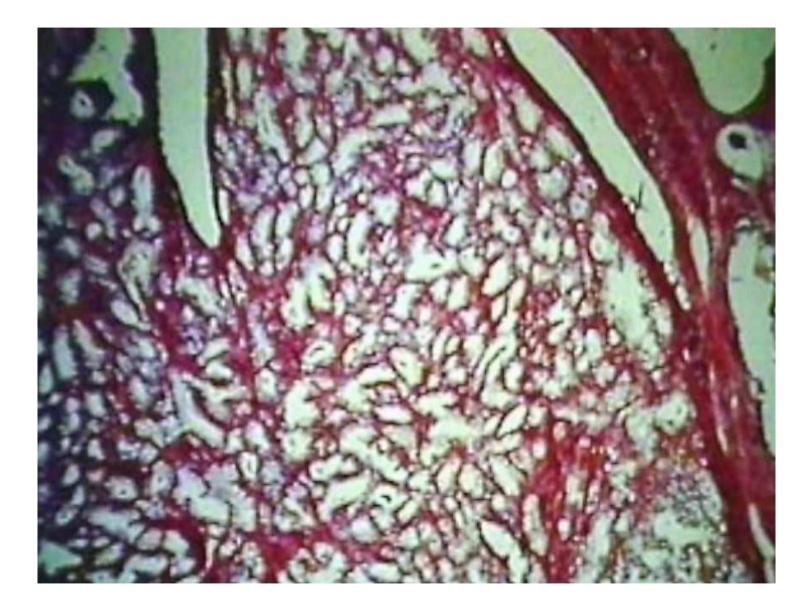
Predominantly poorly-formed/fused/cribriform glands with lesser component of well-formed glands*

Gleason grade group 4: Gleason score 4+4=8; 3+5=8; 5+3=8

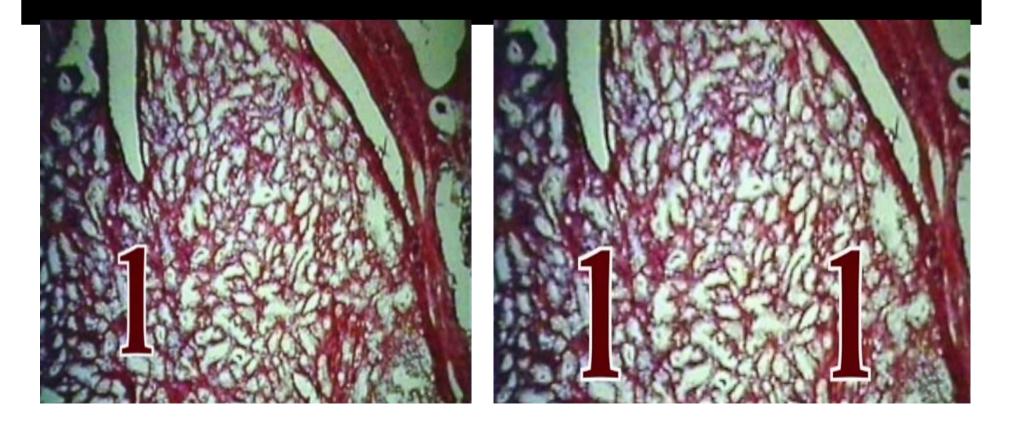
- Only poorly-formed/fused/cribriform glands or
- Predominantly well-formed glands and lesser component lacking glands¹ or
- Predominantly lacking glands and lesser component of well-formed glands¹

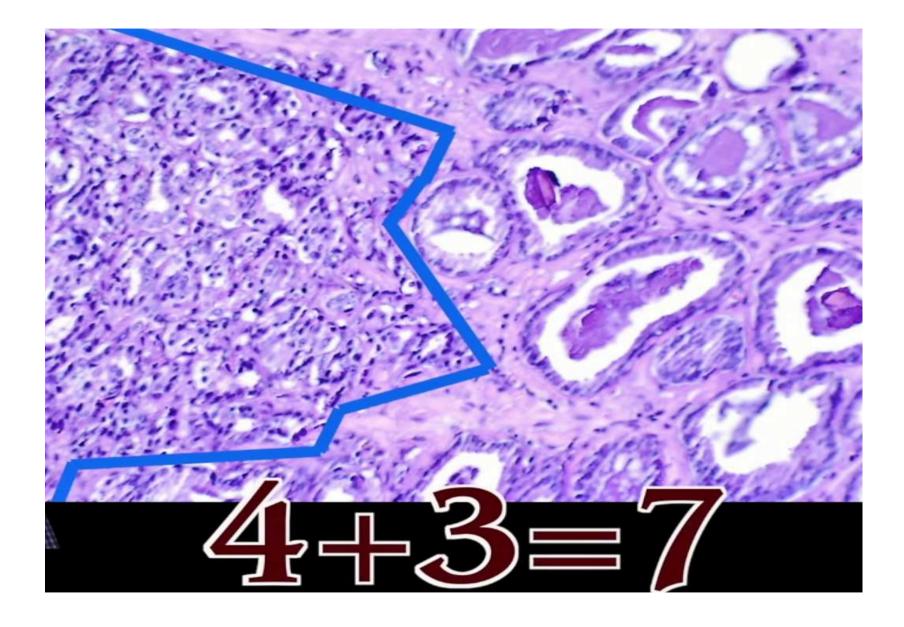
Gleason grade group 5: Gleason score 9-10

Lack gland formation (or with necrosis) with or without poorly formed/fused/cribriform glands²



Gleason score = Gleason grade + Gleason grade





RISK STRATIFICATION

D'amico Risk Stratification

- First one to give concept
- Predicts about chances of recurrence after treatment
- Low risk- T1-T2a and GS ≤6 and PSA ≤10
- Intermediate risk- T2b and/or GS =7 and/or PSA >10-20
- High risk- ≥T2c or PSA >20 or GS 8–10
- Does not take into account other parameters

| PSA Level Q <10 ng/mL +1 | Intermediate |
|-----------------------------|--|
| • 10-20 ng/mL -2 | Risk of Recurrence 25-50% failure at 5 years post treatment. |
| Q >20 ng/mL +3 | |
| Gleason Score | |
| Q<6+1 | |
| 07-2 | |
| 0 28 -3 | |
| Clinical Stage | |
| • 11-12a - 1 | |
| O T2b +2 | |
| Q 2T2c +3 | |

| PARTIN TABLE | S | | |
|----------------------------|---------------------------------------|--|------------------------------------|
| PSA: >10 • | | | |
| Gleason Score: 4+3 • | | | |
| Clinical Stage: T2a | ۲ | | |
| Find Results | | | |
| | | | |
| OC: organ confined (22) | EPE: extraprostatic extension (22) | SV+: seminal vesicle involvement (10) | LN+: lymph node involvement (2) |
| 25(20-31) | 44(36-53) | 20(12-29) | 10(4-18) |
| Numbers represent n | errentage of natients with the snu | ecified PSA, clinical stage, and biopsy | Gleason score who would have |
| 10 (29) | | (EPE), cancer invading into the semin | |

 Uses Gleason score, serum PSA, and clinical stage – to predict whether the tumor will be confined to the prostate

The UCSF-CAPRA score

- UCSF developed the Cancer of the Prostate Risk Assessment score
- 0-2 low risk
 3-5 intermediate risk
 6-10 high risk

| Variable | Specific patient's level | Points to be assigned | | |
|--|---|-----------------------|--|--|
| Age at diagnosis | Under 50 | 00 | | |
| | 50 or older | 1.0 | | |
| | less than or equal to 6 | 00 | | |
| | between 6.1 and 10 | 10 | | |
| PSA at diagnosis (ng/ml) | between 10.1 and 20 | 2 () | | |
| nason score of the biopsy imary/secondary) nical stage (T-stage) roent of biopsy cores involved with cancer | between 20.1 and 30 | 3.8 | | |
| | more than 30 | 40 | | |
| | no pattern 4 or 5 | 0.0 | | |
| ∃leason score of the biopsy primary/secondary) | secondary pattern 4 or 5 | 1.00 | | |
| | primary pattern 4 or 5 | 3.0 | | |
| Clinical stage (T-stage) | T1 or T2 | 0.8 | | |
| | ТЗа | 10 | | |
| Percent of biopsy cores involved with cancer | less than 34 percent | 0.0 | | |
| (positive for cancer) | Under 50 9 0 50 or older 1 # less than or equal to 6 0 0 between 6.1 and 10 1 0 between 10.1 and 20 2 0 between 20.1 and 30 3 # more than 30 4 0 no pattern 4 or 5 0 0 secondary pattern 4 or 5 1 # primary pattern 4 or 5 3 0 T1 or T2 0 # T3a 1 0 set than 34 percent 0 # 34 percent or more 1 0 Your CAPRA Score is 5 This indicates intermediate risk prostate cancer Among 3,000 men with CAPRA 3-5 who und prostatectomy at UCSF since 2000, the likeling • metastasis or dying from prostate cancer 5 years was 1% | 10 | | |
| | This indicates intermediate risk prostate cancer Among 3,000 men with CAPRA 3-5 who underwent radical | | | |
| Calculate Reset | metastasis or dying from pros | state cancer within | | |

NCCN Risk categories

5.1 Very low risk

Includes men with a T1c tumor, PSA level less than 10 ng/mL, PSA density less than 0.15 ng/mL/g, <u>Gleason</u> score 6 or less, and cancer in fewer than three <u>biopsy</u> cores and in half or less of any core.

5.2 Low risk

Includes men with a T1a, T1b, T1c, or T2a tumor, PSA level less than 10 ng/mL, and Gleason score 6 or less.

5.3 Intermediate risk

Includes men with a T2b or T2c tumor, PSA level between 10 and 20 ng/mL, or Gleason score 7. If you meet two or all three conditions, your risk is high.

5.4 High risk

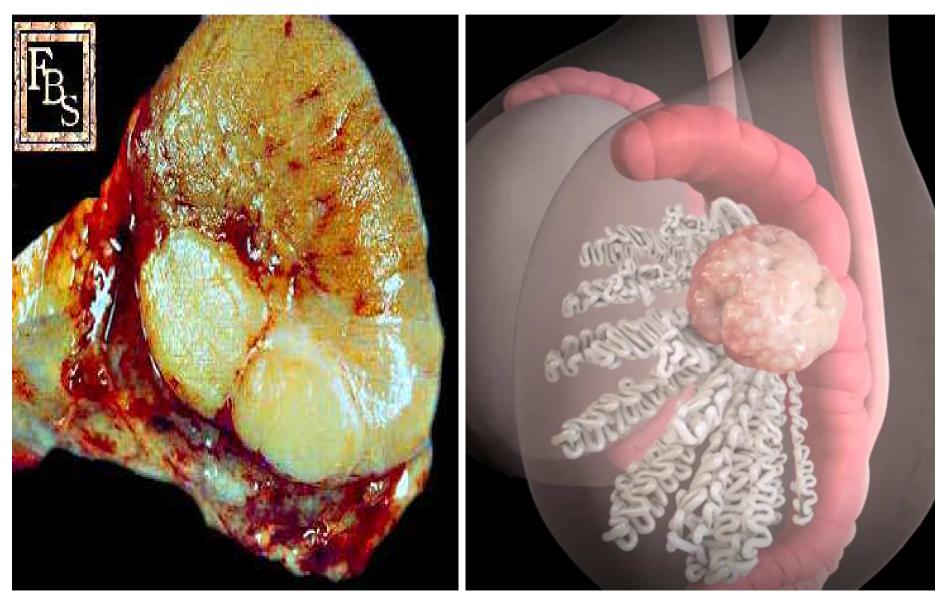
Includes men with a T3a tumor, a PSA level greater than 20 ng/mL, or a Gleason score between 8 and 10. If you meet two or all three conditions, your risk is very high.

5.5 Very high risk

Includes men with a T3b or T4 tumor, primary <u>Gleason</u> grade 5, or more than 4 biopsy cores with Gleason scores between 8 and 10.

5.6 Metastatic disease Includes men with N1 or M1 disease.

TESTICULAR CANCER



American Joint Committee on Cancer (AJCC) TNM Staging System for Testis Cancer (7th ed., 2010)

Primary Tumor (T)*

The extent of primary tumor is usually classified after radical orchiectomy, and for this reason, a pathologic stage is assigned.

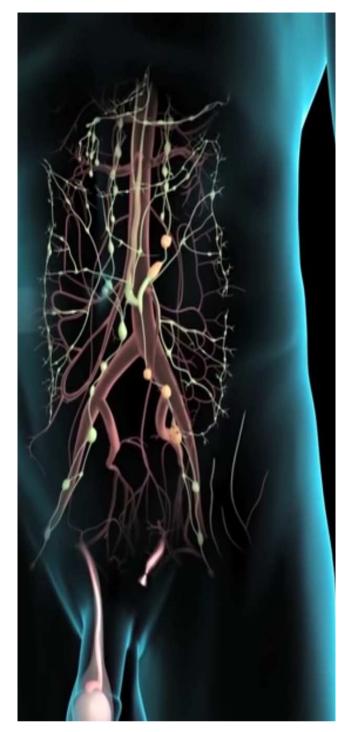
- pTX Primary tumor cannot be assessed
- pT0 No evidence of primary tumor (e.g. histologic scar in testis)
- pTis Intratubular germ cell neoplasia (carcinoma in situ)
- pT1 Tumor limited to the testis and epididymis without vascular/ lymphatic invasion; tumor may invade into the tunica albuginea but not the tunica vaginalis
- pT2 Tumor limited to the testis and epididymis with vascular/ lymphatic invasion, or tumor extending through the tunica albuginea with involvement of the tunica vaginalis
- pT3 Tumor invades the spermatic cord with or without vascular/ lymphatic invasion
- pT4 Tumor invades the scrotum with or without vascular/lymphatic invasion

Regional Lymph Nodes (N) Clinical

- NX Regional lymph nodes cannot be assessed
- N0 No regional lymph node metastasis
- N1 Metastasis with a lymph node mass 2 cm or less in greatest dimension; or multiple lymph nodes, none more than 2 cm in greatest dimension
- N2 Metastasis with a lymph node mass, more than 2 cm but not more than 5 cm in greatest dimension; or multiple lymph nodes, any one mass greater than 2 cm but not more than 5 cm in greatest dimension
- N3 Metastasis with a lymph node mass more than 5 cm in greatest dimension

Pathologic (pN)

- pNX Regional lymph nodes cannot be assessed
- pN0 No regional lymph node metastasis
- pN1 Metastasis with a lymph node mass 2 cm or less in greatest dimension and less than or equal to five nodes positive, none more than 2 cm in greatest dimension
- pN2 Metastasis with a lymph node mass more than 2 cm but not more than 5 cm in greatest dimension; or more than five nodes positive, none more than 5 cm; or evidence of extranodal extension of tumor
- pN3 Metastasis with a lymph node mass more than 5 cm in greatest dimension
- ;), Distant Metastasis (M)
 - M0 No distant metastasis
 - M1 Distant metastasis
 - M1a Nonregional nodal or pulmonary metastasis
 - M1b Distant metastasis other than to nonregional lymph nodes and lung



Serum Tumor Markers (S)

- SX Marker studies not available or not performed
- SO Marker study levels within normal limits
- S1 LDH < 1.5 x N* and hCG (mlu/mL) < 5,000 and AFP (ng/ml) < 1,000
- S2 LDH 1.5-10 x N or hCG (mlu/mL) 5,000-50,000 or AFP (ng/ml) 1,000-10,000
- S3 LDH > 10 x N or hCG (mlu/mL) > 50,000 or AFP (ng/ml) > 10,000

*N indicates the upper limit of normal for the LDH assay.

American Joint Committee on Cancer (AJCC) TNM Staging System for Testis Cancer (7th ed., 2010)

ANATOMIC STAGE/PROGNOSTIC GROUPS

| Group | т | N | м | S (Serum Tumor Markers) |
|------------|-----------|-------|-----|-------------------------|
| Stage 0 | pTis | N0 | мо | S0 |
| Stage I | pT1-4 | N0 | MO | SX |
| Stage IA | pT1 | N0 | MO | S0 |
| Stage IB | pT2 | NO | MO | S0 |
| | PT3 | NO | MO | S0 |
| | PT4 | N0 | MO | S0 |
| Stage IS | Any pT/TX | N0 | MO | S1-3 |
| Stage II | Any pT/Tx | N1-3 | MO | SX |
| Stage IIA | Any pT/TX | N1 | MO | S0 |
| | Any pT/TX | N1 | MO | S1 |
| Stage IIB | Any pT/TX | N2 | MO | S0 |
| | Any pT/TX | N2 | MO | S1 |
| Stage IIC | Any pT/TX | N3 | MO | S0 |
| | Any pT/TX | N3 | MO | S1 |
| Stage III | Any pT/TX | Any N | M1 | SX |
| Stage IIIA | Any pT/TX | Any N | M1a | S0 |
| | Any pT/TX | Any N | M1a | S1 |
| Stage IIIB | Any pT/TX | N1-3 | MO | S2 |
| | Any pT/TX | Any N | M1a | S2 |
| Stage IIIC | Any pT/TX | N1-3 | MO | S3 |
| | Any pT/TX | Any N | M1a | S3 |
| | Any pT/Tx | Any N | M1b | Any S |

Staging and Classification

| Test | Recommendation | GR |
|--|---|----|
| Serum tumour markers | Alpha-fetoprotein hCG LDH | A |
| Abdominopelvic CT | All patients | A |
| Chest CT | All patients | Α |
| Testis ultrasound (bilateral) | All patients | Α |
| Bone scan or MRI columna | In case of symptoms | |
| Brain scan (CT/MRI) | In case of symptoms and patients with metastatic disease with multiple lung metastases and/or high beta-hCG values. | |
| Further investigations | | |
| Fertility investigations: Total testosterone LH FSH Semen analysis | | В |
| Sperm banking | Should be offered | A |

• The mean serum half-life of AFP 5-7 days and hCG is 2-3days

Staging and Classification

| Good-prognosis group | |
|-----------------------------|--|
| Non-seminoma (56% of cases) | All of the following criteria: |
| 5-year PFS 89% | Testis/retroperitoneal primary |
| 5-year survival 92% | No non-pulmonary visceral metastases |
| | AFP < 1,000 ng/mL |
| | hCG < 5,000 IU/L (1,000 ng/mL) |
| | • LDH < 1.5 x ULN |
| Seminoma (90% of cases) | All of the following criteria: |
| 5-year PFS 82% | Any primary site |
| 5-year survival 86% | No non-pulmonary visceral metastases |
| | Normal AFP |
| | Any hCG |
| | Any LDH |

Staging and Classification

| Intermediate prognosis group | |
|---|--|
| Non-seminoma (28% of cases) | Testis/retroperitoneal primary |
| 5-year PFS 75% | No non-pulmonary visceral metastases |
| 5-year survival 80% | AFP 1,000 - 10,000 ng/mL or |
| in lange daaleer 1 dae de adaaleer wat wat ook ak a | hCG 5,000 - 50,000 IU/L or |
| | • LDH 1.5 - 10 x ULN |
| Seminoma (10% of cases) | All of the following criteria: |
| 5-year PFS 67% | Any primary site |
| 5-year survival 72% | Non-pulmonary visceral metastases |
| | Normal AFP |
| | Any hCG |
| | Any LDH |
| Poor prognosis group | |
| Non-seminoma (16% of cases) | Any of the following criteria: |
| 5-year PFS 41% | Mediastinal primary |
| 5-year survival 48% | Non-pulmonary visceral metastases |
| | AFP > 10,000 ng/mL or |
| | hCG > 50,000 IU/L (10,000 ng/mL) or |
| | • LDH > 10 x ULN |
| Seminoma | No patients classified as poor prognosis |

| | Seminoma | | Non-seminoma | | |
|---------------|----------|--------------|--------------|--------------|------|
| | good | intermediate | Good | Intermediate | Poor |
| 5 yr PFS | 82% | 67% | 89% | 75% | 41% |
| 5 yr survival | 86% | 72% | 92% | 80% | 48% |

PENILE CANCER

Natural History

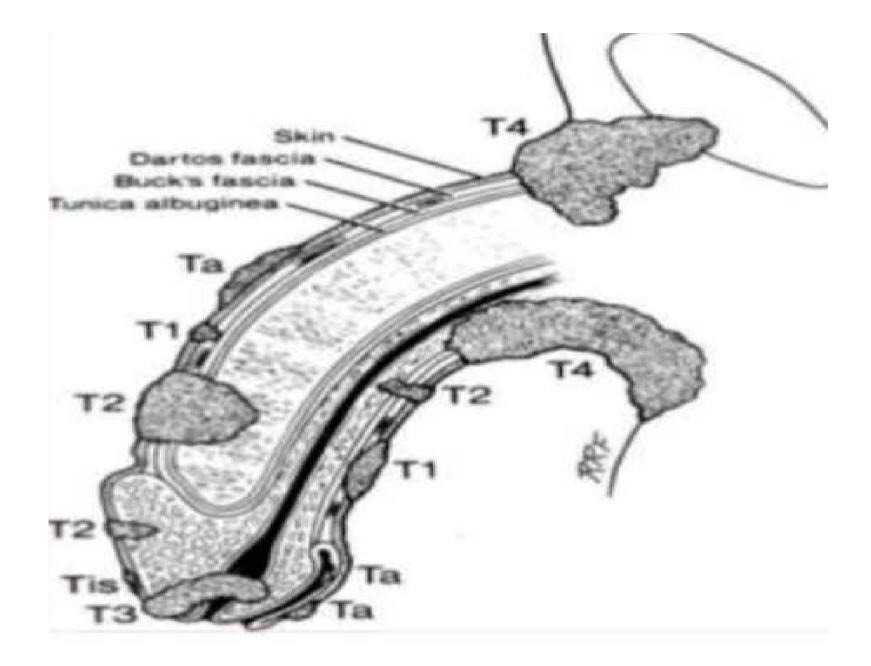
COMMON SITE OF PRIMARY TUMOR

- within the preputial area
- in the glans
- coronal sulcus
- prepuce
- Lesions arising in the skin of the shaft are rare
- slow loco regional progression

American Joint Committee on Cancer (AJCC) TNM Staging System for Penile Cancer (7th ed., 2010)

Primary Tumor (T)

- TX Primary tumor cannot be assessed
- T0 No evidence of primary tumor
- Ta Noninvasive verrucous carcinoma*
- Tis Carcinoma in situ
- T1a Tumor invades subepithelial connective tissue without lymph vascular invasion and is not poorly differentiated (i.e., grade 3-4)
- T1b Tumor invades subepithelial connective tissue with lymph vascular invasion or is poorly differentiated
- T2 Tumor invades corpus spongiosum or cavernosum
- T3 Tumor invades urethra
- T4 Tumor invades other adjacent structures
- *Note: Broad pushing penetration (invasion) is permitted; destructive invasion is against the diagnosis



Regional Lymph Nodes (N)

Clinical Stage Definition*

- cNX Regional lymph nodes cannot be assessed
- cN0 No lymph node metastasis
- cN1 Palpable mobile unilateral inguinal lymph node
- cN2 Palpable mobile multiple or bilateral inguinal lymph nodes
- cN3 Palpable fixed inguinal nodal mass or pelvic lymphadenopathy unilateral or bilateral

Pathologic Stage Definition*

- pNX Regional lymph nodes cannot be assessed
- pN0 No regional lymph node metastasis
- pN1 Metastasis in a single inguinal node
- pN2 Metastasis in multiple or bilateral inguinal lymph nodes
- pN3 Extranodal extension of lymph node metastasis or pelvic lymph node(s) unilateral or bilateral

*Note: Pathologic stage definition based on biopsy or surgical excision.

Distant Metastasis (M)

- M0 No distant metastasis
- M1 Distant metastasis

| ANATOMIC S Stage 0 | TAGE/PRO | N0 | M0 |
|-----------------------|----------|-------|----|
| Stage 0 | Ta | NO | MO |
| Stage I | T1a | N0 | MO |
| Stage II | T1b | N0 | MO |
| | T2 | NO | MO |
| | Т3 | NO | MO |
| Stage IIIA | T1-3 | N1 | MO |
| Stage IIIB | T1-3 | N2 | MO |
| Stage IV | T4 | Any N | MO |
| | Any T | N3 | MO |
| | Any T | Any N | M1 |

MODIFICATIONS IN AJCC 8TH EDITION

Penis

Changes in T category and pathological N category

THANKS

STUDENTS ARE THE REAL

TORCH

BEARERS

