Malignant renal tumors 2. Nephroblastoma (Wilm's tumor)

- Uncommon renal tumor but one of the commonest *childhood* neoplasms (2-5 yrs).
- Sporadic or familial associated with congenital malformations due to **WT-1 or WT-2** genes mutaions on **chr.11**:
 - **WAGR** syndrome.
 - *Denys-Drash syndrome* (DDS).
 - □*Beckwith-Wiedemann syndrome*.

Morphology

• Gross:

- Large, well-circumscribed soft, tan-gray, homogenous tumor.
- Microscopic*:
 - Classic TRIPHASIC components:
 - **Blastemal**: sheets of undismall blue cells
 - **Epithelial****: abortive tubules or glomeruli.
 - Stromal.



Wilm's tumor





Wilm's tumor



Clinical features

Palpable *abdominal mass*, (may cross the midline) + hematuria, abdominal pain or hypertension

Prognosis:

• Generally *good* (currently 90% long term survival).

• Treatment:

Combination of nephrectomy & chemotherapy

Malignant renal tumors

3. Urothelial Carcinoma of the Renal Pelvis

- **5-10%** of renal neoplasms
- Often *small* and present *early* with, due to:
 - Painless hematuria.
 - Pain or mass due to hydronephrosis.
- Linked to Lynch syndrome & analgesic nephropathy.
- In **50%** of renal pelvic tumors there is a preexisting or concomitant bladder urothelial tumor* (field effect).
- Prognosis: Variable, depend on stage & grade.
 Despite removal by nephrectomy → 50% 5 yrs

Urothelial carcinoma of renal pelvis





Diseases of ureters and lower urinary tract

Ureters

- Sites of narrowing:
 - Ureteropelvic junction.
 - Ureterovesical junction.
 - Crossing of iliac vessels.
- In female pelvis, they lie close to uterine arteries and the cervix. Operations of the female genital tract and diseases of the cervix and uterus may affect ureters.

Causes of Ureteral Obstruction

Intrinsic

- Calculi
- Strictures
- Tumors
- Blood clots
- Neurogenic

Extrinsic

- Pregnancy
- Retroperitoneal fibrosis*
- Endometriosis
- Tumors

Urinary bladder **5. Carcinoma of urinary bladder**

- Epidemiology:
 - \circ 7% of cancers & 3% of cancer deaths.
 - M>F. 50-80 yrs.
- Types:
 - Transitional (urothelial) cell carcinoma (>90%).
 - ✤ Squamous cell carcinoma (5-7%).
 - ♣ Adenocarcinoma (~2%).
 - ✤ Others.

Urothelial (transitional cell) neoplasms

- May involve any site of the urinary collecting system from renal pelvis to urethra → Most commonly seen in the bladder.
- Many are **multifocal** at presentation

Urothelial carcinoma - Etiology

- Mostly are sporadic with unknown cause.
- Possible causes:
 - Cigarette smoking (*most important*).
 - Dye and rubber industry
 - Aromatic amines 2 naphthylamine
 - 4-aminobiphenyl
 - Benzidine.
 - Azo dyes.

- Cyclophosphamide
- Phenacetin containing analgesics
- S. Hematobium infestation (SCC+TCC).
- Previous radiation.
- Induction time ~22yrs.

Classification of urothelial neoplasms			
	Papillary (commoner)	Flat	
Non-invasive	 WHO/ISUP grade: Urothelial papilloma Papillary urothelial neoplasms of low malignant potential (PUNLMP) Papillary urothelial carcinoma, low grade Papillary urothelial carcinoma, high grade 	Carcinoma in-situ By definition <i>high</i> grade	
Invasive	Papillary and invasive More in high grade	Flat and invasive	

Morphologic patterns of bladder tumors



Papillomapapillary carcinoma



Invasive papillary carcinoma



Flat noninvasive carcinoma (CIS)



Flat invasive carcinoma





Exophytic tumor (Most arise from the **lateral** or posterior walls)

Papilloma



-Small papillary fronds lined by **normal-appearing** urothelium -Some have endophytic growth (**inverted papilloma**)





Resembles papilloma but larger & is covered by thicker urothelium

Low-grade papillary urothelial carcinoma



Mild degree of atypia & mitosis with *preserved polarity of cells; the cells are evenly spaced and cohesive*

High-grade papillary urothelial carcinoma



Increased atypica & mitosis with loss of polarity & dyscohesive cells



Carcinoma in situ (CIS) or flat urothelial carcinoma



Defined by the presence of any cytologically malignant cells within a flat urothelium (high grade by definition).
 Accounts for less than 1-3% of urothelial neoplasms → seen in association with invasive carcinoma (45-65%)

Invasive urothelial cancer

- May be flat or papillary, low or high grade:
 - < 10% of low-grade cancers are invasive.</p>
 - 80% of high-grade carcinomas (including CIS) are invasive.
- May be superficial (to lamina propria) or deeper (to muscularis propria).
 - The extent of the invasion is the most important prognostic factor

Non-invasive vs invasive TCC



Other bladder tumors

- <u>Squamous cell carcinoma (5%)</u>, linked to:
 - Schistosomiasis
 - Chronic bladder irritation.
 - Tobacco smoking
- <u>Adenocarcinoma</u> (2%), linked to:
 - Urachal remnants.
 - Bladder exstrophy.
 - Cystitis glandularis.
- Sarcomas.
- Paraganglioma (*micturition syncope*).
- Carcinoid.

Clinical features & prognosis

- *Painless hematuria* is the dominant symptom.
- Lesions that invade the ureteral or urethral or ifices cause urinary tract obstruction.

Prognosis:

- Overall 5-year survival is 57%.
- Depends on:
 - The depth of invasion of the lesion (stage).
 - Histologic grade.

Urothelial (Transitional) Cell Carcinoma

	Low grade	High grade
Gross appearance	Papillary	Nodular flat, papillary
Invasion	<10%	80%-90%
Adjacent dysplasia & CIS	none	Frequent
Recurrences	50%	80%-90%
Progression of disease	<10%	65%
10 year survival	>90%	40%