

#### Miss Eva Bolton Consultant Urologist (Bladder and Renal) Imperial College Healthcare NHS Trust



- Haematuria is the presence of red blood cells in the urine
- It can either be:
  - Macroscopic
    - (Visible / gross haematuria )
  - Microscopic
    - (Non-visible / dipstick positive haematuria)







# The History



- Hx
  - Painful or painless
  - How long is the history
  - Any previous episodes
  - Associated LUTS
  - Associated fever/rigor
- PMH
  - Stones
  - RCC/TCC
  - UTI
- FH
  - Stones
  - RCC/TCC
- SH
  - Alanine dyes
  - Benzen derivatives
  - Smoker



#### **Examination and Investigations**



• O/E

- What does the urine look like
- Retention?
- Any palpable renal or bladder mass

• Ix

- Bedside
  - Urine dip
- Lab
  - MSU
  - FBC, U+E
- Flexible cystosocpy
- Imaging
  - CT urogram/USKUB



#### Location, Location, Location!





# When to refer? – significant health care burden

• 27% of all Urologic evaluation

#### Non-visible haematuria

- +ve urine dip with
  - High WCC
  - Dysuria
- Or clinical suspicion of TCC/RCC
- Micro haematuria noted in 6.5% of healthy participants in screening study (2.4-31.1%)

#### Visible haematuria

- No Age /Risk Stratification Decision
- Haematuria with no associated UTI or continues despite treatment of UTI
- Or clinical suspicion of TCC/RCC
- Acute referral if clots/claret haematuria or Hb drop



#### Microscopic haematuria

- Ignore samples with a trace of blood. Abnormal only if 1+ of blood or more. If painless
  must have 1+ of blood or more on at least 2 out of 3 occasions.
- Do not attribute microscopic haematuria to aspirin or warfarin.
- Investigate as per the flow chart above.
- Refer patients aged 60 or over with unexplained non-visible haematuria and either dysuria or leucocytosis via 2ww pathway for investigation for bladder cancer.



#### PROSPECTIVE ANALYSIS OF 1,930 PATIENTS WITH HEMATURIA TO EVALUATE CURRENT DIAGNOSTIC PRACTICE

M.H. Khadra, R.S. Pickard, M. Charlton, P.H. Powell, D.E. Neal

1,930 patients -prospectively study in hematuria clinic (October 1994 and March 1997)

Basic demographics, history and examination, routine blood tests, urinalysis and cytology. All patients underwent plain abdominal radiography, renal ultrasound, IVP and flexible cystoscopy.

RESULTS: 1,194 males and 736 females ,Age of 58 years (range 17 to 96)

Overall, 61% of patients had no basis found for hematuria, 12% had bladder cancer, 13% had urinary tract infection and 2% had stones. Kidney and upper tract tumors were noted in 14 patients (0.7%), including 4 who presented with microscopic hematuria. If only ultrasound or IVP had been performed 4 of these cases would have been missed. Of 982 patients presenting with microscopic hematuria 51 had cancer. Bladder cancer was found in 7 patients younger than 40 years.

•At age 50 to 59 years, malignancy was identified in 20.4 versus 1.9 percent of men with macroscopic versus microscopic hematuria, respectively, and in 8.9 versus 1.9 percent of women with macroscopic versus microscopic hematuria.

•At age 60 to 69 years, malignancy was found in 28.9 versus 7.9 percent of men with macroscopic versus microscopic hematuria, and 21.1 versus 4.5 percent of women with macroscopic versus microscopic hematuria



# Follow-up after negative initial investigations

\*\*No published guidelines

- Microhaematuria patient
  - Yearly Urine Analysis
  - Discontinue if –ve for 2 years
  - Re-evaluate within 3 to 5 years if persistent Microhaematuria
- Consider full evaluation if recurrence of Gross Haematuria Davis R J et al 2012



#### **Bladder Cancer Guidance**

Recommendations	<ul> <li>Refer people using a suspected cancer pathway referral (for an appointment within 2 weeks) for bladder cancer if they are:</li> <li>aged 45 and over and have: <ul> <li>unexplained visible haematuria without urinary tract infection or</li> <li>visible haematuria that persists or recurs after successful treatment of urinary tract infection, or</li> </ul> </li> <li>are aged 60 and over and have unexplained non-visible haematuria and either dysuria or a raised white cell count on a blood test. [new 2015]</li> </ul>
	Consider non-urgent referral for bladder cancer in people aged 60 and over with recurrent or persistent unexplained urinary tract infection. [new 2015]

#### **Renal Cancer Guidance**

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Recommendations	<ul> <li>visible haematuria that persists or recurs after successful treatment of urinary tract infection. [new 2015]</li> </ul>



#### **Bladder Cancer**



#### Background

- 2<sup>nd</sup> commonest urological malignancy
- >90% Transitional Cell Carcinoma (TCC)
- 1-7% SCC

Associated with chronic inflammation Schistosoma haematobium, long term catheter

#### • 2% Adenocarcinoma

Rare, poor prognosis

1/3 originate in urachus

Long term complication of bowel implantation into urinary tract



### **Transitional Cell Carcinoma**

- >90% of bladder cancers
- Can involve renal pelvis, ureter, bladder
- Single or multifocal
- Classified as superficial or muscle invasive





## **Bladder Caner Risk factors**

- Cigarette Smoking
  - Major cause in developing world
  - 2-5 fold increased risk of bladder Ca + recurrence
- Male sex
- Age
- Occupational Exposure
  - Aromatic hydrocarbons rubber, dye, hairdressers, etc
- Drugs cyclophosphamide
- Pelvic radiotherapy

#### WARNING

#### Cigarettes cause bladder cancer.

Toxic chemicals in tobacco smoke damage the lining of the bladder causing cancer. The most common sign is blood in the urine.

You have the will. There is a way.



**Health Canada** 



## **Bladder Tumour - Staging**

- Ta non-invasive papillary cancer
- Tis carcinoma in situ high grade
- T1 tumour invades submucosa
- •
- T2a superficial muscle
- T2b deep muscle
- T3a perivesical tissue microscopically
- T3b perivesical tissue macroscopically
- T4a prostate, uterus, vagina
- T4b pelvic wall or abdominal wall



Diagram showing the T stages of bladder cancer © CancerHelp UK



### Bladder Cancer Staging

- N1 single <2cm
- N2 single >2 to 5 cm, multiple < 5cm
- N3 > 5cm
- M0-No
- M1 Yes



#### **Bladder Cancer – Presentation**

- Painless visible haematuria -
  - 20% patients with VH will have malignancy
- Asymptomatic Non Visible Haematuria
  - 5% of patients with NVH will have malignancy
- Irrigative LUTS urgency, suprapubic pain
- Recurrent UTI
- Pain, Weight loss, lymphoedema



National 'blood in pee' campaign 13 October-23 November 2014



## Investigations

- Clinical assessment history, examination, urinalysis, urine cytology
- Upper tract imaging
  - USS KUB
  - CT Urogram/MRI
  - IVU
- Direct visualisation of bladder -CYSTOSCOPY





# Flexible cystoscopy









#### Bladder Cancer – Diagnosis and Staging



- Transurethral Resection of Bladder Tumour (TURBT)
  - GA or Spinal anaesthetic
  - Resect Tumour incl muscle specimen
  - +/- Intravesical Mitomycin
  - 3 way catheter, irrigation
  - Usually overnight stay in hospital
- Staging CT urogram + chest
- MDT



#### Bladder Cancer – Management

- Results discussed in MDT meeting
- Further treatment depends on grade and stage of tumour ?muscle invasive ?metastatic
  - Muscle invasive Cystectomy /Radiotherapy/chemotherapy
  - Superficial surveillance cystoscopies, intra-vesical chemotherapy



### Management of Superficial TCC

- Intravesical Chemotherapy
  - Mitomycin C (MMC), single dose, weekly course. Inhibits DNA synthesis. Reduces risk of recurrence.
  - Bacille Calmette-Guerin (BCG)- immune stimulant. Given as 6 wk course via catheter. Reduces progression. More effective than MMC but more toxic.
  - BCG toxicity fever, myalgia, irritative symptoms, may need anti-TB therapy for severe systemic symptoms
  - BCG contraindicated in immunosuppressed, pregnant/lactating, after traumatic catheterisation



#### Flexible Cystoscopy Surveillance

- Flexible cystoscopy
- Day unit procedure
- Local anaesthetic lignocaine gel
- Risks: Bleeding, infection





# Management of muscle invasive TCC – radical cystectomy

- 5 year survival 3% for MIBC
- Radical Cystectomy
  - Ileal conduit formation
  - Neo-bladder formation
- High risk surgery
  - 25% have major complication (death, reoperation, sepsis, thromboembolism), ITU care required

• Stoma care



## **Ileal Conduit**

- Stoma Care
- Complications- ileus, urinary leak, stricture, enteral leak, stoma problems









### Management of MIBC

- Radical External Beam
   Radiotherapy
  - Pts unfit/ unwilling to have surgery
  - Results inferior to radical cystectomy
  - May use neo-adjuvant chemotherapy
  - Complications radiation cystitis, proctitis,

- Palliative treatment
  - Radiotherapy for metastatic bone pain
  - Symptom control
- Use of chemotherapy in locally advanced / metastatic disease



#### **Renal Cancer**



#### **Renal Tumours**

- Benign v Malignant
- Primary v Secondary

#### <u>Benign</u>

Adenoma (Von Hippel Lindau)

Haemangioma

Angiomyolipoma (CT Dx)

*Juxtaglomerular tumour (↑BP)* 



### Malignant Parenchymal

- Renal Cell Carcinoma (RCC)
- Wilms Tumour
- Sarcomas rare, poor prognosis.
- Secondaries
  - breast
  - Lung
  - malignant melanoma
  - *lymphomas*



### Malignant Renal Pelvis

- Transitional cell carcinoma
- Present with haematuria / obstruction
- Association with bladder & ureteric tumours
- Squamous cell carcinoma metaplasia secondary to chronic irritation





### Renal Cell Carcinoma

- Incidence 3% of cancers
- Age mainly over 50 yrs
- Sex 2 M: 1 F
- Geography
- Predisposing factors genes (VHL), APKD
- Microscopic adenocarcinoma
- Macroscopic solid lump, occasionally cystic.





# RCC – clinical

- Presentation
  - incidental finding
  - haematuria, pain, +/- mass
- Diagnosis
  - cross-sectional imaging. 95% solid renal masses=RCC
- Treatment
  - Radical surgery
  - Nephron sparing surgery
  - Survaillance





### **Renal Cell Carcinoma**

#### • Uncommon symptoms

Iron deficiency anaemia

Polycythaemia

Hypertension

Hypercalcaemia due to parathormone-like protein production

PUO

Elevated ESR

Secondary lesions – cannonball lesion on chest X-ray



#### **RCC Staging** Gerota's Adrenal fascia gland Stage I Aorta Tumor <7 cm in greatest dimension and Kidney limited to kidney; 5-year survival, 95% Inferior Stage II vena cava Tumor >7 cm in greatest dimension and limited to kidney; 5-year survival, 88% Stage III Tumor in major veins or adrenal gland, Lymph' nodes tumor within Gerota's fascia, or 1 regional lymph node involved; 5-year survival, 59% Stage IV Tumor beyond Gerota's fascia or >1 regional lymph node involved; 5-year survival, 20%



## Renal Cell Carcinoma

- Investigations
  - US
  - CT
  - Arteriography (MR Angiogram)
  - FBC
  - U&Es
  - GFR
  - Chest X-ray



## Laboratory Findings

- Anaemia
- Haematuria
- 个 ESR
- Paraneoplastic syndrome erythrocytosis hypercalcaemia
  - hypertension

# Imaging - CT

- Gold Standard
- Enhances with IV contrast
- Heterogeneous pattern
- Pre-operative staging





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#### Treatment

- Localised disease: *Stage I,II,IIIA =* Nephrectomy/Parital nephrectomy
- Laparoscopic (robotic) v Open
- En bloc removal: ipsilateral adrenal, proximal ureter, Gerota's fascia
- Curative
- 2-3% recurrence rate



#### **Radical Nephrectomy**







#### Laparoscopic versus Radical Nephrectomy













# **Nephron Sparing Options**

Ablative therapies



Partial nephrectomy (gold standard)



ROBOTIC ASSISTED PARTIAL NEPHRECTOMY



#### Metastatic Disease

- 30% present with advanced disease
- Radiation therapy *effective palliation in 2/3*
- Chemotherapy poor
- Immunotherapy TK inhibitors, interferon, interleukin 2- adjunct to debulking surgery







#### Any Questions?