Analysis of urine and the urinary sediment.

urine analysis

• Urine analysis, – one of the oldest laboratory procedures in the practice of medicine.

Urine analysis and microscophy

• Is an array of tests performed on urine, and one of the most common methods of medical diagnosis.



Courtesy of the National Library of Medicine

Reason for urinalysis:

- General evaluation of health
- Diagnosis of disease or disorders of the kidneys or urinary tract
- Diagnosis of other systemic disease that affect kidney function
- Monitoring of patients with diabetes
- Screening for drug abuse (eg. Sulfonamide or aminoglycosides)

TEST

Collection of urine specimens

- Improper collection---- may invalidate the results
- Containers for collection of urine should be wide mouthed, clean and dry.
- Analysed within 2 hours of collection else requires refrigeration.



Types of urine sample

Sample type	Sampling	Purpose
Random specimen	No specific time most common, taken anytime of day	Routine screening, chemical & FEME
Morning sample	First urine in the morning, most concentrated	Pregnancy test, microscopic test
Clean catch midstream	Discard first few ml, collect the rest	Culture
24 hours	All the urine passed during the day and night and next day I st sample is collected.	used for quantitative and qualitative analysis of substances
Postprandial	2 hours after meal	Determine glucose in diabetic monitoring
Supra-pubic aspired	Needle aspiration	Obtaining sterile urine



a





b

a: clean catch urine collection method

in children

b: Suprapubic aspiration of urine. c: Urine storage and transportation kit

Urinalysis; measurements

- Urinalysis consists of the following measurements:
 - Macroscopic or physical examination
 - Chemical examination
 - Microscopic examination of the sediment
 - Urine culture

TEST



Physical examination of urine

Examination of physical characteristics:

- Volume
- Color
- Odor
- pH and
- Specific gravity
 - The refractometer or a reagent strip is used to measure specific gravity

TEST

Physical examination continued...

- Normal- 1-2.5 L/day
- Oliguria- Urine Output < 400ml/day Seen in
 - Dehydration
 - Shock

Volume

- Acute glomerulonephritis
- Renal Failure
- Polyuria- Urine Output > 2.5 L/day Seen in
 - Increased water ingestion
 - Diabetes mellitus and insipidus.
- Anuria- Urine output < 100ml/day Seen in renal shut down

Color • Normal- pale yellow in color due to pigments urochrome, urobilin and uroerythrin.

- Cloudiness may be caused by excessive cellular material or protein, crystallization or precipitation of non pathological salts upon standing at room temperature or in the refrigerator.
- Colour of urine depending upon it's constituents.

Physical examination continued

Color • Abnormal colors:

- Colorless diabetes, diuretics.
- Deep Yellow concentrated urine,

excess bile pigments, jaundice

Blue Green	<u>Pink-Orange-</u> Red	Red-brown-black
Methylene Blue	Haemoglobin	Haemoglobin
Pseudomonas	Myoglobin	Myoglobin
Riboflavin	Phenolpthalein	Red blood cells
	Porphyrins	Homogentisic Acid
	Rifampicin	L -DOPA
		Melanin
		Methyldopa



	Pathological	Non pathological
White	Chyle Pus	Phosphates
Yellow to Orange	Bilirubin Urobilin	Concentrated urine Carrots Senna Riboflavin Acriflavine sulfasalazine
Pink to Red	Haemoglobin Myoglobin Porphyrins Red blood cells	Beets(anthocynin) Aminopyrine Methyldopa Food color Bromosulfonphthalein Pyridium Senna

	Pathological	Nonpathological
Red to Brown to Purple	Porphobilinogen Uroporphyrin	
Brown to Black	Homogenistic acid Melanin Myoglobin Methaemoglobin Phenol Porphyrins	Chloroquine Iron compounds Levodopa Metronidazole Quinine
Blue to Green	Biliverdin Pseudomonas infection	Acriflavine Azure A Methylene blue Vit B Phenyl salicylate Amitryptiline

ODOUR

TEST

- Fruity/sweet odour- presence of ketones.
- Pungent smell- presence of bacteria/ specimen contaminated with bacteria.
- Sweaty feet- Isolvaleric acidemia
- Misty/mousy odour- Phenylketonuria.
- > Maple syrup- Congenital metabolic disorder.
- Fishy odour/Rancid butter- Hypermethioninemia

рΗ

- Concentration ability of kidney to maintain normal hydrogen ion concentration
- Normal pH 4.6 to 8.0
- Average- 6.0

• PROCEDURE

- Dip the litmus paper strips in the urine, remove and read the color change immediately.
- Blue litmus turns red acid
- Red litmus turns blue alkaline

TEST

• Decrease in pH

- High protein intake
- Ingestion of cranberries
- Respiratory acidosis
- Metabolic acidosis
- Uremia
- Severe diarrhoea
- Starvation
- UTI caused by E.coli

• Increase in pH

- Diet high in vegetables and citrus fruits
- Respiratory alkalosis
- Metabolic alkalosis
- Vomiting
- UTI caused by Proteus and Pseudomonas

рН

- Reflects ability of kidney to maintain normal hydrogen ion concentration in plasma & ECF
- Urine pH ranges from 4.5 to 8
- Normally it is slightly acidic lying between 6 6.5.
- Tested by:
 - litmus paper
 - pH paper
 - dipsticks
- Acidic Urine –Ketosis (diabetes, starvation, fever), systemic acidosis, UTI- E.coli, acidification therapy
- Alkaline Urine after meal, systemic alkalosis, UTI proteus, alkalization therapy

Specific gravity

- It is measurement of urine density which reflects the ability of the kidney to concentrate or dilute the urine relative to the plasma from which it is filtered.
- Measured by:
 - urinometer
 - refractometer
 - dipsticks



Specific gravity

• Normal :- 1.001-1.040.

S.G	Osmolality (mosm/kg)
1.001	100
1.010	300
1.020	800
1.025	1000
1.030	1200
1.040	1400

- Increase in Specific Gravity Low water intake, Diabetes mellitus, Albuminuruia, Acute nephritis.
- Decrease in Specific Gravity Absence of ADH, Renal Tubular damage.
- Fixed specific gravity (isosthenuria)=1.010

- A sample of well-mixed urine (usually 10-15 ml) is centrifuged in a test tube at relatively low speed (about 2000-3,000 rpm) for 5-10 minutes which produces a concentration of sediment (cellular matter) at the bottom of the tube.
- A drop of sediment is poured onto a glass slide, a thin slice of glass (a coverslip) is place over it ond observed under microscope

TEST

Microscopic examination of urine

- A variety of normal and abnormal cellular elements may be seen in urine sediment such as:
 - Red blood cells
 - White blood cells
 - Mucus
 - Various epithelial cells
 - Various crystals
 - Bacteria
 - Casts

Abnormal findings

- Per High Power Field (HPF) (400x)
 - > 3 erythrocytes
 - > 5 leukocytes
 - > 2 renal tubular cells
 - > 10 bacteria
- Per Low Power Field (LPF) (200x)
 - > 3 hyaline casts or > 1 granular cast
 - > 10 squamous cells (indicative of contaminated specin
 - Any other cast (RBCs, WBCs)
- Presence of:
 - Fungal hyphae or yeast, parasite, viral inclusions
 - Pathological crystals (cystine, leucine, tyrosine)
 - Large number of uric acid or calcium oxalate crystals

- Hematuria is the presence of abnormal numbers of red cells in urine due to any of several possible causes.
 - glomerular damage,
 - tumors which erode the urinary tract anywhere along its length,
 - kidney trauma,
 - urinary tract stones,
 - acute tubular necrosis,
 - upper and lower urinary tract infections,
 - nephrotoxins
- WBC in high numbers indicate inflammation or infection somewhere along the urinary or genital tract



Red blood cells in urine appear as refractile disks



White blood cells in urine

- Urinary casts are cylindrical aggregations of particles that form in the distal nephron, dislodge, and pass into the urine. In urinalysis they indicate kidney disease.
- They form via precipitation of Tamm-Horsfall mucoprotein which is secreted by renal tubule cells.

Types of cast seen :

- Acellular cast: Hyaline casts, Granular casts, Waxy casts, Fatty casts, Pigment casts, Crystal casts.
- <u>Cellular cast</u>: Red cell casts, White cell casts, Epithelial cell cast
- The most common type of cast- hyaline casts are solidified Tamm-Horsfall mucoprotein secreted from the tubular epithelial cells and seen in fever, strenuous exercise, damage to the glomerular capillary.
- Red blood cells may stick together and form red blood cell casts. Such casts are indicative of glomerulonephritis, with leakage of RBC's from glomeruli, or severe tubular damage
- White blood cell casts are most typical for acute pyelonephritis, but they may also be present with glomerulonephritis. Their presence indicates inflammation of the kidney.



Hyaline Cast



Granular Cast



Red blood cell cast in urine



White blood cell cast in urine



A variety of normal and abnormal crystals may be present in the urine sediment

- The chemical analysis of urine us undertaken to evaluate the levels of the following componen:
 - Protein
 - Glucose
 - Ketones
 - Occult blood
 - Bilirubin
 - Urobilinogen
 - Bile salts

- The presence of normal and abnormal chemical elements in the urine are detected using dry reagent strips called dipsticks.
- When the test strip is dipped in urine the reagents are activated and a chemical reaction occurs.
- The chemical reaction results in a specific color change.
- After a specific amount of time has elapse, this color change is compared against a reference color chart provided by the manufacturer of the strips.







The dipstick method of chemical analysis of urine

LEUKOCYTES 2 minutes	NEGATIVE			TRACE	SMALL +	MODERATE + +	LARGE + + +	
NITRITE 60 seconds	NEGATIVE			POSITIVE	POSITIVE	(Any degree of	f uniform pink color is positive)	
UROBILINOGEN 60 seconds	NORMAL 0.2	NORMAL 1		mg/dL 2	4	8	(1 mg = approx	«. 1EU)
PROTEIN 60 seconds	NEGATIVE	TRACE		mg/dL 30 +	100 + +	300 + + +	2000 or more + + + +	
pH 60 seconds	5.0	6.0		6.5	7.0	7.5	8.0	8.5
BLOOD 60 seconds	NEGATIVE	NON- HEMOLYZED TRACE	147 - 4 17 - 4 19 - 4	NON- Hemolyzed Moderate	HEMOLYZED TRACE	SMALL +	MODERATE + +	LARGE +++
SPECIFIC GRAVI 45 seconds	TY 1.000	1.005		1.010	1.015	1.020	1.025	1.030
KETONE 40 seconds	NEGATIVE		mg/dL	TRACE 5	SMALL 15	MODERATE 40	LARGE 80	LARGE 160
BILIRUBIN 30 seconds	NEGATIVE			SMALL +	MODERATE + +	LARGE + + +		
GLUCOSE 30 seconds	NEGATIVE		g/dL (%) mg/dL	1/10 (tr.) 100	1/4 250	1/2 500	11000	2 or more 2000 or more
	LEUKOCYTES 2 minutes NITRITE 60 seconds UROBILINOGEN 60 seconds PROTEIN 60 seconds BLOOD 60 seconds BLOOD 60 seconds SPECIFIC GRAVI 45 seconds KETONE 40 seconds BILIRUBIN 30 seconds GLUCOSE 30 seconds	LEUKOCYTES 2 minutesNEGATIVENITRITE 60 secondsNEGATIVEUROBILINOGEN 60 secondsNORMAL 0.2PROTEIN 60 secondsNEGATIVEPH 60 seconds5.0BLOOD 60 secondsNEGATIVESPECIFIC GRAVITY 45 seconds1.000KETONE 40 secondsNEGATIVEBILIRUBIN 30 secondsNEGATIVEGLUCOSE 30 secondsNEGATIVE	LEUKOCYTES NEGATIVE NITRITE NEGATIVE G0 seconds NORMAL UROBILINOGEN NORMAL G0 seconds 0.2 PROTEIN NEGATIVE G0 seconds NEGATIVE PROTEIN NEGATIVE G0 seconds NEGATIVE PH 5.0 60 seconds NEGATIVE BLOOD NEGATIVE G0 seconds NEGATIVE BLOOD NEGATIVE 60 seconds NEGATIVE SPECIFIC GRAVITY 1.000 45 seconds 1.000 KETONE NEGATIVE 40 seconds NEGATIVE BILIRUBIN NEGATIVE 30 seconds NEGATIVE GLUCOSE NEGATIVE 30 seconds NEGATIVE	LEUKOCYTES NEGATIVE NITRITE NEGATIVE 60 seconds NEGATIVE UROBILINOGEN NORMAL 60 seconds NORMAL 0.2 NORMAL PROTEIN NEGATIVE 60 seconds NEGATIVE PH 5.0 60 seconds NEGATIVE SPECIFIC GRAVITY 1.000 45 seconds NEGATIVE KETONE NEGATIVE 40 seconds NEGATIVE BILIRUBIN NEGATIVE 30 seconds NEGATIVE Subseconds NEGATIVE Subseconds NEGATIVE	LEUKOCYTES NEGATIVE TRACE NITRITE NEGATIVE POSITIVE 60 seconds NORMAL NORMAL mg/dL 2 NORMAL 0.2 NORMAL mg/dL 2 PROTEIN NEGATIVE TRACE mg/dL 60 seconds NEGATIVE TRACE mg/dL 30 40 seconds NEGATIVE TRACE mg/dL 30 9H 5.0 6.0 6.5 8LOOD NEGATIVE HEMOLYZED HEMOLYZED 9H 5.0 6.0 6.5 8LOOD NEGATIVE HEMOLYZED HEMOLYZED SPECIFIC GRAVITY 1.000 1.005 1.010 KETONE NEGATIVE mg/dL TRACE 40 seconds NEGATIVE mg/dL TRACE 80 seconds NEGATIVE mg/dL TRACE 30 seconds NEGATIVE g/dL (%) 1/10 (r.) 30 seconds NEGATIVE g/dL (%) 1/10 (r.)	LEUKOCYTES 2 minutes NEBATIVE TRACE SMALL + NITRITE 60 seconds NEGATIVE POSITIVE POSITIVE UROBILINOGEN 60 seconds NORMAL 0.2 NORMAL 0.2 MORMAL 1 mg/dL 2 4 PROTEIN 60 seconds NEGATIVE TRACE mg/dL 30 100 +++ PH 60 seconds 5.0 6.0 6.5 7.0 BLOOD 60 seconds NEGATIVE HEMOLYZED TRACE HEMOLYZED WODERATE HEMOLYZED WODERATE SPECIFIC GRAVITY 45 seconds 1.000 1.005 1.010 1.015 KETONE 40 seconds NEGATIVE mg/dL TRACE 5 SMALL 5 BILIRUBIN 30 seconds NEGATIVE g/dL (%) mg/dL 1100 114 250	LEUKOCYTES NEGATIVE TRACE SMALL MODERATE NITRITE NEGATIVE POSITIVE POSITIVE POSITIVE Any degree of the seconds UROBILINOGEN NORMAL NORMAL mg/dL mg/dL 4 8 PROTEIN NORMAL NORMAL mg/dL 100 300 60 seconds NEGATIVE TRACE mg/dL 100 300 60 seconds NEGATIVE TRACE mg/dL 100 300 60 seconds NEGATIVE TRACE mg/dL 100 300 60 seconds 6.0 6.5 7.0 7.5 BLOOD 6.0 6.5 7.0 7.5 BLOOD NEGATIVE HEMOLIZZED HEMOLIZZED SMALL 60 seconds NEGATIVE HEMOLIZZED SMALL NODERATE SPECIFIC GRAVITY 1.000 1.005 1.010 1.015 1.020 KETONE NEGATIVE mg/dL TRACE SMALL MODERATE MODERATE SUBLIRUBIN NEGATIVE mg/dL TRACE SMAL	LEUKOCYTES 2 minutes REBATIVE TRACE SMALL MODERATE LARCE NITRITE 60 seconds NEBATIVE POSITIVE POSITIVE POSITIVE (Any degree of uniform pink color is positive for seconds UROBILINOGEN 60 seconds NORMAL 0.2 NORMAL 0.2 mg/dL mg/dL 4 8 (I mg = approx PROTEIN 60 seconds NEGATIVE TRACE mg/dL 30 100 4 8 (I mg = approx PROTEIN 60 seconds NEGATIVE TRACE mg/dL 30 100 4 8 (I mg = approx PBOTEIN 60 seconds NEGATIVE TRACE mg/dL 30 100 100 200 200 gr more 4+++ PH 60 seconds 5.0 6.0 6.5 7.0 7.5 8.0 BLOOD 60 seconds NEGATIVE HEMOLYTER HEMOLYTER SMALL MODERATE SPECIFIC GRAVITY 45 seconds 1.000 1.010 1.015 1.020 1.023 SUBJURDIN 40 seconds NEGATIVE mg/dL TRACE SMALL MODERATE LARGE SUBJURDIN 30 seconds

Proteins in urine:

- Detected by heat coagulation or dipstick method
- Urine proteins come from plasma protein and Tomm-Horsfall (T-H) glycoprotein
- healthy individuals excrete <150 mg/d of total protein and <30 mg/d of albumin.
- Plasma cell dyscrasias (multiple myeloma) can be associated with large amounts of excreted light chains in the urine, which may not be detected by dipstick. The light chains produced from these disorders are filtered by the glomerulus and overwhelm the reabsorptive capacity of the proximal tubule and Bence Jones proteinuria occurs

Chemical analysis of urine **EVALUATION OF PROTEINURIA** PROTEINURIA ON URINE DIPSTICK Quantify by 24-h urinary excretion of protein and albumin or first morning spot albumin-to-creatinine ratio Microalbuminuria Macroalbuminuria Nephrotic range 30-300 mg/d or 300-3500 mg/d or > 3500 mg/d or 30-300 mg/g 300-3500 mg/g > 3500 mg/g Go to + RBCs or RBC casts on urinalysis Fig. 44-2 In addition to disorders listed under microalbuminuria consider Consider Myeloma-associated kidney Nephrotic syndrome disease (check UPEP) Early diabetes Diabetes Essential hypertension Intermittent proteinuria Amyloidosis Postural proteinuria Early stages of Minimal change disease glomerulonephritis Congestive heart failure FSGS (especially with RBCs, Fever Membranous glomerulopathy RBC casts) Exercise

Proteins in urine

- Normal- upto 150 mg/24 hours or 10mg/100ml in single sample.
- Methods-
- Heat and acetic acid test- The test is based on the principle of heat coagulation and precipitation of proteins by acetic acid.
- **Sulphosalicylic acid test-** Sulphosalicylic acid neutralizes protein cation, resulting in precipitation of protein.

Causes of proteinuria

TEST

- Pre-renal
- >Addison's disease
- > Fever
- ➤ Eclampsia
- > Hypertension
- ➢ Haemoglobinuria
- ➢ Rhabdomyolysis

Renal → All cases of glomerulonephritis → Nephrotic syndrome → Pyelonephritis

- Post renal
- Lesions of renal pelvis, urethra (cystitis, prostatitis)
 Severe UTI
Cause of Proteinuria as Related to Quantity



• MINIMAL PROTEINURIA (<0.5 gm/day)

➢ Exercise

> Fever

Emotional stress

>HTN

>Renal tubular dysfunction

Polycystic kidneys

≻ Lower UTI

MODERATE PROTEINURIA (0.5-3 gm/day)

Chronic glomerulonephritis
CCF
Pyelonephritis
Pre-eclampsia
Multiple myeloma

MARKED PROTEINURIA (> 3gm/day)

Acute glomerulonephritis
Chronic glomerulonephritis, severe
Nephrotic syndrome
Diabetic nephropathy, severe
Renal amyloidosis
Lupus nephritis

Classification of Proteinuria

TYPE	PATHOPHYSIOLOGIC FEATURES	CAUSE		
Glomerular	Increased glomerular capillary permeability to protein	Primary or secondary glomerulopathy		
Tubular	Decreased tubular reabsorption of proteins in glomerular filtrate	Tubular or interstitial disease		
Overflow	Increased production of low-molecular- weight proteins			

Selected Causes of Proteinuria by Type

Glomerular Primary glomerulonephropathy □ Minimal change disease □Idiopathic membranous glomerulonephritis □Focal segmental glomerulonephritis Membranoproliferative glomerulonephritis □IgA nephropathy

Secondary glomerulonephropathy ✓ Diabetes mellitus ✓ Collagen vascular disorders (e.g., lupus nephritis) ✓ Amyloidosis ✓ Preeclampsia ✓ Infection (e.g., HIV, hepatitis B and C, poststreptococcal illness, syphilis, malaria and endocarditis) ✓ Gastrointestinal and lung cancers ✓Lymphoma, chronic renal transplant rejection Glomerulonephropathy associated with the following drugs: ≻Heroin ➢NSAIDs Gold components ➢ Penicillamine ≻Lithium Heavy metals

Tubular Hypertensive nephrosclerosis Tubulointerstitial disease due to:

Uric acid nephropathy
Acute hypersensitivity interstitial nephritis
Fanconi syndrome
Heavy metals
Sickle cell disease
NSAIDs, antibiotics

Overflow

Hemoglobinuria
Myoglobinuria
Multiple myeloma
Amyloidosis

Selective proteinuria

- When LMW proteins like albumin (MW- 66000) or transferrin (MW-76000) are selectively excreted through kidney.
- \succ Eg- all causes of nephrotic syndrome.

Non-selective proteinuria

When HMW protein like globulin, fibrinogen in addition to LMW protein are excreted through kidney

Microalbuminuria

- Urinary albumin excretion between 30-300 mg/day.
- Cannot be detected by dipstick methods.
- Strong predictor of development of diabetic nephropathy.
- Can be detected 10-15 years before development of diabetic nephropathy.
- Significant risk marker of cardiovascular ds.
- Measured by nephelometry and radioimmunoassay

Diagnostic relevance microalbuminuria

- In diabetic patients for early diagnosis of nephropathy.
- In hypertensive patients as indicator of end organ damage

Bence Jones proteins

- BJ protein is abnormal LMW globulin consisting of light chains of Ig either Lambda or Kappa chains.
- Characteristic feature- PPT at 40^o C to 60_o C and redissolves at higher temperature (100^oc) & reappears when the urine is cooled.
- Conditions a/w BJ proteinuria:
- Multiple myeloma
- Plasmacytoma
- Waldesnstrom macroglobinaemia

SUGARS IN URINE

 This is a non-specific test useful for semiquantitation of marked glucosuria.

Benedict's qualitative test

 Principle- Aldehyde group of reducing sugar reduces Cupric ions in Benedict's reagent to cuprous oxide.

Detects all sugars except sucrose.

The final color of the solution depends on how much of this precipitate was formed, and therefore the color gives an indication of how much reducing sugar was present.

Increasing amounts of reducing sugar
Green yellow orange red

Sugars detected by Benedict's test-

- ➢ Glucose
- ➢ Galactose
- ➢ Lactose
- ➢ Fructose
- ➢ Maltose
- > Pentose
- False +
- >Ascorbic acid, Creatinine, Uric acid
- ➤ Salicylates
- >X-ray contrast

COLORIMETRIC REAGENT STRIP TEST

- **Principle**: this test is based on a double sequential enzyme reaction.
- One enzyme, glucose oxidase, catalyzes the formation of gluconic acid and hydrogen peroxide from the oxidation of glucose.
- A second enzyme, peroxides catalyzes the reaction of hydrogen peroxide with potassium iodide chromogen to oxidize the chromogen to colors ranging from green to brown.





TESTS AND READIN	G TIME			the strength			
LEUKOCYTES 2 minutos	NEGATAR		TRACE	5HALL	5000H431E	LANCE	
NITRITE 60 seconds	RERATINE		Posimet	Pasilar	(Bry degree of an door	a pirti culto: n pominu)	1
UROBILINOGEN 60 seconds	KÓRYAL 3.5	улин	*511	•		(1 mg = antimer and)	L.C.
PROTEIN 60 seconds	RECAILLY	TRACE	18(30) 34 *	120	***	2000 07 A MR	
pH 60 seconda	50	a.	6.5	2.5	7.5	+.5	1.5
BLOOD 6D seconds	REGATINE	HEINOS BALL	HING THE TOTAL	FERDING TELE	enent	MDDERUTE	Linese
SPECIFIC GRAVIT 45 seconds	1400 L	1408	1,914	1475	1.83	1.035	1.530
KETONE 40 seconds	BIGATINE	nçır.	TRACY	swig	MOOR RATE 40	Mag and	ra#\$\$
BILIBUBIN 30 seconds	SEGRITAR		tamit	MODERATE	uns:		
GLUCOSE 30 seconds	NECATING	19, 154 15 ga	120 120	11 250	12 500	1000	2003 or mano



KETONES IN URINE (ketonuria):

• TYPES

• Acetone, diacetic acid (acetoacetic acid), betahydroxybutyric acid.



TEST

• Causes of Ketonuria:

DKA > Fever > Anorexia \succ Gastrointestinal disturbances ➢ Fasting ➢ Starvation Severe vomiting

OCCULT BLOOD IN URINE:

- Red blood cells / haemoglobin.
- Haematuria- when 5 or more intact RBCs/HPF.

Causes of Haemoglobinuria

- Malaria- black water fever.
- Hemolytic streptococcal septicaemia.
- Incompatible blood transfusion.
- Drugs- Sulphonamides, phenylhydralazine.
- PNH

Causes of Haematuria

• Renal

- ≻ Neoplasms
- Calculi
- ≻ TB
- > Pyelonephritis
- > Hydronephrosis
- ➢ Oxaluria
- ≻ Acute GN
- Polycystic kidney ds

Post-Renal

- > Ureter- calculus, neoplasm
- Urinary bladder- neoplasm, TB, Cystitis, calculus.
- > Prostate- BPH, Neoplasm
- General
- > Embolism of kidney from SBE.
- > Malignant HTN kidney
- ➢ Haemophilia
- Leukemia

BILIRUBIN METABOLISM



Bile salts

• Primary bile acids

- Cholic acid and chenodeoxycholic acid (CDCA)- synthesized from cholesterol in the liver, conjugated with glycine or taurine, and secreted into the bile.
- Secondary bile acids
- Deoxycholate and lithocholate, are formed in the colon as bacterial metabolites of the primary bile acids.
- Sodium taurocholate and sodium glycocholate are found in urine.

Bile pigments

- Normal urine-
- ➤ Urochrome
- Traces of Urobilin
- Abnormal urine
- ➢ Bilirubin
- > Urobilinogen
- ➢ Biliverdin
- ➤ Urobilin

Causes of increased urobilinogen

- Cirrhosis
- Haemolytic jaundice
- Paralytic enterocolitis
- Hepatic congestion

LABELLING

- Sample container-for identification of sample.
- Cytology requisition form-for identification of individual patient sample.

CENTRIFUGATION

Basically of 2 types-

- I Normal.
- II Cytospin-A device that spins cells in a fluid suspension .
- Orawbacks-distortion of cellular morphology due to air drying artifacts and loss of cells by absorption of fluid into the filter card.
- In this process, urine sample is taken in a conical tube and centrifuge at a rate of 2000rpm for 10-15 minutes.

PAPANICOLAOU STAIN

• Done by two methods-

1 Automated stainer-- large scale slides.

Takes 30 minutes for staining.

2 Manual staining using copplin jar-

For small scale slides.

Takes less than 7 minutes to stain.

- Staining objectives-
 - Well stained nuclear chromatin.
 - II Differential counterstaining i.e. staining the cytoplasm of different cell types into different colours and intensity.
 III Retaining cytoplasmic transparency.

Urine Cytology

- INTERPRETATION-
- 1 Normal-Normal constituents of urine.
- 2 Abnormal-Any variation from normal-

I cellular components

II Acellular components.

CELLS derived from-

- Urothelial and its variants.
- Renal tubules.
- Adjacent organs-like prostrate.
- Cells extragenous to the urinary tract-RBCs.

Urine Cytology....

A Cells-

TEST

- -Erythrocytes-Hematuria
- -Leucocytes-Infective etiology
- -Epithelial cells
- **B** Casts-
 - -Hyaline cast
 - -Red cell cast
 - -Granular cast
 - -Epithelial cast
 - -waxy cast
 - -Fatty cast

C Crystals-Some are common in Acidic urine -Some are common in Alkaline urine

D Bacteria-Normal urine is free from bacteria.

E Yeast.

F Malignant cells.

G Artifacts.
CELLS

Erythrocytes

- usually appear as hourglass appearance.
- presence of red cells 1-2RBCs/HPF is not considered abnormal.
- in hypotonic urine red cells swell up causing lysis -releasing Hb in urine-lysed cells are referred as *ghost cells*.
- when the red cells are swollen/crenated sometimes mistaken for WBCs and yeast cells



Leucocytes

- Normal up to 1-2WBCs/HPF.
- Larger than red cells and smaller than renal epithelial cells.
- Usually spherical-singly/clumps.
- Mostly neutrophils-presence of characteristics granules and lobulation.
- Addition of 2%acetic acid to slide accentuated the nuclei of cells.
- Presence of many white cells in clumps is strongly suggestive of acute urinary tract infection.





Figure 22-14A Urine sediment in inflammation. A low-power view of urine sediment containing numerous leukocytes.

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Epithelial cells

- Any site in genitourinary tract from PCT to the urethra or from the vagina.
- Normally a few cells from these sites can be found.
- A marked increase indicates *inflammation* of that proportion of urinary tract from which the cell is derived.
 TYPES
- Renal tubular
- Transitional
- Squamous

Renal tubular epi. cells

- Larger than white cells
- Large round nucleus.
- May flat/ cuboidal/ columnar.
- Increase no indicates tubular damage.



Transitional epithelial cells

- 2 to 4 times larger than white cells.
- Round/ pear shaped/ may have tail like projection.
- Line the urinary tract from pelvis of kidney to upper portion of urethra.



Squamous epithelial cells

- Line urethra and vagina.
- Have little diagnostic significance.





- Usually not found in fresh urine but appear when urine strands for a while.
- Many of crystal found in urine have little clinical significance except in case of metabolic disorders.
- Crystals are identified by their appearance and their solubility characteristics.

TYPES-

- Acidic urine crystals
- Alkaline urine crystals

CRYSTALS IN ACIDIC URINE

- URIC ACID.
- AMORPHOUS URATES
- CALCIUM OXALATE
- CYSTINE
- LEUCINE
- TYROSINE
- SULPHA

TEST

CRYSTALS IN ALKALINE URINE

- TRIPLE PHOSPHATE
- CALCIUM CARBONATE
- CHOLESTROL
- Amorphous phosphates
- Ammonium biurate

TEST

Uric acid crystals

- Most characteristics form are diamond or rhombic prism.
- Presence of uric acid crystals in urine is a normal appearance.
- Increase in-gout
 - -AFI
 - -Chronic nephritis
 - -high purine metabolism



Calcium oxalate crystals

- Octahedral or envelope shaped crystal
- Can be present in normal urine after ingestion of various oxalate rich foods.
- Pathological- DM

 -Liver disease
 -Severe chronic renal disease



Amorphous urates

- Urates salts of sod, pot. and calcium
- Having a granular appearance
- Present in urine as non crystalline amorphous forms.



• No clinical significance.

Hippuric acid crystals

- Elongated prism like.
- Rarely seen in urine.
- No clinical significance.



Cystine crystals

- Refractile hexagonal plate swith equal or unequal sides.
- Frequently have layered or laminated appearance.
- Soluble in ammonia.
- Can be detected chemically by Sodcyanide-sod. Nitropruside test.
- Always Pathological (Cystinosis).



Leucine crystals

- Highly refractile having spheroid with radical and concentric striations.
- Clinically very significant.
- Maple syrup disease
- Serious liver disease



Tyrosine crystals

- very fine needle likes occurring in sheaves or clusters.
- Clinically significance
- Severe liver disease
- tyrosinosis





Cholesterol crystals

- large or flat plates with notched corners
- Presence of excessive in urine indicates tissue breakdown





CHOLESTEROL CRYSTAL

Sulfa drugs crystals

- precipitate as sheets of needles usually with eccentric binding
- May be history of sulfa drugs medication



Alkaline urine crystals

Triple phosphates crystals-

- prism like with three to six sides
- Frequently found in normal urine
- Pathological

 -chronic pyelitis
 -cystitis
 -enlarged prostate





Amorphous phosphates

- granular particles with no definite shape
- No clinical significance.



Calcium carbonate

- appearing as dumbbell or spherical or large granular mass.
- No clinical significance.





- Presence of casts are frequently associated with proteinuria.
- Have nearly parallel sides with rounded or blunts ends.
- Always renal in origin and indicates intrinsic renal disease
- Casts are more or less circular with thicker in middle.

• meaning renal hematuria

• Always pathological.



White cell cast





Granular casts

- degeneration of cellular casts or direct aggregation of serum proteins.
- Almost always indicate significant renal disease.
- May be fine granular or coarse granular casts.





Epithelial cells casts

- result as statis and desquamation of renal tubular epithelial cells.
- Indicates tubular injury.





Waxy casts

• smooth

homogenous appearance.

- Results from degeneration of granular casts.
- Found in acute and chronic renal disease.



Fatty casts

- Appear as a few fat droplets or compose almost entirely of fat droplets of various sizes.
- Found in fatty degeneration of tubular epithelial.





Hyaline cast

• Damage to glomerular capillary membrane, fever, orthostatic proteinuria, and emotional stress or strenuous exercise.





- When accompanied with white cells usually indicates UTI.
- Occurs as rod or chains or cocci.







Fibres

Starch crystals





Enterobius vermicularis

Schistosoma Haematobioum







- Usually ovoid cells with budding.
- Not dissolve in 2%acetic acid solution and not stained with Eosin.


Malignancy

- Urinary bladder
- Renal pelvis
- Kidney
- Ureter
- Adjacent organs.

Types

- Low grade papillary tumours.
- High grade papillary tumours.





(left) and one showing slight nuclear enlargement and hyperchromasia (right). (A-C: High magnification.)



Figure 23-14B Cancer cells in urinary sediment. Cancer cells, appearing singly and in clusters, in voided urine sediment (ThinPrep).

Figure 7-23D Mitotic abnormalities in cancer cells. Carcinoma of bladder, voided urine sediment with a tumor cell metaphase containing numerous chromosomes. (A,B: High magnification; D: oil immersion.) (A and B Courtesy of Dr. Carlos Rodriguez, Tucuma\$AA\$n, Argentina.)



Figure 33-10 Prostatic carcinoma in voided urine. A small cluster of small cancer cells with relatively large, hyperchromatic nuclei. The details of the nuclear structure are not visible. The prostatic origin of the cluster is not secure.



Figure 23-40C Renal pelvic carcinoma. Voided urine sediment (scanning power) containing small cancer cells corresponding to the tumor of renal pelvis shown in D.



Figure 23-38A Choriocarcinoma of urinary bladder, accompanied by a flat carcinoma in situ (not shown). Cancer cells in voided urine resembling cells of urothelial carcinoma.