# COMMON ABNORMAL LABS

LAURA GUNDER MCCLARY, DHSc, MHE, PA-C

Non-Declaration Statement: I have no relevant relationships with ineligible companies to disclose within the past 24 months. (Note: Ineligible companies are defined as those whose primary business is producing, marketing, selling, re-selling, or distributing healthcare products used by or on patients.)

### OBJECTIVES

1) Identify the most common abnormal lab and their etiology for a complete blood count (CBC)

2) Identify the most common abnormal lab results and their etiology for a basic metabolic panel (BMP)

3) Identify the most common abnormal lab results and their etiology for a urinalysis (UA)

## Leukopenia

- WBC < 4\*10<sup>9</sup> /L High susceptibility to infections Pancytopenia causes:
- bone marrow tumor
- aplastic state of bone marrow
  - ionizing radiation
  - chemotherapy of tumors (cytostatics)
  - intoxication with benzene, myelotoxic drugs

(levomycetine, NSAIDs)

- B12 deficiency
- > overactive spleen.

## Neutropenia

- Benign ethnic neutropenia
- Familial neutropenia
- Congenital neutropenia
- Infection
- Medications
- Nutritional deficiencies (vitamin B12, folate, copper)
- Hematologic malignancies
- Rheumatologic disorders
- Autoimmune neutropenia
- Aplastic anemia
- Chronic idiopathic neutropenia



### LEUKOCYTOSIS

### Table 4. Selected Conditions Associated with Elevations in Certain White Blood Cell Types

<i>White blood</i> cell line	Conditions that typically cause elevations
Basophils	Allergic conditions, leukemias
Eosinophils	Allergic conditions, dermatologic conditions, eosinophilic esophagitis, idiopathic hypereosinophilic syndrome, malignancies, medication reactions, parasitic infections
Lymphocytes	Acute or chronic leukemia, hypersensitivity reaction, infections (viral, pertussis)
Monocytes	Autoimmune disease, infections (Epstein-Barr virus, fungal, protozoan, rickettsial, tuberculosis), splenectomy
Neutrophils	Bone marrow stimulation, chronic inflammation, congenital, infection, medication induced, reactive, splenectomy

Distinguishing features	Evaluation
Pregnancy, obesity, race, age	Reference appropriate WBC count by age or pregnancy trimester Compare WBC count to recent baseline (if available)
Fever, system-specific symptoms Physical examination findings	Obtain system-specific cultures and imaging (e.g., sputum cultures, chest radiography) Consider empiric antibiotics Consider use of other biomarkers, such as CRP and procalcitonin
Exercise, physical stress (e.g., postsurgical, febrile seizures), emotional stress (e.g., panic attacks), smoking	Confirm with history
Rheumatic disease, inflammatory bowel disease, granulomatous disease, vasculitides, chronic hepatitis	Obtain personal and family medical history Consider erythrocyte sedimentation rate and CRP levels, specific rheumatology laboratories Consider subspecialist consultation (e.g., rheumatology, gastroenterology)
Corticosteroids, beta agonists, lithium, epinephrine, colony-stimulating factors	Confirm with history; consider discontinuation of medication, if warranted
Hemolytic anemia, immune thrombocytopenia, bone marrow suppression recovery, colony- stimulating factors	Complete blood count differential; compare with baseline values (if available) Examine peripheral smear Consider reticulocyte and lactate dehydrogenase levels Consider flow cytometry, bone marrow examination, hematology/ oncology consultation
History of trauma or sickle cell disease	Confirm with history
Hereditary/chronic idiopathic neutrophilia, Down syndrome, leukocyte adhesion deficiency	Obtain family, developmental history Consider hematology/oncology, genetics, and immunology consultations
	<ul> <li>Fever, system-specific symptoms Physical examination findings</li> <li>Exercise, physical stress (e.g., postsurgical, febrile seizures), emotional stress (e.g., panic attacks), smoking</li> <li>Rheumatic disease, inflammatory bowel disease, granulomatous disease, vasculitides, chronic hepatitis</li> <li>Corticosteroids, beta agonists, lithium, epinephrine, colony-stimulating factors</li> <li>Hemolytic anemia, immune thrombocytopenia, bone marrow suppression recovery, colony-stimulating factors</li> <li>History of trauma or sickle cell disease</li> <li>Hereditary/chronic idiopathic neutrophilia, Down syndrome,</li> </ul>

Table 3. Nonmalignant Causes of Neutrophilia

NOTE: After patient characteristics, causes are listed in approximate order of frequency.

*CRP* = *C*-reactive protein; *WBC* = white blood cell.

Information from references 1 through 7, 9, and 10.

### History:

23-year-old male

Over the past week noted increasing fatigue, sore throat, earaches, headaches, and episodic fever and chills

Unable to run his customary 20 miles per week

### **Physical Exam:**

Erythematous throat and tonsils Swollen cervical lymph nodes No other organomegaly WBC: 14.4 x 10<sup>3</sup>/uL (4.5-10.5)

RBC: 5.25 x 10<sup>6</sup>/uL (4.2-5.4)

HGB: 15.4 g/dL (14-18)

HCT: 46.1 % (38-48)

MCV: 87.9 fL (80-100)

MCH: 29.3 pg (26-34)

MCHC: 33.4 g/dL (31-37)

RDW: 12.2 % (<14)

Neutrophils: 24 % (65-75) Lymphocytes: 73 % (35-45) Eosinophils: 3 % (0-5) Basophils: 0 % (0-1) Monocytes: 0 % (0-7)

Platelets: 333 x 10<sup>3</sup>/uL (150-400)



#### Normal: < 1%



#### Clincially significant: > 10%

Band neutrophils are slightly less mature than segmented neutrophils and have indented, unsegmented "C" or "S" shaped nuclei.

Band neutrophils normally account for approximately 5-10% of peripheral blood leukocytes.

#### **LEFT SHIFT:**

-Cell population has "shifted towards immature precursors " -Stab cell- nucleus looked like a Shepard's crook (Stab in German) -neutrophils counted left to right with bands being on the left



### **Differential Diagnosis:**

-Seizures -Toxic ingestions -metabolic abnormalities -inflammatory processes -Tissue damage -leukemia -Corticosteroid use -autoimmune diseases -chemotherapy

Myeloblast. Promyelocyte Myelocyte Metamyelocyte

#### LEFT SHIFT-acute infection

### Segmented Neutrophil



Even with normal WBC, Pts with moderate and high BANDEMIA on admission had significantly increased odds of having positive cultures, including blood cultures and of in hospital mortality

https://www.amjmed.com/article/S0002-9343(12)00450-0/abstract

BANDEMIA is highly predictive of a serious infection, suggesting that clinicians who do not appreciate the value of band counts may delay diagnosis or overlook severe infections.

Degree of bandemia, especially in the setting of concurrent tachycardia or fever, is associated with greater likelihood of negative clinical outcomes.

https://www.sciencedirect.com/science/article/abs/pii/ S0735675715001795

Bandemia is a superior indicator of infection relative to both WBC and Temp. -Abnormal temperature (hypo/hyperthermia) had a

sensitivity of 67% for culture proven bacteremia

-Abnormal WBC count had a sensitivity of 48% for culture proven bacteremia

-Bandemia had a sensitivity of 82% for culture proven bacteremia

-In patients with bacteremia and hypotension or septic shock, 33% had normal temperature and 21% had normal WBC count on initial evaluation

https://doi.org/10.1016/j.jemermed.2010.05.038

### **HISTORY:**

32-year-old male with no significant past medical history presents to the emergency department with abdominal pain. He states the pain began a few days ago in the right lower quadrant of the abdomen, and now feels as though it is spreading to the mid-abdomen.

### СВС

WBC 18.4 x 10 /mcl 61% neutrophils 11% lymphocytes 6% monocytes 2% eosinophils, 20% bands

HGB/HCT 14.6/44.6 g/dl PLT 356 x 10 /mcl





### Anemia

- Reduction in one or more of the major RBC measurements: Hgb, Hct or RBC count
- A low hemoglobin concentration and/or low hematocrit are the parameters most widely used to diagnose anemia.
- Caveats for normal:
  - Causes of lower values: intense physical activity, pregnancy, older age
  - Causes of higher values: smoking, hemoconcentration (hypovolemia), high altitude

#### Anemia evaluation in outpatients (nonpregnant adults)



This algorithm addresses anemia in healthy outpatients, which is often an incidental finding or may be identified when a CBC is performed to evaluate mild symptoms such as fatigue. It is not appropriate for individuals who are acutely ill with fever, bleeding, neurologic symptoms, or any severe cytopenia (hemoglobin <7 to 8 g/dL; platelet count <50,000/microL, absolute neutrophil count [ANC] <1000/microL). Consider the history, CBC, MCV, and reticulocyte count (if available) simultaneously. Refer to UpToDate for details of testing for specific causes of anemia.

### History:

70 year old female.

Symptoms of dyspnea on exertion, easy fatigability, and lassitude for past 2 to 3 months. Denied hemoptysis, GI, or vaginal bleeding. Claimed diet was good, but appetite varied. **Physical Exam:** 

Other than pallor, no significant physical findings were noted. Occult blood was negative.

### CBC

(with microscopic differential)
RBC 3.71 x 10 <sup>12</sup> /L
HGB 5.9 g/dL
HCT 20.9 %
MCV 56.2 fL
MCH 15.9 pg
MCHC 28.3 g/dL
RDW 20.2

WBC 5.9 x 10 <sup>9</sup> /I
N 82 %
L 13
M 1
E 4
B 0
PLT 383 x 10 <sup>9</sup> /L



### HISTORY:

37 year old male.

Lifelong history of a seizure disorder, treated since age two.

At a routine check with his neurologist, he complained of fatigue, exertional dyspnea, and lightheadedness over the past 2-3 months.

He appeared pale, but otherwise his physical exam was within normal limits. He was found to have a decreased hemoglobin, and was referred to Hematology Clinic.

#### CBC

(with microscopic differential)	WBC 6.2 x 10 <sup>9</sup> /L
	N 73 %
RBC 1.26 x 10 <sup>12</sup> /L	L 21
HGB 5.7 g/dL	M 1
HCT 16.3 %	E 4
MCV 130 fL	B 1
MCH 45.2 pg	PLT 219 x 10 <sup>9</sup> /L
MCHC 34.9 g/dL	
RDW 18.1	



## THROMBOCYTOPENIA

- Norm platelet count >150,000 per μl
- Thrombocytopenia <50,000 has 个 risk of major hemorrhage
- Causes: bone marrow depression, immune diseases and/or infections (particularly viral infections)
- Petechiae are often noted with a platelet count < 50,000</li>
- Decrease in # of platelets

- Most common cause of abnormal bleeding
- Causes:
  - Bone marrow damage (RT, chemo)
  - Idiopathic
  - Chemicals (benzene, insecticides)
  - Complication of viral diseases
  - Myelodysplastic syndrome
  - Drugs (thiazide diuretics, alcohol)
  - Malignancy (marrow infiltration)
    - Multiple myeloma
    - Acute leukemias
    - Lymphomas
    - Myeloproliferative disorders

21 yr old with no significant medical history complains of hematuria, bleeding gums while brushing his teeth.

On Physical exam, petechiae are noted in the oral cavity.

WBC 8,000 /mm3 Hematocrit 35% Platelet count 13,000 PT 13 s PTT 28 s

Normal smear VS. thrombocytopenia

### Hyponatremia

- Na < 120 mEq per L
- Most common electrolyte abnormality seen in medical practice
- Classified based on volume status
- In adults, the most common causes of hyponatremia are thiazide diuretic use, postoperative state with infusion of IV fluids, syndrome of inappropriate antidiuretic hormone secretion (SIADH), psychogenic polydipsia, exercise-associated hyponatremia, and unintentional water intoxication.



### Hypernatremia

- Na > 145 mEq per L
- Associated w/ increased m&m in inpatient setting
- Caused by net water loss
- Risk factors: impaired thirst mechanism, restricted access to water, AMS, intubated patients, infant, elderly



\*—The diagnosis of diabetes insipidus usually requires a combination of water deprivation and a trial of desmopressin. With water deprivation, patients with diabetes insipidus will have increased plasma osmolality but not urinary osmolality. In patients with central diabetes insipidus, urinary osmolality will increase by approximately 200 mOsm per kg after receiving desmopressin.<sup>35</sup>

#### Table 2. Differential Diagnosis and Treatment of Hypernatremia

Condition	Diagnosis	Treatment
Hypovolemic hypernatremia		
Body fluid loss (e.g., burns, sweating)	Clinical	Free water replacement
Diuretic use	Clinical	Stop diuretic
Gastrointestinal loss (e.g., vomiting, diarrhea, fistulas)	Clinical	Free water replacement
Heat injury	Elevated temperature, myoglobinuria, elevated creatinine level	Intravenous fluids, supportive care
Osmotic diuresis (e.g., hyperosmolar nonketotic coma, mannitol use, enteral feeding)	Elevated glucose level; sodium level often elevated after correction	Correct glucose level, stop causative agent
Post-obstruction	Clinical	Supportive care
Euvolemic hypernatremia		
Central diabetes insipidus	Clinical history of central nervous system insult; urinary concentration after administration of desmopressin	Treatment is rarely required unless thirst is impaired
Fever	Clinical	Treat underlying cause
Hyperventilation/mechanical ventilation	Clinical	Adjust ventilation
Hypodipsia	Clinical	Increase free water consumption
Medications (e.g., amphotericin, amino- glycosides, lithium, phenytoin [Dilantin])	Medication review	Stop causative medication
Nephrogenic diabetes insipidus	History of nephrotoxic medication use (amphotericin, demeclocycline [Declomycin], foscarnet, lithium, methoxyflurane), failure to concentrate urine after administration of desmopressin	Stop causative medication
Sickle cell disease	Hemoglobin electrophoresis	Treat underlying disease
Suprasellar and infrasellar tumors	Magnetic resonance imaging	Treat underlying disease
Hypervolemic hypernatremia		
Cushing syndrome	24-hour urinary cortisol and adrenocorticotropic hormone levels, dexamethasone suppression test	Treat underlying disease
Hemodialysis	Clinical history	Treat underlying disease
Hyperaldosteronism	History of hypertension and hypokalemia, plasma aldosterone-to-renin ratio, <sup>3</sup> history of hypertension and hypokalemia	Treatment usually not needed for hypernatremia
latrogenic (e.g., salt tablet or salt water ingestion, saline infusions, saline enemas, intravenous bicarbonate, enteral feedings)	Recent administration of hypertonic saline, enteral feedings, sodium bicarbonate infusion, or hypertonic dialysis	Stop causative medication, rapid free water replacemen

Information from references 3, 12, 33, and 34.

# Hypokalemia

- < 3.6 mmol/L
- Can results from decreased intake, increased translocation into cell, or most often due to increased losses due to GI tract or urine

#### Major causes of hypokalemia

Decreased potassium intake
Increased entry into cells
An elevation in extracellular pH
Increased availability of insulin
Elevated $\beta$ -adrenergic activity - stress or administration of beta agonists
Hypokalemic periodic paralysis
Marked increase in blood cell production
Hypothermia
Chloroquine intoxication
Increased gastrointestinal losses
Vomiting
Diarrhea
Tube drainage
Laxative abuse
Increased urinary losses
Diuretics
Primary mineralocorticoid excess
Loss of gastric secretions
Nonreabsorbable anions
Renal tubular acidosis
Hypomagnesemia
Amphotericin B
Salt-wasting nephropathies - including Bartter's or Gitelman's syndrome
Polyuria
Increased sweat losses
Dialysis
Plasmapheresis



# Hyperkalemia

- >5.2 mmol/L
- Commonly triggered by medications affecting K+ homeostasis, illness or dehydration
- The presence of ECG changes or rapid rise is potentially life-threatening
- IV calcium will stabilize cardiac membrane
- Insulin, bicarb will allow for intracellular shift of K+
- Kayexalate, dialysis allow for removal of K+

#### Major causes of hyperkalemia

Pseudohyperkalemia         Metabolic acidosis         Insulin deficiency, hyperglycemia, and hyperosmolality         Increased tissue catabolism         Beta blockers         Exercise         Hyperkalemic periodic paralysis         Other         Overdose of digitalis or related digitalis glycosides         Red cell transfusion         Succinylcholine         Arginine hydrochloride         Activators of ATP-dependent potassium channels (eg, calcineurin inhibitors, diazoxide, minoxidil, and some volatile anesthetics)         Reduced aldosterone secretion         Reduced response to aldosterone         Reduced distal sodium and water delivery         Effective arterial blood volume depletion         Acute and chronic kidney disease         Other         Selective impairment in potassium secretion         Gordon's syndrome         Uretojejunostomy	Increased potassium release from cells
Insulin deficiency, hyperglycemia, and hyperosmolality Increased tissue catabolism Beta blockers Exercise Hyperkalemic periodic paralysis Other Overdose of digitalis or related digitalis glycosides Red cell transfusion Succinylcholine Arginine hydrochloride Activators of ATP-dependent potassium channels (eg, calcineurin inhibitors, diazoxide, minoxidil, and some volatile anesthetics) <b>Reduced urinary potassium excretion</b> Reduced response to aldosterone Reduced distal sodium and water delivery Effective arterial blood volume depletion Acute and chronic kidney disease Other Urotaroiniument in potassium secretion	Pseudohyperkalemia
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Gordon's syndrome	Other
Uratarojojupostomy	Selective impairment in potassium secretion
Ureterojejunostomy	Gordon's syndrome
	Ureterojejunostomy

# URINALYSIS GROSS EXAMINATION: urine colors

PINK OR RED & CLOUDY	RBCs
PINK OR RED & CLEAR	Hemoglobin or myoglobin Porphyrins
DARK AMBER	Presence of conjugated bilirubin
BLACK	Melanin (metastatic malignant melanoma)
DARK YELLOW / VARIATIONS OF YELLOW	Hydration status
DARK ORANGE	Pyridium Rifampin

# URINALYSIS GROSS EXAMINATION: turbidity

- Can be seen in presence of infection or contamination
- Lipids
- RBCs

# URINALYSIS GROSS EXAMINATION: Urine odor

- Most commonly a result of production of ammonia by bacteria
- Presence of ketones can cause sweet or fruity odor
- Certain rare disorders can cause others:
  - Maple syrup urine disease- maple syrup odor
  - PKU- musty or mousy odor
- Foods: i.e. asparagus

# URINALYSIS DIPSTICK / CHEMICAL ANAYLSIS

CHARACTERISTIC	NORMAL	ABNORMAL	INTERPRETATION
SPECIFIC GRAVITY	1.005- 1.030		Low: hypovolemia, diabetes insipidus, High: hypervolemia, glucosuria, proteinuria, contrast, mannitol
рН	5.0-7.0		Aids in identification of acid-base disorders
HEME	(-)	(+)	RBCs, abnormal intravascular hemolysis, extensive burns, rhabdomyolysis, MI, Ascorbic acid (Vitamin C) can yield false-negative
LEUKOCYTE ESTERASE	(-)	(+)	Marker for presence of WBCs due to infection/inflammation Concentrated urine, proteinuria or glucosuria may produce false-negative
NITRITES	(-)	(+)	Bacteria
PROTEIN	(-)	(+)	Rhabdomyolysis, pregnancy, pre-eclampsia, eclampsia, multiple myeloma (Bence-jones proteins), Kidney disease, drugs/toxins,
GLUCOSE	(-)	(+)	Diabetes, pregnancy, renal glycosuria
KETONES	(-)	(+)	Inadequate carbohydrate intake, frequent vomiting/diarrhea, DKA,

# URINALYSIS: MICROSCOPIC EVALUATION

RBCS Normal: 0-2/hpf	UTI, stones, benign or malignant urinary tract neoplasms, trauma, prostatitis, glomerular or tubular disease, pyelonephritis, toxins, multisystem diseases i.e. SLE, blood dyscrasias i.e. Sickle cell disease, Rhabdomyolysis, menstrual contamination
	Infection or inflammation of genitourinary system
WBCS Normal: 0-3/hpf	<b>Pyelonephritis-</b> Present with moderate/heavy proteinuria, pyuria, WBC casts, and hematuria <b>Cystitis-</b> Present with pyuria, hematuria, and smaller amounts of protein. <u>No casts</u>
EPITHELIAL CELLS Clean: < 5/hpf	Squamous Epithelial Cells- most frequently seen, clinically least significant Transitional Epithelial Cells- large numbers typically after instrument procedures, suspect pathology if no h/o instrumentation i.e. catherization Renal Tubular Epithelial Cells- increased number indicates tubular necrosis or damage

ТҮРЕ	INTERPRETATION
RBC CASTS	Proliferative glomerulonephritis
WBC CASTS	Kidney infection, inflammation, or trauma
RENAL TUBULAR EPITHELIAL CASTS	Intrinsic renal tubular disease: ATN, acute interstitial nephritis, proliferative glomerulonephritis
GRANULAR CASTS	ATN, patients w/ischemic or toxic injury to tubular epithelial cells
HYALINE CASTS	Non-pathologic- strenuous exercise, dehydration, fever, emotional stress, diuretic therapy
	Pathologic- acute glomerulonephritis, pyelonephritis, chronic renal disease
WAXY CASTS	Severe urine stasis in renal tubules; often present in chronic renal failure
BROAD CASTS	Advanced CKD, ESRF

### Microscopic examination: crystals

- Calcium oxalate or calcium phosphate- form in alkaline urine, usually benign. Can be seen in ethylene glycol poisoning
- Magnesium ammonium phosphate- constituents of struvite stones. Occurs with increased ammonia production and elevated urine pH in the setting of UTI with a urease- producing organism i.e. Proteus or Klebsiella
- **Uric acid-** seen in diseases causing hyperuricemia (*gout*)
- Cystine- seen in cystinuria, an amino acid disorder

### Microscopic examination: microorganisms

- Bacteria- UTI, bacterial contamination
- Yeast
- **Parasites-** most common *Trichomonas vaginalis*

**History:** 24 F with no significant PMHx presents w/chief complaint of dysuria. She admits fever, vomiting, & flank pain.

Vitals: BP: 120/85 mmHg, HR: 115 bpm, RR: 22, Temperature: 103 F

**PE:** Non-toxic appearing, (+) R CVA tenderness, Abdomen is soft, non-tender and non-rigid without peritoneal signs.

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CBC: Elevated WBCs, otherwise WNL
CMP: WNL
Urinalysis: (+) Nitrites, (+) LE, (+) WBCs > 10/hpf, (+) RBCs
Urine Pregnancy: (-)
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