



Health Professionals

Things to never miss in the office

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Presenter Disclosure

• Faculty / Speaker's name: Brett Houston / Leonard Minuk

- Relationships with commercial interests:
 - Grants/Research Support: None
 - Speakers Bureau/Honoraria: None
 - Consulting Fees: None
 - Other: None





Learning Objectives

1. To review the red flags, diagnosis and approach to thrombotic thrombocytopenic purpura (TTP)

2. To review the common presentations of acute leukemia, and the immediate action plan





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- 44F with newly identified bruising / petechiae
- Vaginal bleeding since insertion of IUD one month prior; otherwise no bleeding

- Medical history: PCOS
- Medications: none





WBC (x10 ⁹ /L)	15
Differential (x10e9)	Neutrophils 11.9 Lymphocytes 2.73
Hemoglobin (g/L)	68
MCV (fL)	98
Reticulocyte count	330
Platelets (x10 ⁹ /L)	11

- Creatinine 93
- ALT 10
- Total bilirubin 42
- Indirect bilirubin 33
- LDH 975
- Haptoglobin < 0.01

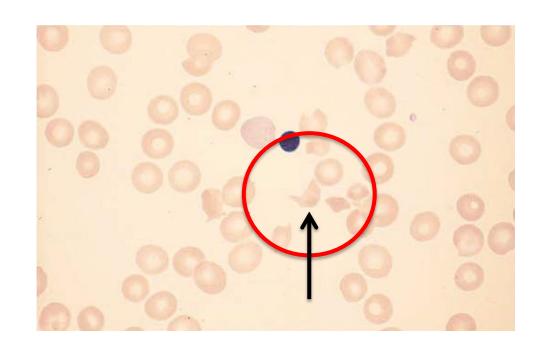




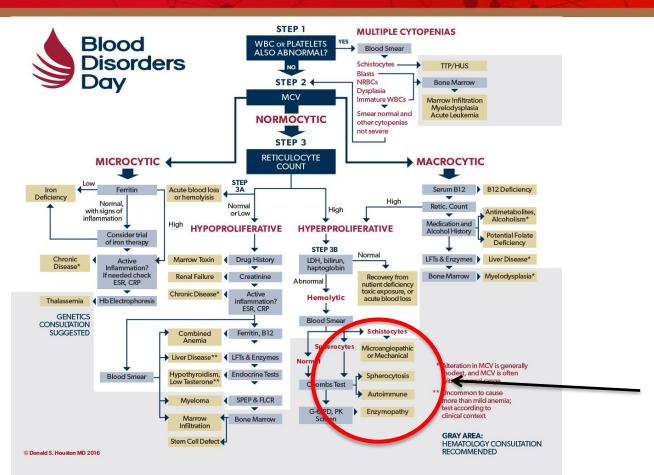
Peripheral film: increased polychromasia; 4-10 schistocytes / hpf

Schistocytes

- = Fragments
- = Microangiopathy



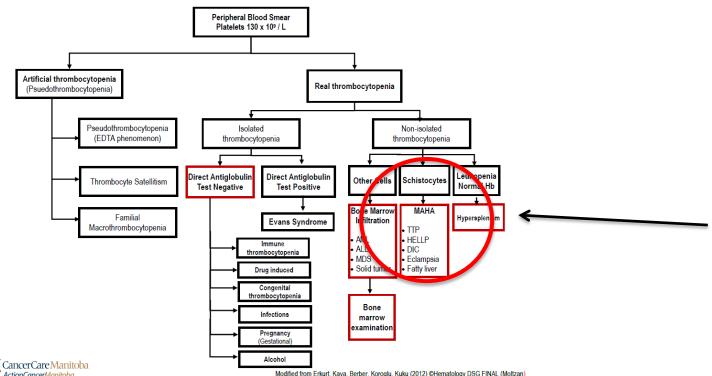








Work-Up of THROMBOCYTOPENIA



ActionCancer/Manitoba

Modified from Erkurt, Kaya, Berber, Koroglu, Kuku (2012) ©Hematology DSG FINAL (Moltzan)

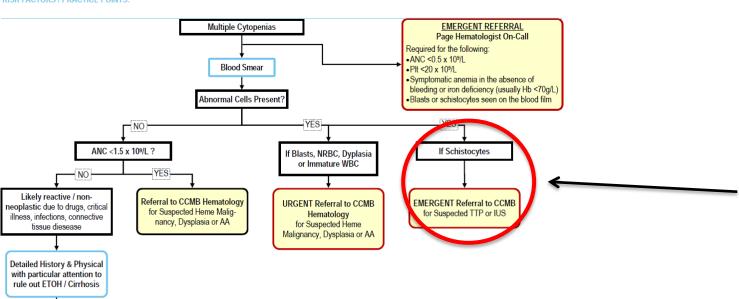
Pathways are subject to clinical judgment and actual practice patterns may not always follow the proposed steps in this pathway.





Work-up of PANCYTOPENIA

RISK FACTORS / PRACTICE POINTS



Referral to CCMB Hematology if persistent / symptomatic cytopenia





TTP - Clinical manifestations

- Classic pentad:
 - Fever
 - Thrombocytopenia
 - Microangiopathic hemolytic anemia
 - Neurologic symptoms
 - Renal insufficiency

Present in <10% of patients at diagnosis





TTP – Clinical manifestations

- *Thrombocytopenia (usually <30)
- *Thrombotic microangiopathy
- Neurologic symptoms (60%)
- Abdominal pain (mesenteric ischemia) (35%)
- Cardiac ischemia (25%)
- Renal failure (10-25%)





TTP - Investigations

- CBC
- Reticulocyte count
- Peripheral blood film
- Creatinine
- Bilirubin, LDH, haptoglobin





TTP – Differential diagnosis

- Malignant hypertension
- Pre-eclampsia / HELLP
- Hemolytic uremic syndrome
- Disseminated intravascular coagulation

- → Blood pressure
- $\rightarrow \beta$ -hcg
- → Complement testing
- → INR, aPTT, fibrinogen, d-dimer





TTP - Management

- Hematology referral (STAT → call)
- Plasma exchange

- Prior to plasma exchange, mortality in TTP was 90%
- With plasma exchange, survival in TTP is now 90%





Case #1 Revisited

- Despite feeling entirely well, 1 L of FFP was transfused and she sent to Winnipeg by ambulance
- Central line inserted in ER upon arrival
- Plasma exchange initiated within 4 hours
 - Hemoglobin, reticulocyte count, platelets and LDH normalized over the next few days
 - Currently under observation





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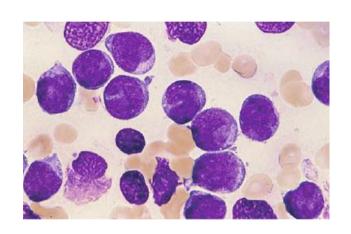
- 59F seen in clinic with increased bruising
- Otherwise well

- Past medical history: none
- Medications: none





WBC (x10e9)	69
Differential (x10e9)	Neutrophils 0.7 Lymphocytes 17.9 Monocytes 1.4 Myelocytes 0.69 Blasts 48.3
Hemoglobin (g/L)	109
MCV (fL)	84
Platelets (x10e9)	27

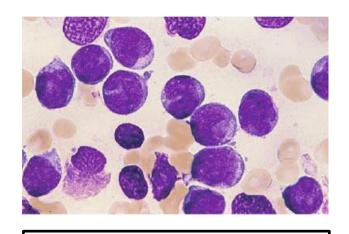


^{*} Prior CBC normal





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IMPRESSION:

Leukocytosis (with blasts)

Pancytopenia

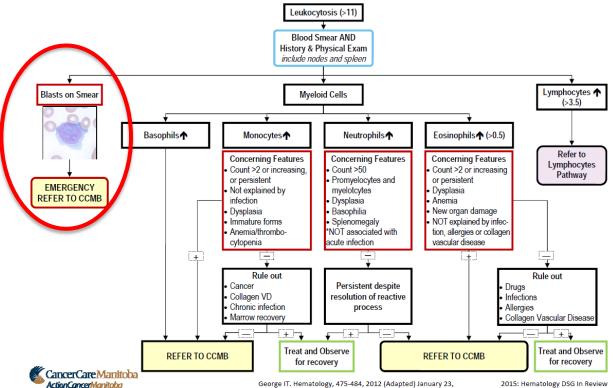
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Work-Up of LEUKOCYTOSIS

RISK FACTORS: add info herePRACTICE POINTS: add info here







- 24M seen in clinic with progressive fatigue, night sweats and weight loss
- Recently antibiotics for ?pneumonia
- No infectious source, no bleeding

- Past medical history: none
- Medications: none





WBC (x10 ⁹ /L)	1.9
Differential (x10 ⁹ /L)	Neutrophils 1.19 Lymphocytes 0.71 Monocytes 0.02 Blasts 0.02
Hemoglobin (g/L)	97
MCV (fL)	94
Platelets (x10 ⁹ /L)	153

^{*} Prior CBC normal





WBC (x10 ⁹ /L)	1.9
Differential (x10 ⁹ /L)	Neutrophils 1.19 Lymphocytes 0.71 Monocytes 0.02 Blasts 0.02
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IMPRESSION:

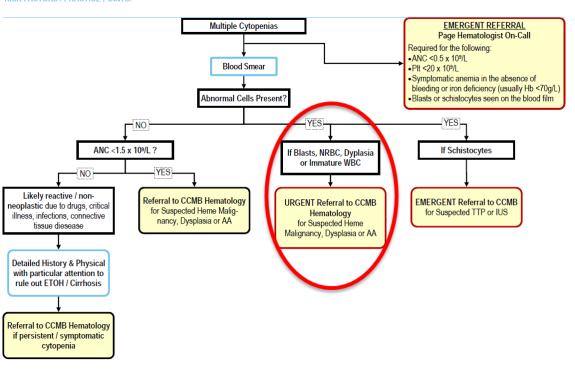
Pancytopenia





Work-up of PANCYTOPENIA

RISK FACTORS / PRACTICE POINTS:







Acute leukemia – Clinical manifestations

- Constitutional symptoms
- Fatigue, malaise
- Anemia (pallor, heart failure)
- Thrombocytopenia (bleeding)
- Leukopenia / leukocytosis (infection)





Acute leukemia - Diagnosis

- Leukocytosis (blasts) OR pancytopenia
 - Peripheral blood OR bone marrow blasts >20%

- Delayed treatment is associated with reduced survival
- Early mortality related to bleeding and infection





Diagnosis – Take home points

- Present with constitutional symptoms, infections, bleeding
- CBC can show leukocytosis OR pancytopenia
- Differential and peripheral film are very informative

• New blasts are always bad; call HEMATOLOGY ON CALL





Acute leukemia - Complications

- White blood cells (infection, leukostasis)
- Hemoglobin (symptomatic anemia, CHF)
- Platelets (bleeding)

- DIC (bleeding, thrombosis)
- Tumor lysis syndrome





Acute leukemia - Assessment

- History & physical
 - Focal infection
 - Bleeding (intracranial, GI, mucocutaneous)
 - Thrombosis (DVT, PE)
 - Organ dysfunction (head to toe)
 - Leukostasis





Acute leukemia - Assessment

- Laboratory tests
 - CBC, differential, blood film
 - Electrolytes, Ca/Mg/PO4, albumin
 - Creatinine
 - Liver enzymes
 - LDH, uric acid
 - DIC screen (INR, aPTT, fibrinogen, d-dimer)





Case #2 / #3 Revisited

- Both patients were transferred to HSC leukemia service within 24 hours
- Baseline investigations (bone marrow, MUGA)
- Started induction chemotherapy





Take home messages

- TTP and acute leukemia are medical emergencies with acute life-threatening complications
- Maintain a high index of suspicion
 - TTP anemia / thrombocytopenia
 - Acute leukemia leukocytosis / pancytopenia
- If you suspect these, please call hematology on-call (at any hour!)





