



FOR  
Health Professionals

# Parsing Out Platelets

How I Approach Thrombocytopenia

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# Disclosures

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# Objectives

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1. Be aware of conditions commonly associated with thrombocytopenia in clinical medicine
2. Use an algorithm to guide investigation of a patient with thrombocytopenia
3. Be aware when urgent hematologic referral is indicated for a patient with thrombocytopenia

# Referral to Hematology

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Dear Dr.

Please assess this 35 year old female with thrombocytopenia, incidentally noted on bloodwork. She is otherwise asymptomatic.

Results:

|   |           |
|---|-----------|
| WBC $6.0 \times 10^9/L$ , normal differential | (4.5 -10) |
| Hb 135 g/L, normal indices                    | (120-160) |
| Platelet $40 \times 10^9/L$                   | (130-400) |

# Definition

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- Thrombocytopenia- platelet count that is less than normal
- Normal is 130 to 400 x 10<sup>9</sup>/L
- Clinical definition is a platelet count of less than 100 x 10<sup>9</sup>/L
- Treatment usually not indicated until platelet count is less than 30 x 10<sup>9</sup>/L
- Very important that the blood film is examined to ensure it is true thrombocytopenia

# Pseudothrombocytopenia

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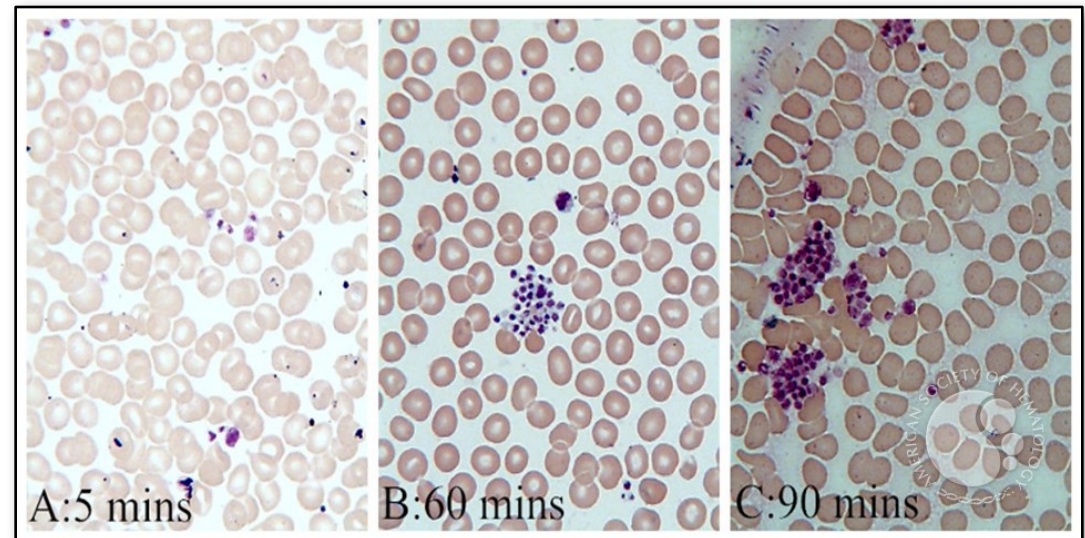
- Platelets are recorded as falsely low by the hematology analyzer
- Causes include:
  - EDTA-induced platelet aggregation
  - Platelet satellitism
  - Familial Macrothrombocytopenia



# EDTA-Induced Platelet Aggregation

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- Time-dependent
- *In vitro*
- Antibody crosslink



# Platelet Satellitism

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- Platelets adhere to WBC leaving the automated platelet count low
- Also the result of EDTA

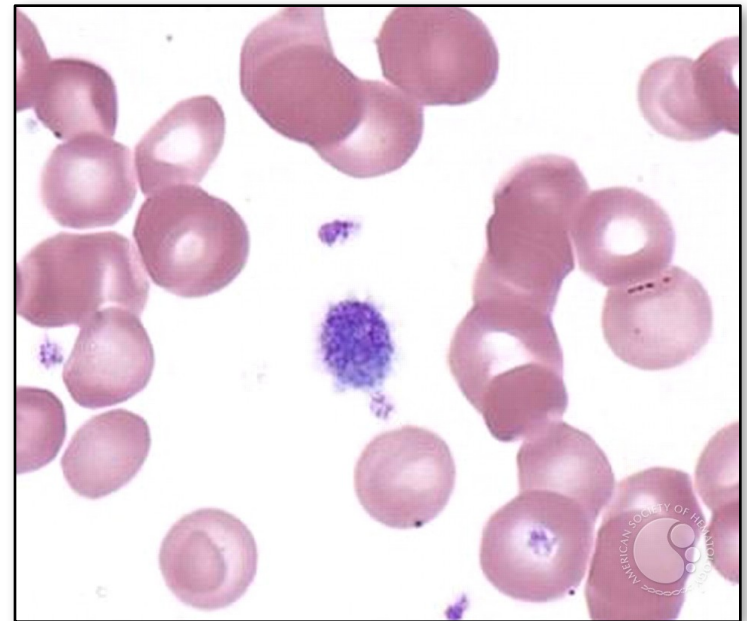




# Macrothrombocytopenia

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- Most are familial
- Large platelets counted as red cells by the hematology analyzer



# Causes of True Thrombocytopenia

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- Decreased bone marrow production of platelets
- Sequestration of platelets
- Increased destruction of platelets
  - Immune
  - Non-immune

# Decreased Bone Marrow Production

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- Rare cause of isolated thrombocytopenia- alcohol
- Usually associated with leukopenia, anemia, and/or abnormal bone marrow pathology- pancytopenia algorithm
- Referral to hematology consideration of bone marrow examination

# Increased Sequestration of Platelets

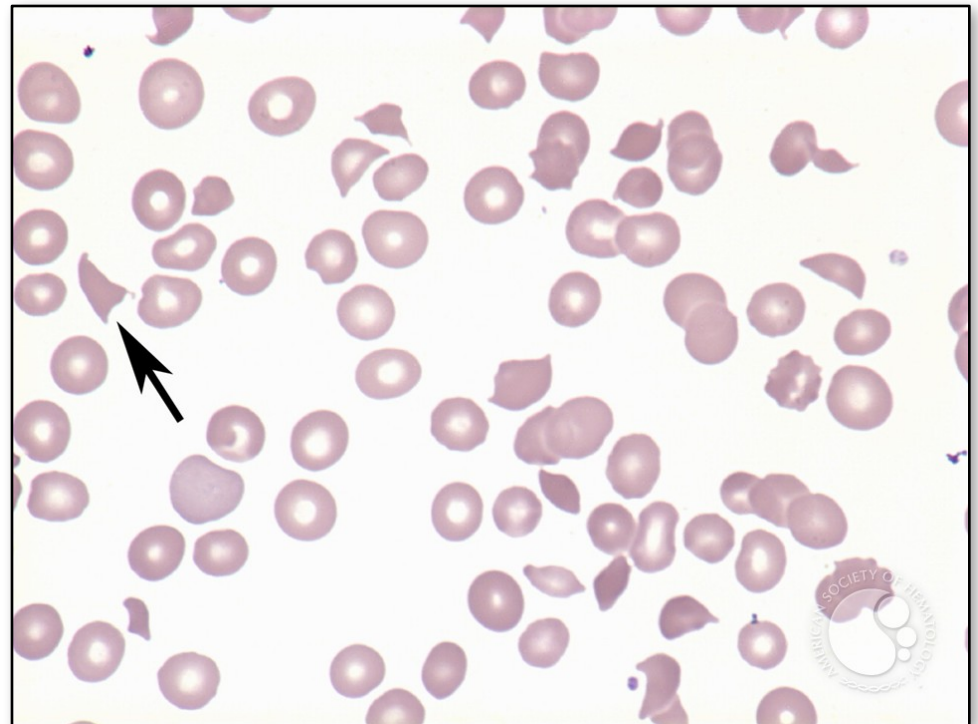
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- Rare cause of isolated thrombocytopenia
- Usually associated with leukopenia (and normal hemoglobin)
- Causes are those of hypersplenism
  - Portal hypertension
  - Other causes of splenomegaly
- Need imaging of liver/spleen

## Non-immune Destruction of Platelets

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- Causes of microangiopathic hemolytic anemia
- TTP, HUS, DIC
- Urgent hematology consult

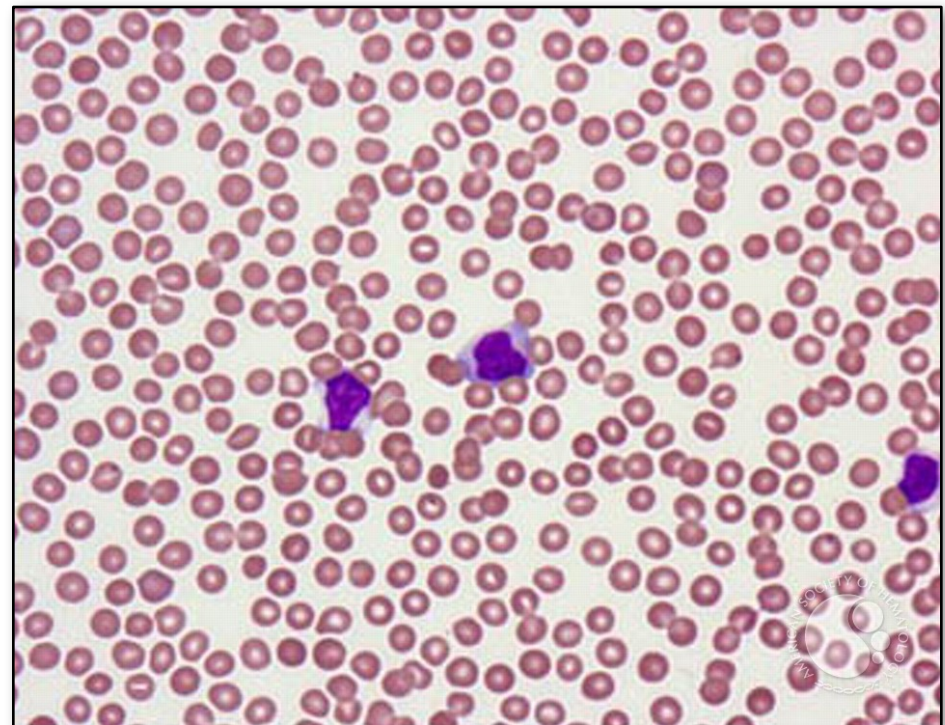




# Immune Destruction of Platelets

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- Primary
- Secondary
  - Autoimmune diseases
  - Lymphoproliferative Diseases
  - Infections
    - HIV
    - Hepatitis B and C
    - EBV, CMV



# Workup of Isolated Thrombocytopenia

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- **History:** viral illness, autoimmune disease, LPD, bleeding
- **Medications:** amiodarone, beta lactams, carbamazepine, GpIIb/IIIa inhibitors, heparin, ibuprofen, mirtazepine, phenytoin, rifampin, quinidine, quinine, TMP/SMX, vancomycin- need to stop offending drugs
- **Physical Exam:** hepatosplenomegaly, lymphadenopathy, autoimmune disease, petechiae/bleeding

# Workup of Isolated Thrombocytopenia

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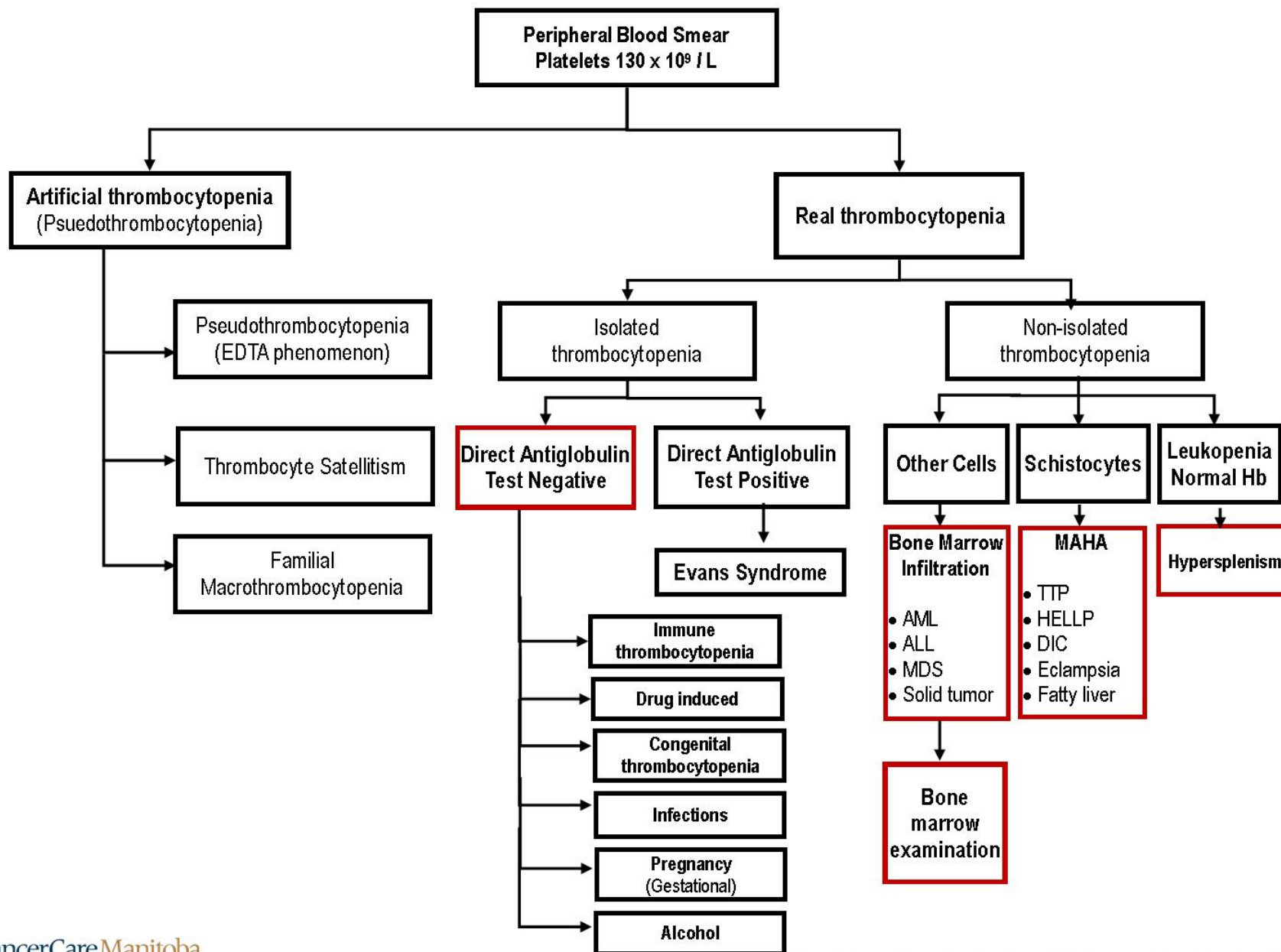
- If results do not make sense, repeat CBC first
- CBC, retic count, INR, direct antiglobulin test, ABO and Rh
- Review of peripheral blood smear
- Renal function, liver enzymes, (ANA/autoimmune panel)
- HIV, Hepatitis B and Hepatitis C serology (EBV, CMV)

# Referral of Immune Thrombocytopenia

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- Emergent referral if platelet count less than  $30 \times 10^9/L$  and/or platelet count less than  $50 \times 10^9/L$  with bleeding
- Urgent referral if platelet count between  $30$  and  $50 \times 10^9/L$
- Routine referral if platelet count between  $50$  and  $100 \times 10^9/L$
- Referral not required if platelet count greater than  $100 \times 10^9/L$

# Work-Up of THROMBOCYTOPENIA





# Take Home Messages

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- Peripheral blood film review rules out pseudothrombocytopenia and the presence of other abnormal cells
- Most cases of isolated thrombocytopenia are immune
- Need to think about medications as the cause but relatively few medications cause significant thrombocytopenia
- Urgent referral required if platelet count less than  $30 \times 10^9/L$  and/or platelet count less than  $50 \times 10^9/L$  with bleeding

# References

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# Questions?

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**2. Which of the following drugs is not likely to cause thrombocytopenia?**

- A. Quinine
- B. Acetaminophen
- C. Ibuprofen
- D. Amiodarone
- E. Heparin

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