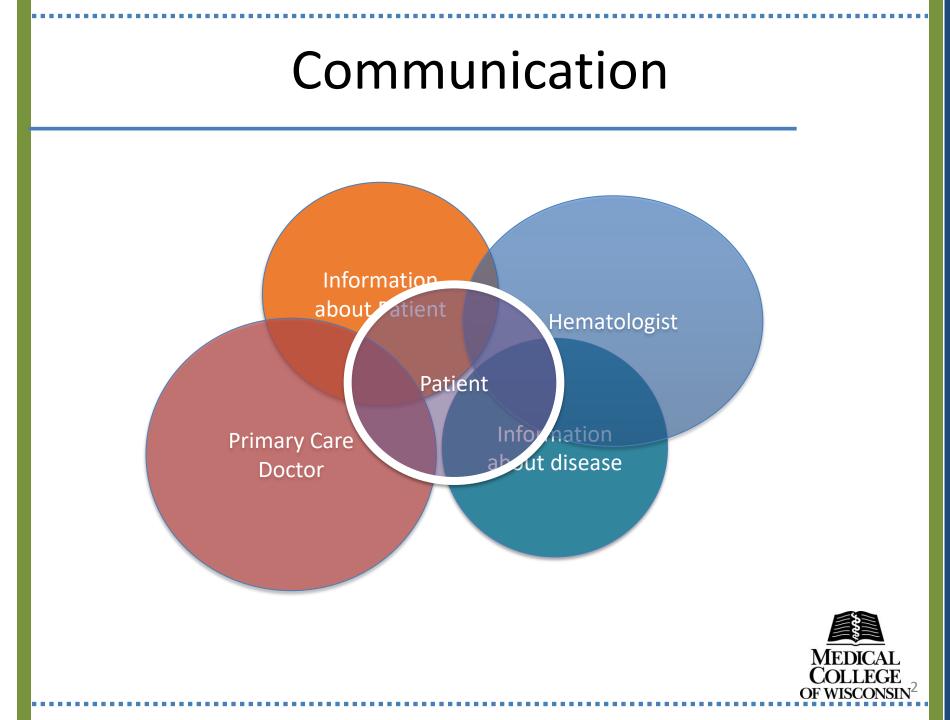


#### What's Routine? General Medical Care in Patients with MPNs

Laura C. Michaelis, MD Joyce Niblack Memorial Conference 2017



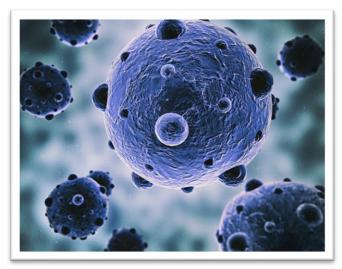


## Today's Goals

- Review unique risks faced by MPN patients
- Review targets for optimizing
  - Optimizing cardiac health
  - Minimizing infection risks
  - Limiting other cancers
- Discuss strategies for increasing provider-toprovider communication



### **Overall Risks**



#### Risk of Infections: Bacterial, Viral,

Fungal





<sup>1</sup>Hultcrantz M et al., JCO Vol 33; No 20; July 2015



#### **Risk of Heart Attack**

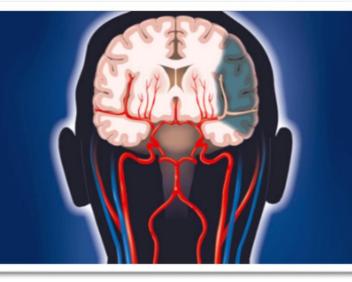


## **Overall Risks**



Risk of Blood clots in the legs, lung, liver and skull veins

# Risk for arterial clots that cause stroke





## Control what you can





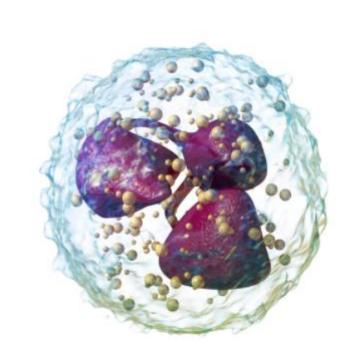
## Infections

- Large studies show increased rate of fatal infections
- Why might that be?
  - Disease  $\rightarrow$  irregularities in immune system
  - Low blood counts: Neutropenia
  - Splenic dysfunction
  - Treatment with JAK inhibitors





#### Neutropenia



- Determination
  - Blood smear examined
  - Percentage neutrophils among total white blood cells calculated
- Treatment
  - If asymptomatic, can watch
  - If repeated infections, then prophylaxis recommended

## **Preventing Infection**

• Prophylaxis only in selected patients

NCCN: "antibiotic prophylaxis for recurrent infections is recommended"

- Bacterial: Flouroquinolone antibiotic
- Viral: Acyclovir or other
- Fungal: Fluconazole, Posaconazole, Voriconazole
- Vaccination
  - Influenza vaccine to patient and family
  - If immunocompromised (i.e. no spleen, on JAK inhibitor)
    - Avoid live-attenuated virus vaccination, i.e. no Zoster vaccination
    - 7-days contact avoidance from family members

### Infections related to treatment

- Most common:
  - Bacterial infections of respiratory and urogenital tract
  - Frequency decreases with length on therapy
  - Severe infections no more common than patients on placebo
- Likely increased risk for rare infections
  - For example, tuberculosis reactivation
  - Shingles (Herpes zoster)



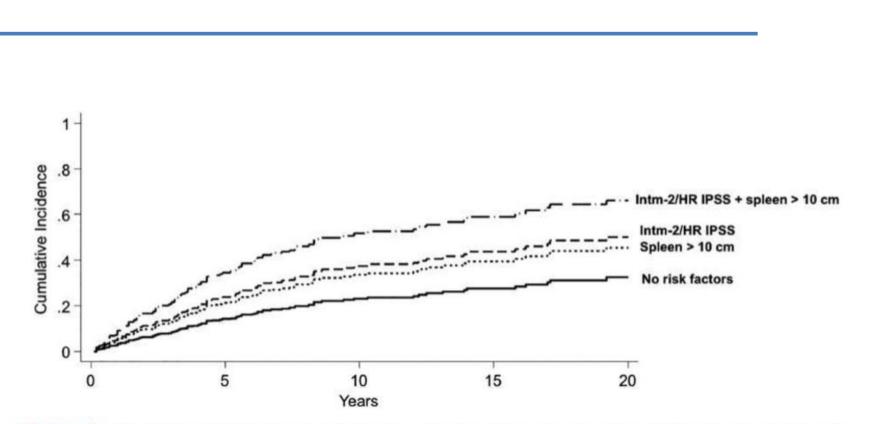


Figure 2. Cumulative incidence of infectious complications according to IPSS and splenomegaly.



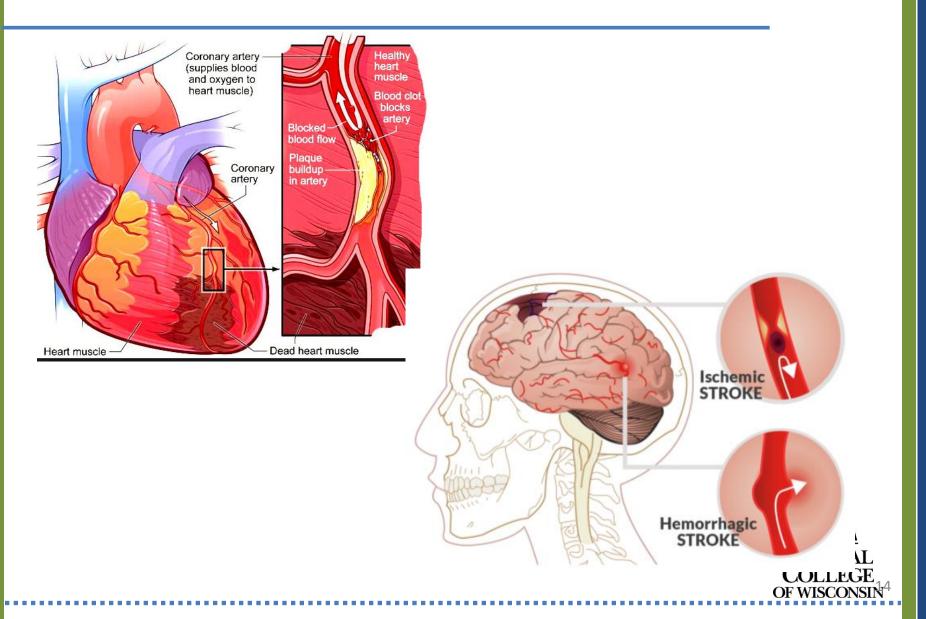
Polverelli et al., AJH 92: 37-41; 2017

## Care while on therapy

- Avoid starting therapy while actively infected
- Ensure no tuberculosis risk prior to starting treatment
- Watch for signs/symptoms of zoster
- Rule out Hep B prior to starting therapy
- No routine recommendations for prophylaxis against infections

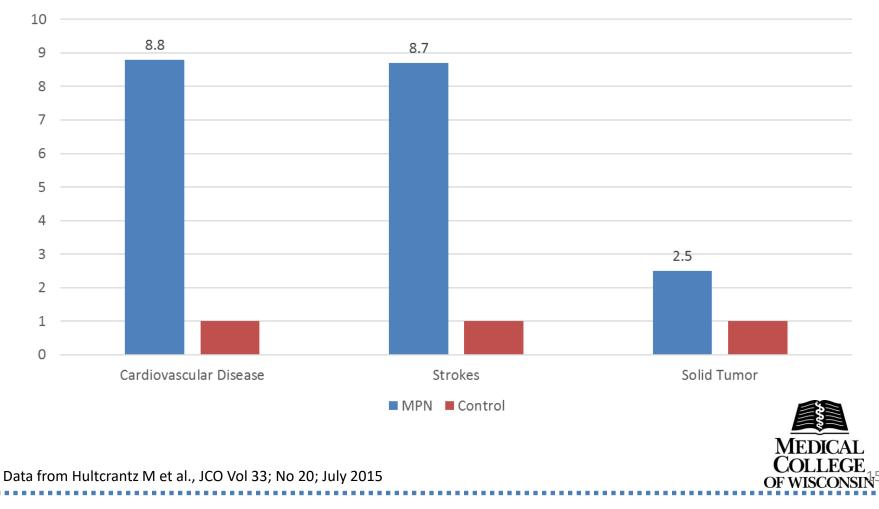


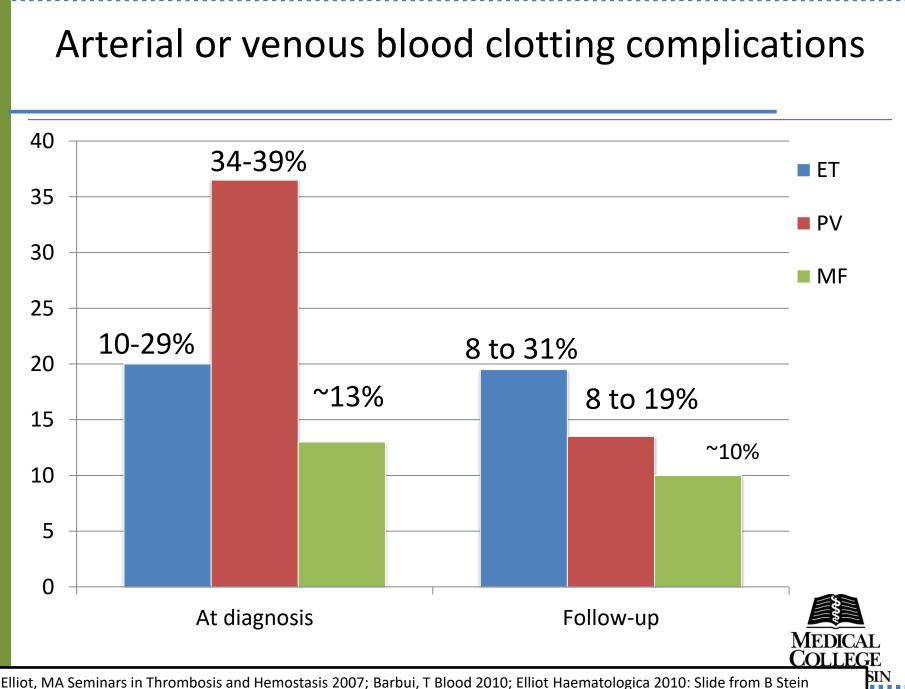
#### **Arterial Clotting Events**



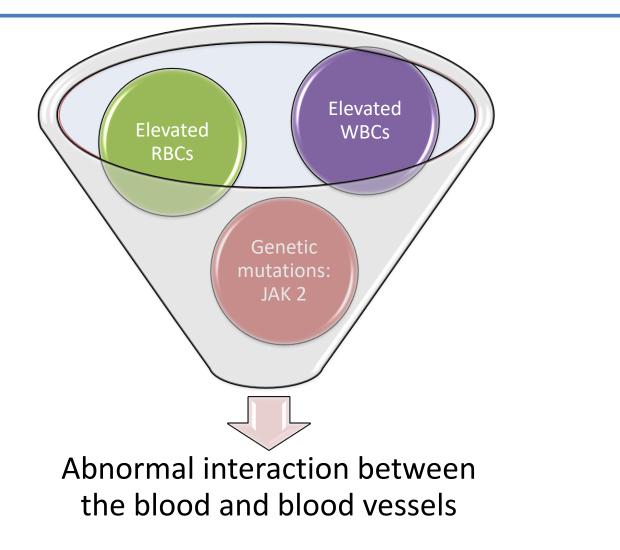
## Younger Individuals

Cause-Specific Mortality Rates when Aged 18-49 at diagnosis





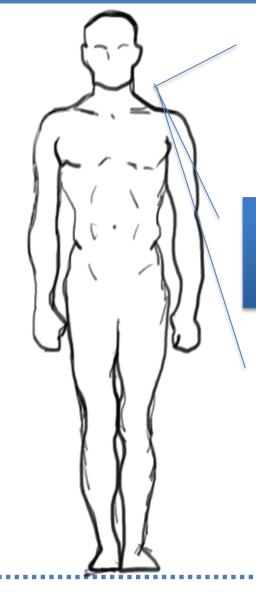
#### Why more common?





## Cardiac Risk factors

- Age
- Smoking
- Total Cholesterol
- "Good cholesterol"
- Diabetes
- Blood pressure
- Activity level
- Weight



60 yo male; no diabetes, smoker, total cholesterol of 210; good cholesterol of 30; blood pressure of 140/70 mm/hg

Risk of heart attack in next 10 years: 23%

> If he is not a smoker, takes medication or diets to reduce cholesterol by 30 points and lowers his BP to 130/70: Risk drops to 13%

### **Blood Pressure Targets**

Blood Pressure Category	Systolic mm Hg (upper #)		Diastolic mm Hg (lower #
Normal	less than 120	and	less than 80
Prehypertension	120 – 139	or	80 – 89
High Blood Pressure (Hypertension) Stage 1	140 – 159	or	90 – 99
High Blood Pressure (Hypertension) Stage 2	160 or higher	or	100 or higher
Hypertensive Crisis Emergency care needed)	Higher than 180	or	Higher than 11

Guidelines: American Heart Association



## Targets

- Complete abstinence from smoking
- Heart-healthy eating patterns
  - American Heart Association
    - Use up the calories you take in
    - Choose high-nutrient foods: Fruits and veges, whole grains, low-fat dairy, poultry and fish without the skin, nuts and legumes, non-tropical vegetable oils
- Keep your body-mass index between 18-24.9
  AHA has a calculator



### Cholesterol?

#### **Metrics**

- Total cholesterol = HDL + LDL + 20 percent of your triglyceride level.
- HDL (good) cholesterol: High is better
- LDL (bad) cholesterol: Low is better
- Targets are complicated depend on your age, prior history, presence or absence of diabetes
- Frequency of testing: discuss with PCP

#### Interventions

- Dietary interventions
  - Fish, Omega-3 fatty acids
  - Oatmeal, oat bran, high soluble fiber
- Exercise
  - Boost HDL, decreases danger of LDL
- "Statins"
  - Lower LDL
- Nicotinic Acid "Niacin"
  - Raise HDL
- Fibrates
  - Lower triglycerides

#### Exercise

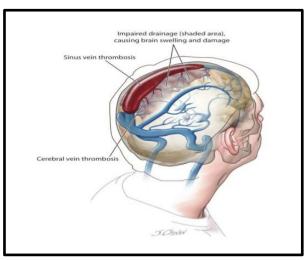
- Aerobic Exercise
  - Lowers cholesterol and BP
  - Increases endurance
  - Lower resting heart rate
  - Weight loss and maintenance
  - Stress relief
  - Improved sleep
- 30 min; 5-7 days a week

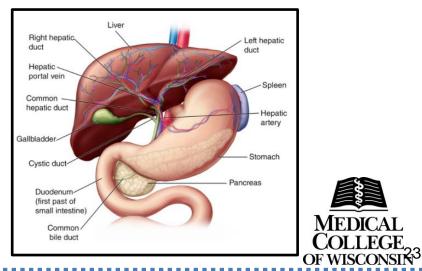


#### What about Venous Clots?

- Most common locations
  - Legs (deep venous thrombosis "DVT")
  - Lungs (pulmonary embolism "PE")
- Can also occur in

abdominal vasculature and brain drainage





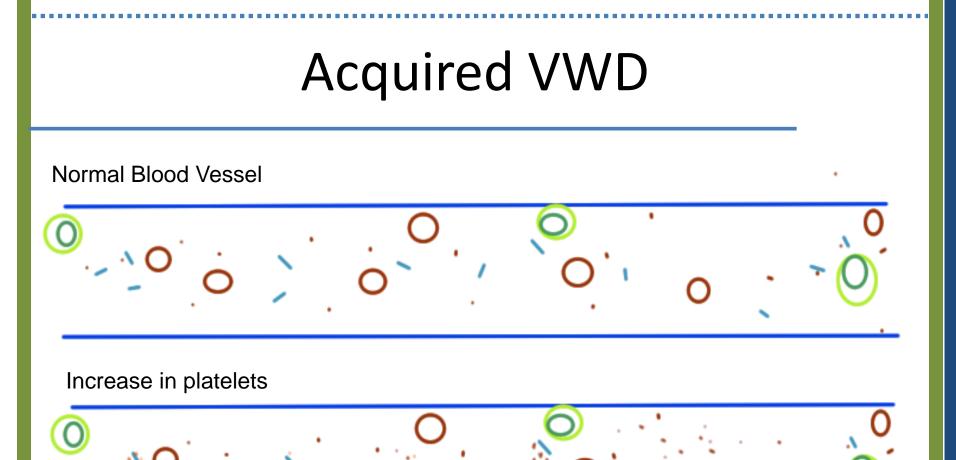
## Treatment of PE/VTE or Other

Protection from Blood Clots: "Prophylaxis"	Initial Treatment	Extended Treatment		
MPN Disease Control				
Around the time of surgery	Blood Thinning medications for at least 3-6 months	Consider extended treatment for abdominal clots, SVT, life threatening clots, second clot		
	Usually pause aspirin while on blood thinner unless risk >benefit			

## What about bleeding?

- Can occur in patients with very high platelet count, i.e. over 1.5 million/uL
- Called acquired Von Willebrand's disease
- Treatment
  - Avoid aspirin
  - Lower platelet count







## Modifying Surgical Risk

Planning	Assessment by hematologist Optimize blood counts Especially platelets if splenectomy planned
Preoperative	Discontinue ASA
Postoperative	Anticoagulation – LMWH Clinical vigilance re hemorrhage US of abdominal veins

OF WISCONSIN

## Other cancers?

- Slightly increased risk for solid tumors
  - Italian Study
    - Roughly 4 times as likely to acquire lymphoma-type diseases
  - Danish Study
    - Incidence of solid tumors slightly higher in patients with ET, PV and CML
  - Swedish Study
    - Increased incidence of thyroid and parathyroid cancers and skin cancers
  - MD Anderson Study
    - Statistically significant increase in solid tumors



### **Preventative Health**

- Everybody
  - No tobacco
  - Good sun protection;
  - Regular skin evaluations
  - Healthy diet; Weight control
  - Limit alcohol
- Age-specific (start date depends on family history)
  - Mammogram, colonoscopies, PAP smears, prostate exams
  - Low-dose CT scans if smoker



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## **Provider Collaboration**

- Communication between providers critical
- Lots of ways you can facilitate
  - Keep your own records to bring back and forth
    - i.e. information about labs, clinical trials
  - Ask about sharing EMR
  - Encourage communication
  - Work with ancillary staff
  - Engage in your care



Thank You for all you've done to advance this field

Happy to take any questions

THE REAL PROPERTY.