

Syncope: What To Do When the Lights Go Out?

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Disclosure

I have no financial interests or relationships to disclose.

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Learning Objectives

Upon completion, the participant will be able to:

1. Conduct an appropriate diagnostic evaluation for the patient with syncope.
2. Recognize potentially life-threatening EKG's that are associated with syncope.
3. Describe patients with syncope that are "high risk" for subsequent fatal arrhythmia events.
4. Be familiar with the 2017 AHA/ACC guideline for syncope.



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Your Patient *"Passed Out"...* Your Differential Diagnosis Is:

CV causes

- **Autonomic**
 - Carotid sinus syncope
 - Cough
 - Defecation
 - Excessive vagal tone (athletes, adolescents)
 - Micturition
 - Postprandial
 - Sneeze
 - Swallow
 - Valsalva

Non-CV causes

- **Metabolic**
 - Alcoholism
 - Carbon monoxide
 - Drug-induced
 - Hyperventilation
 - Hypoglycemia
 - Hypothyroid
 - Hypoxia/asphyxiation
 - pheochromocytoma

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Your Patient *"Passed Out"...* Your Differential Diagnosis Is:

CV causes

- **Orthostatic hypotension**
 - Adrenal insufficiency
 - Autonomic insufficiency
 - CNS dz, alcoholism, DM
 - Dehydration
 - Drugs
 - Antihypertensives
 - Drugs of abuse
 - Vasodilators
 - CNS drugs
 - Hemorrhage
 - idiopathic

Non-CV causes

- **CNS**
 - Basilar artery migraine
 - Narcolepsy
 - Seizure
 - Subarachnoid hemorrhage
 - Subclavian steal
 - Vertebrobasilar insufficiency
 - Increased intracranial pressure

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Your Patient *"Passed Out"...* Your Differential Diagnosis Is:

CV causes

- **Obstructive lesions**
 - Aortic dissection
 - Aortic, mitral or pulmonary stenosis
 - Atrial myxoma
 - Cardiac tamponade
 - Congenital heart disease
 - Left ventricular dysfunction
 - Pulmonary embolism
 - Pulmonary hypertension

Non-CV causes

- **Psychiatric**
 - Anxiety disorder
 - Breath-holding spells
 - Conversion reaction
 - Drug-induced
 - Anticonvulsants
 - Antihistamines
 - Antiparkinson drugs
 - Cholinesterase inhibitors
 - TCA
 - Pseudoseizure

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Your Patient *"Passed Out"...* Your Differential Diagnosis Is:

CV causes

- **Dysrhythmias**
 - Bradyarrhythmia
 - Heart blocks, Sick sinus syndrome,
 - Meds: Beta and Ca-Channel blockers, digoxin, cholinesterase inhibitors
 - Tachyarrhythmias
 - Supraventricular and ventricular
 - Torsades de pointes
 - Drug induced
 - TCA, digoxin, antiarrhythmics
 - Prolonged Q-T syndrome
 - Congenital
 - Drug-induced
 - Myocardial infarction
 - Pacemaker failure, ICD malfunction

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What Do You Really Want to Know?

- Dangerous etiology vs. Benign etiology

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*“Those Who Suffer from Frequent and Strong Faints
Without Any Manifest Cause Die Suddenly”*

Hippocrates (460 - 375 BC)



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What Do You Really Want to Know?

- Dangerous etiology vs. Benign etiology
- Medical-legal peril vs. None

Admit vs. discharge to home?

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What Do You Really Want to Know?

- Dangerous etiology vs. Benign etiology
- Medical-legal peril vs. None

Admit vs. discharge to home?

What work-up is needed?

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Syncope

- Definition - *very useful*
- Pathophysiology - *very useful*
- H & P elements - *that are useful*
- Tests/
Clinical Policies - *that might be useful*
- EKG's - *that are worrisome!*
- The outpatient evaluation

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Syncope - Definition

- “A brief loss of consciousness associated with an inability to maintain postural tone that spontaneously and completely resolves without medical intervention”

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Syncope - Definition

- “A **brief** loss of consciousness associated with an inability to maintain postural tone that spontaneously and completely resolves without medical intervention”
- **Brief:** not asleep/intoxicated, not post-ictal

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Syncope or Too Much to Drink (or Dead)?



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Syncope - Definition

- “A **brief** loss of consciousness associated with an inability to maintain postural tone that **spontaneously** and completely resolves without medical intervention”
 - **Brief:** not asleep/intoxicated, not post-ictal
 - **Spontaneous:** no intervention needed....
so rarely=> “hypoglycemia”

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Syncope - Definition

- “A **brief** loss of consciousness associated with an inability to maintain postural tone that **spontaneously** and **completely** resolves without medical intervention”
 - Brief: not asleep/intoxicated, not post-ictal
 - Spontaneous: no intervention needed....
so rarely=> “hypoglycemia”
 - **Completely:** no neurologic deficit, return to baseline

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Syncope- Pathophysiology

- Global cerebral hypoperfusion

Forget the “TIA” diagnosis/eval

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Syncope

- Definition - *very useful*
- Pathophysiology - *very useful*
- H & P elements - *that are useful*
- Tests/
Clinical Policies - *that might be useful*
- EKG's - *that are worrisome!*
- The outpatient evaluation

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Syncope - History

High - risk

- Older age
- (+) CV diagnosis
- (+) CHF
- (+) Family Hx
- No prodrome
- Supine position
- Assoc. with exertion

(think structural outflow obstruction)

Lower - risk

- younger age
- (-) CV diagnosis
- (-) CHF
- (-) Family Hx
- (+) prodrome
- (-) exertion

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Syncope - History Age Does Matter!!!

- Methods: prospective study, ED pts (+) syncope
- Results: 477 patients, 97% f/u @14 days
 - 80 (18%) patients with “serious event”

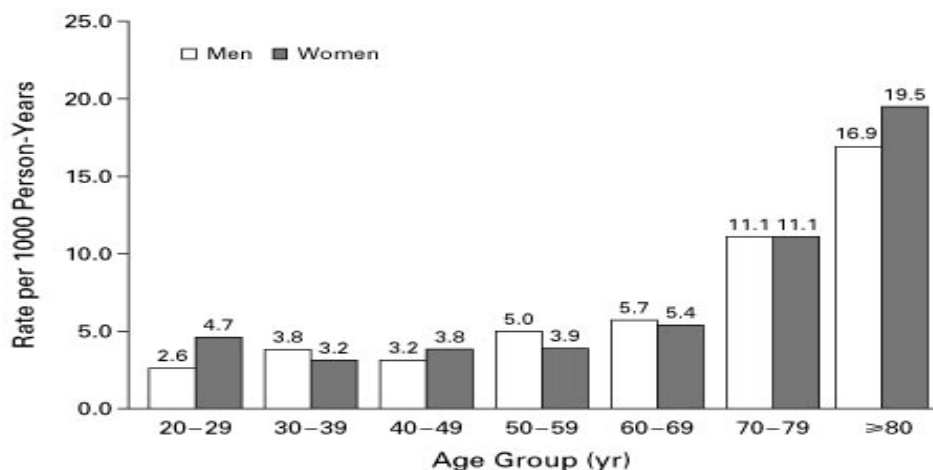
<u>AGE</u>	<u># (%) serious outcome</u>	<u>Odds Ratio</u>
< 40	4/141 (3%)	1.0
40-59	15/112 (13%)	2.7
60- 79	30/115 (26%)	3.8
80+	31/109 (28%)	3.9

Note: 63/80 events were noted in ED

Sun, BC, et al. J Am Ger Soc, 2007

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Incidence Rates of Syncope According to Age and Sex



Soteriades, E. et al. N Engl J Med 2002;347:878-885

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Syncope - History

High - risk

- Older age ✓
- (+) CV diagnosis
- **(+) CHF**
- (+) Family Hx
- No prodrome
- Supine position
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Lower - risk

- younger age
- (-) CV diagnosis
- (-) CHF
- (-) Family Hx
- (+) prodrome
- (-) exertion

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Data to Support Risk Factors

- **Methods:** Retrospective study, Kaiser data, 02-06
 - 22,189 patients with 23,951 syncope episodes
 - 307 deaths in 30 days

- **Results:**

	<u>Hazard Ratio</u>
-CHF: (age 18-59)	14.3
(age 60-79)	3.1
(age 80+)	2.3
-Diabetes	1.5
-Seizure	1.6
-Dementia	1.4

30-day Death rate:

0.2% < 60yrs without CHF, 2.5% all ages with CHF

Derosé SF, et al. Acad Emerg Med 2012

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Syncope - History

High - risk

- Older age
- (+) CV diagnosis
- **(+) CHF**
- (+) Family Hx
- No prodrome
- Supine position
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(think structural outflow obstruction)

Lower - risk

- younger age
- (-) CV diagnosis
- (-) CHF
- (-) Family Hx
- (+) prodrome
- (-) exertion

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Syncope - Step 2 in History The Episode....

Syncope vs. Seizure

- Ask observers
 - Duration
 - Interventions required
 - Seizure activity, tongue biting, incontinence

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Syncope vs. Seizure

Don't confuse with ***“convulsive syncope”***

- 0.03% of all blood donors
- 12% of all syncope
- Men > women
- Individual variable response to global cerebral hypoperfusion

Lin JT, et al. Ann Neurol 1982 11(5): 525-8.

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Syncope vs. Seizure

Suggests seizure

- Tongue biting
- Head turning/posturing
- No memory of LOC
- LOC assoc. with stress
- Cyanosis observed
- Limb jerking observed
- Postictal confusion
- Postictal headache

Suggests syncope

- Presyncopal/prodrome
- Warmth before spell
- Remembered LOC
- Prolonged sitting or standing
- Any chest pain
- Palpitations
- Dyspnea

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Syncope History-

Step 1: Risk Factors

Step 2: The Episode

Step 3: **What Are the Meds?**

- Syncope Clinic, Duke Univ
 - 70 pts - 13% of syncope due to meds

Hanlon, JT, et al. Arch Intern Med, 1990

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Syncope History-

Step 3: **What Are the Meds?**

- Drugs that cause hypotension
 - Alpha blockers, diuretics
- Drugs that cause bradyarrhythmias
 - B-blockers, Ca⁺ channel blockers,
 - **Alzheimer meds?**
- Drugs that cause prolonged QT

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Cholinesterase Inhibitors Do Increase Risk of Syncope!!!

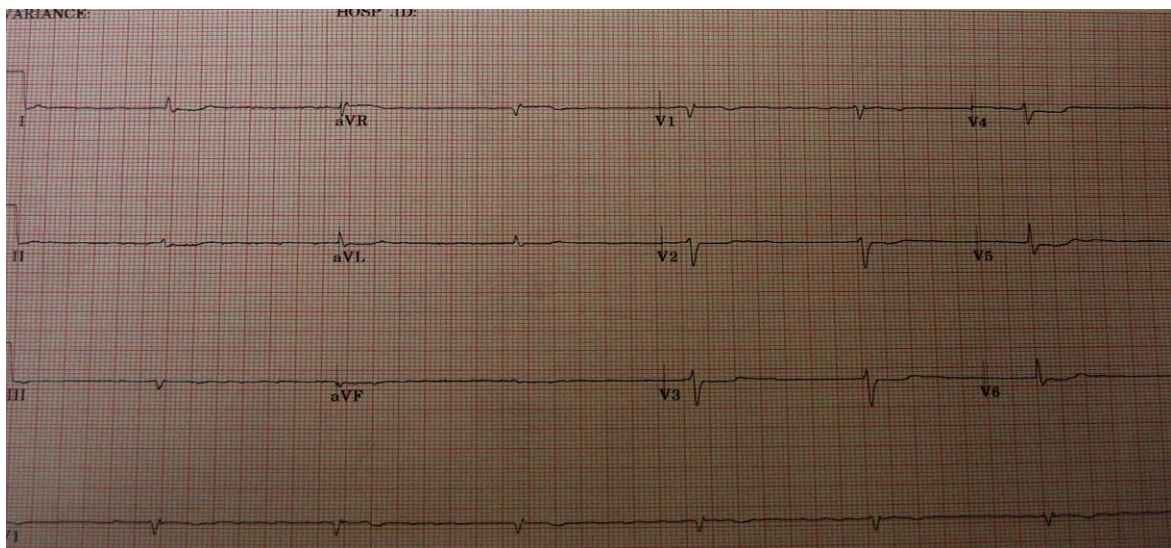
- **Methods:** population-based cohort study,
 - Location: Ontario, Canada; 2002-2004
 - Patients with dx: dementia
- **Results:**

	Controls n=61,499	(+) cholinesterase n= 19,803
Hospital visits for syncope	1.76x (1.57-1.98)	
Bradycardia		1.69x (1.32-2.15)
Pacer insertion		1.49x (1.12-2.00)
Hip fracture		1.18x (1.04-1.34)

Gill, SS, et al. Arch Intern Med, May 11, 2009

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Oct 11, 2009 - 85 y/o Near Syncope Meds: Verapamil, Digoxin, Aricept, Exelon Patch



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Syncope History

Step 3: What Are the Meds?

- Drugs that cause hypotension
 - Alpha blockers, diuretics
- Drugs that cause bradyarrhythmias
 - B-blockers, Ca⁺ channel blockers,
 - *Alzheimer meds?*
- Drugs that cause prolonged QT

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Drugs That Can Cause Prolong QTc/Torsades

Antiarrhythmics

- Amiodarone
- Disopyramide
- Sotalol
- Flecainide
- Ibutilide
- Procainamide
- Propafenone
- Quinidine

Anti-anginal

- Ranolazine

Macrolides

- Erythromycin
- Clarithromycin
- **Azithromycin**

Quinolones

- Ciprofloxacin
- Gatifloxacin
- **Levofloxacin**
- Moxifloxacin

Antifungal

- **Fluconazole**
- Itraconazole

TCA's/SSRI's

- Amitriptyline
- Desipramine
- Doxepin
- Fluoxetine
- Imipramine
- Paroxetine
- Sertraline
- Trazodone
- Venlafaxine

Anti-emetic

- **ondansetron**
- **droperidol**

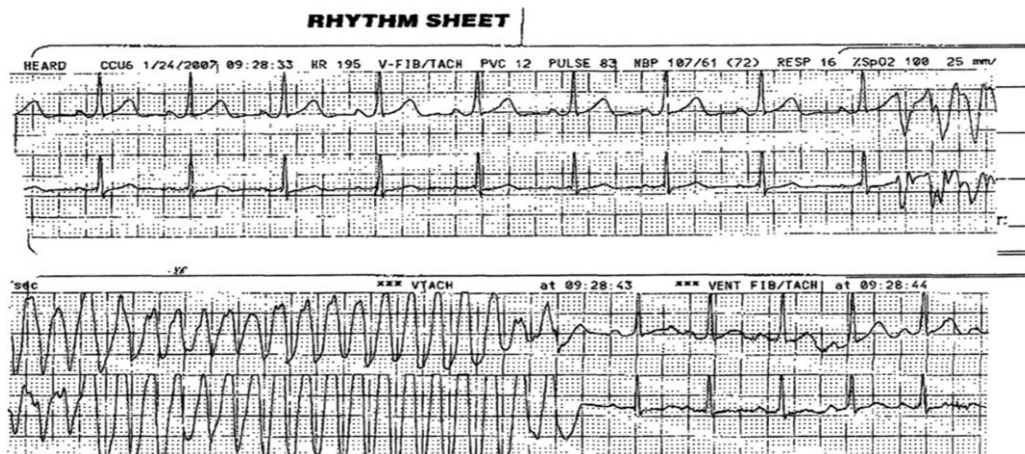
Antipsychotic

- Droperidol
- Haloperidol
- Pimozide
- **Quetiapine**
- Risperidone
- Paliperidone
- Thioridazine
- Chlorpromazine
- **Citalopram**

www.crediblemeds.org

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Azithromycin-induced Polymorphic Ventricular Tachycardia (VT) in a 24-year-old Woman with No Structural Heart Disease and a Normal ECG. The Arrhythmias Resolved with Stopping the Drug.



Zhenjiang Yang et al. *Circ Arrhythm Electrophysiol.*
2017;10:e003560



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Is Azithromycin Associated with Arrhythmia and Death?

Yes

- **Ray WA, et al. NEJM 2012**
Tennessee Medicaid cohort
CV death HR 2.49 (1.38-4.5)
compared to amoxicillin
- **Rao GA, et al. Ann Fam Med 2014**
VA data base
All death HR 1.62 (1.15-2.3)
compared to amoxicillin

No

- **Svanstrom, et al. NEJM 2013**
Danish database
No difference vs. PCN
- **Trifiro G, et al. CMAJ 2017**
European database
No difference vs. Amoxicillin

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Syncope History-

Step 1: Risk Factors

Step 2: The Episode

Step 3: What Are the Meds?

Next--- The Physical Exam

But, what is useful?

- Vitals Signs
- Cardiac Exam
- Abdominal Exam
- Neuro Exam

??????????

- *Orthostatics?*
- *Carotid sinus massage?*

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Orthostatic Hypotension

- Defined: 20mmHg drop in BPsys **standing**
- May indicate:
 - volume depletion
 - Cardiac pump failure
 - Medications
 - Autonomic insufficiency

However, it is present in asymptomatic pts
40% in age >70
23% in age <60

Atkins, D, et al. Am J Med, 1991

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Orthostatic Hypotension

- Defined: 20mmHg drop in BPsys standing

Not to be confused with....

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Orthostatic Hypotension

- Defined: 20mmHg drop in BPsys standing

Not to be confused with....

Postural (orthostatic) tachycardia syndrome (POTS)

- Defined: Increase HR ≥ 30 bpm
 - No orthostatic hypotension
 - Frequent symptoms upon standing (*lightheaded, palpitations, tremulousness, blurred vision, weakness*)
 - **Will feel pre-syncope, but do not pass out**

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Postural (Orthostatic) Tachycardia Syndrome (POTS)

- Pathophysiology: dysautonomia, ?autoimmune?
- Common: 1-3 million Americans
 - 5-10x more common than orthostatic hypotension
 - Women:Men: 5:1

POTS incidence rises significantly post-COVID-19

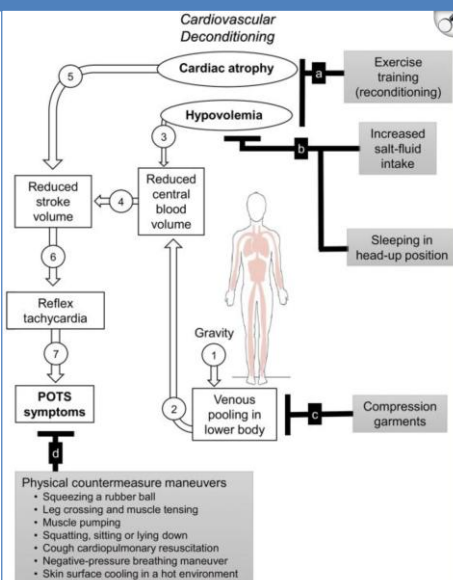


(Pixabay)

The incidence rate of postural orthostatic tachycardia syndrome has risen significantly since the COVID-19 pandemic, according to a study in the European Heart Journal -- Quality of Care and Clinical Outcomes. Researchers using TriNetX data found that post-pandemic, the incidence rate increased from 1.42 to 20.3 per 1 million person-years. They emphasized the need to screen for the condition in post-COVID patients. **Full Story:** [Healio](#)

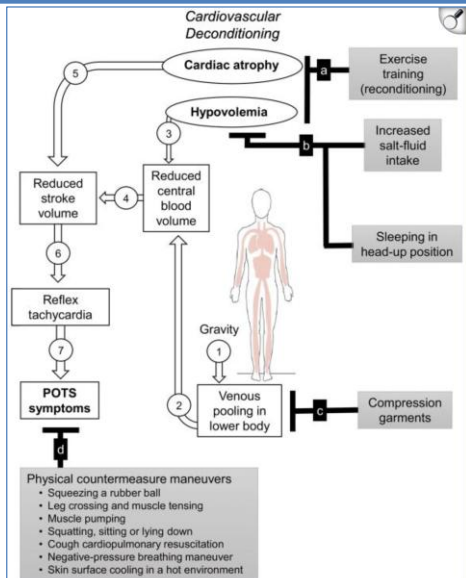
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Postural (Orthostatic) Tachycardia Syndrome (POTS)



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Postural (Orthostatic) Tachycardia Syndrome (POTS)



Drug therapy

- ✓ midodrine,
- ✓ ivabradine,
- ✓ bisoprolol,
- ✓ fludrocortisone,
- ✓ droxidopa,
- ✓ desmopressin,
- ✓ propranolol,
- ✓ modafinil,
- ✓ methylphenidate,
- ✓ melatonin

All sample sizes 10-50 subjects.

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Carotid sinus hypersensitivity

- First described by Ibn Sina (Avicenna) 980-1037
- Asystole > 3 sec after 5-10 sec of **carotid message** *OR* **drop of 50mm Hg BPsyst**
- Suggested as common cause of syncope and falls in the elderly
- Pacing is effective for bradyarrhythmia
Pacing **not** effective for vasodepressive



- **How common is it?**
- **Is it the cause of the syncopal episode?**

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Carotid Sinus Hypersensitivity

- Methods: 272 pts., age >65, single practice in GB

- Results: (+) CSH in 107 (37%)

- Pts with (+) hx of syncope,
falls, dizziness (n= 192)

41%

Pts with no hx of syncope
no falls, dizziness (n=80)

35%

Kerr, SRJ, et al. Arch Intern Med, 2006

■ **How common is it?** *Very common, in elderly*

■ **Is it the cause of the syncopal episode?** ???

*The presence of CSH (and orthostatics)
does **not** preclude looking for other causes*

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Syncope History- Step 1: Risk Factors Step 2: The Episode

Step 3: What Are the Meds? Step 4--- The Physical Exam But, What's Useful?

- Vitals Signs
- Cardiac Exam
- Abdominal Exam
- Neuro Exam

- Orthostatics
- Carotid sinus massage?

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Syncope History-

Step 1: Risk Factors

Step 2: The Episode

Step 3: What Are the Meds?

Step 4--- The Physical Exam

Step 5 – the tests....
But which ones?

Ask the question: “How are you feeling?”

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Syncope: The Dangerous Causes

- ACS
- Aortic dissection
- PE
- AAA
- Ectopic pregnancy
- GI bleed
- SAH

➤ Routine troponin¹,
➤ R/O MI protocols²,
➤ Head CT³
have low/no yield!

¹Hing, R, et al, 2005

²Link, MS, et al, 2001

³Giglio P, et al, 2005

“Be a sniper, don’t use a shotgun!!”
A. Mattu

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Syncope: What Diagnostic Studies Are Needed (to Risk Stratify)? ACEP 2007

- **Answer: EKG only**
 - Note: the yield is < 5%
 - But it is low-cost, non-invasive
 - And can potentially identify life-threatening conditions
- ACC/AHA agree (2006, 2017)
 - **“Routine and comprehensive testing is not useful” (Class III recommendation)**
- **All other studies are guided by H & P**

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ARS Question 1: ACEP (2007) and AHA/ACC (2017) Policies States Which of the Following Tests Should Be Performed in All Patients with Syncope?

- A. EKG
- B. EKG + CBC
- C. EKG + CBC + troponin
- D. EKG + CBC + troponin + head CT
- E. Head CT

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Same Holds True for Elderly!!

- Methods: 2106 admits for syncope, age > 65
 - Yale Med Center, 7/1/02 - 12/31/06, retrospective
- | <u>Test</u> | <u>#(%)</u> | <u>Helped determine etiology</u> |
|------------------|-------------|----------------------------------|
| EKG | 2081 (99%) | 72 (3%) |
| Telemetry | 2001 (95%) | 95 (5%) |
| Cardiac enzymes | 1991 (95%) | 9 (0.5%) |
| Head CT | 1327 (63%) | 7 (0.5%) |
| Echo | 821 (39%) | 13 (2%) |
| Carotid U/S, EEG | | <1% |

Estimated costs nationally = \$6 billion/year

Mendu ML, et al, Arch Intern Med, July 27, 2009

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Syncope

- Definition - **very useful**
- Pathophysiology - **very useful**
- H & P elements - **that are useful**
- Tests/:
Clinical Policies - **that might be useful**
- EKG's - **that are worrisome!**
- The outpatient evaluation

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Syncope and EKG Findings

- **Bradycardias, conduction abnormalities (blocks)**
- Atrial and ventricular tachycardias
- Wolff-Parkinson-White (WPW)
- Prolonged QTc/Long QT syndrome
- Brugada Syndrome
- Hypertrophic Cardiomyopathy
- Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

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Syncope and EKG findings

1st degree A-V Block, Mobitz 1, Mobitz 2 and 3rd degree A-V block*

* Admit/Cardiology

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Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, **Mobitz 2** and **3rd Degree A-V Block***

- **Bradycardias**, conduction abnormalities (blocks)
 - LBBB:
 - RBBB:
 - Bifascicular blocks
 - Trifascicular blocks

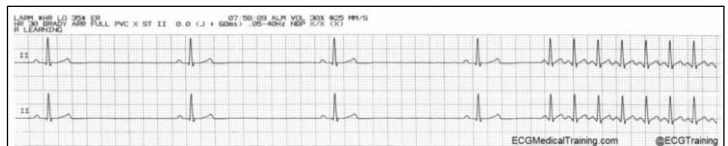
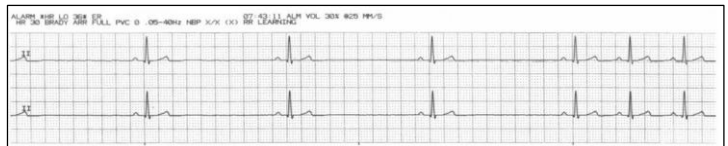
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Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, **Mobitz 2** and **3rd Degree A-V Block***

- **Bradycardias**, conduction abnormalities (blocks)
 - **Sinus node dysfunction (“sick sinus syndrome”)***
*(includes tachycardia-bradycardia syndrome)**

- LBBB
- RBBB
- Bifascicular block
- Trifascicular block



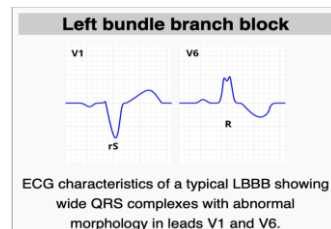
*** Admit/Cardiology**

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Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, **Mobitz 2** and **3rd Degree A-V Block***

- **Bradycardias, conduction abnormalities (blocks)**
 - Sinus node dysfunction (“sick sinus syndrome”) *(includes tachycardia-bradycardia syndrome)*
 - **LBBB: no issue** *(except if new + CP = AMI)**
 - **RBBB:**
 - **Bifascicular blocks**
 - **Trifascicular blocks**



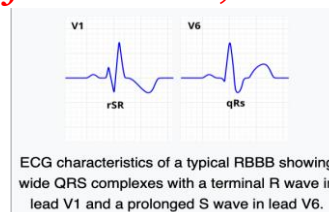
*** Admit/Cardiology**

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Syncope and EKG Findings

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 - **LBBB: no issue** *(except if new + CP = AMI)**
 - **RBBB: no issue** *(except if ST elev V1,2 = Brugada)**
 - **Bifascicular blocks**
 - **Trifascicular blocks**



*** Admit/Cardiology**

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Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, Mobitz 2 and 3rd Degree A-V Block*

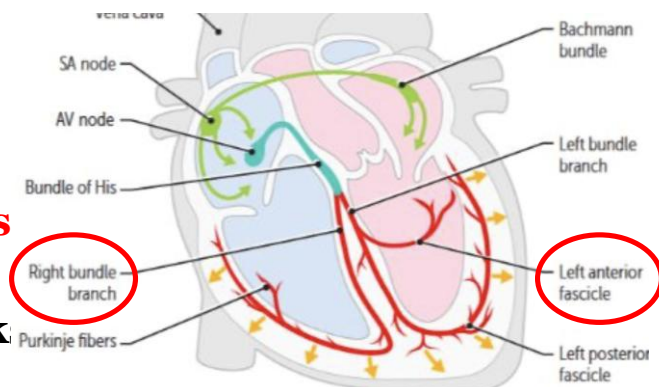
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 - **LBBB: no issue** *(except if new + CP = AMI)**
 - **RBBB: no issue** *(except if ST elev V1 = Brugada)**
 - OR**
 - **Bifascicular blocks = RBBB + L ant. fascicular block***
 - **Trifascicular blocks**

* Admit/Cardiology

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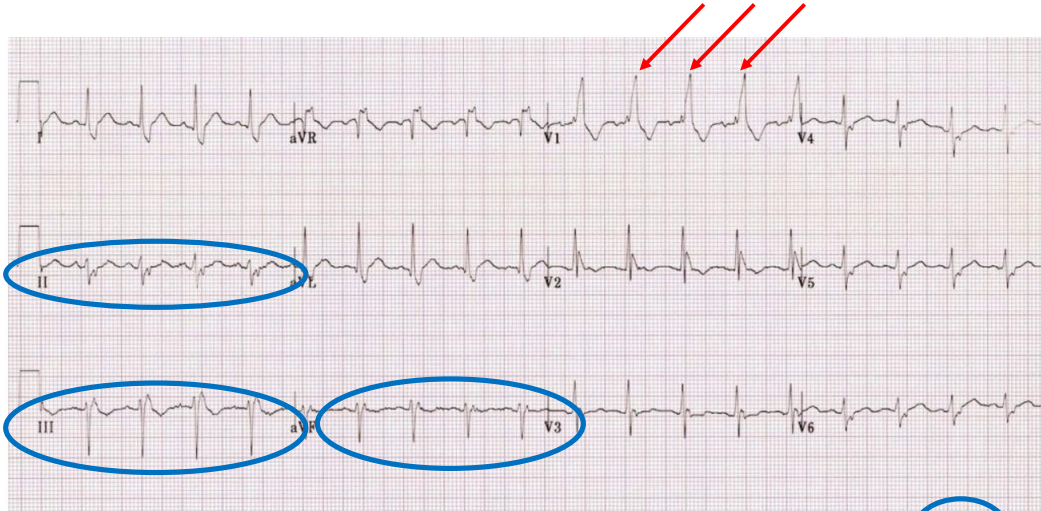
Syncope and EKG Findings

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 - Sinus node dysfunction (“sick sinus syndrome”) *(includes tachycardia-bradycardia syndrome)*
 - LBBB,
 - RBBB
 - **Bifascicular blocks**
 - **Trifascicular block**



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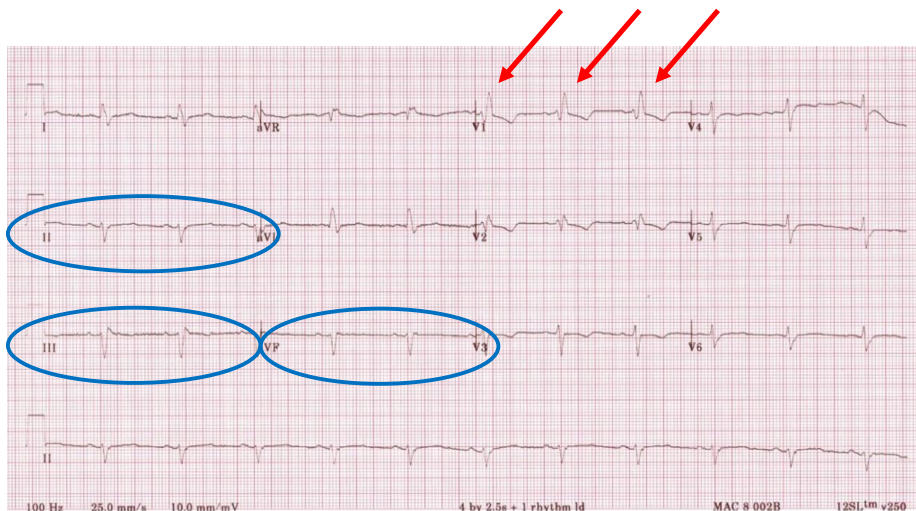
Bifascicular Blocks



Typical bifascicular block pattern: RBBB combined with LAFB (manifested as LAD)

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Bifascicular Blocks



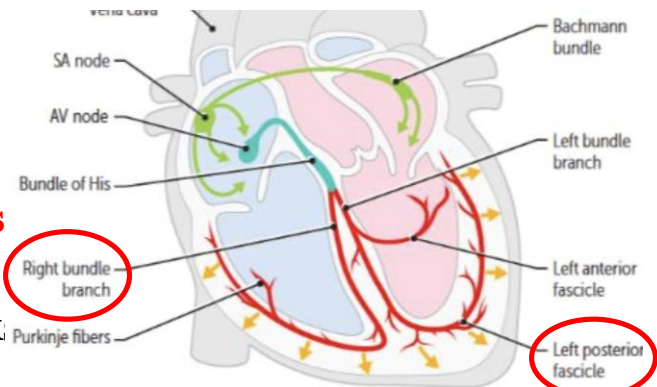
RBBB with LAFB

- RBBB pattern in precordial leads with RSR' complex in V1-2
- Prominent LAD indicating LAFB

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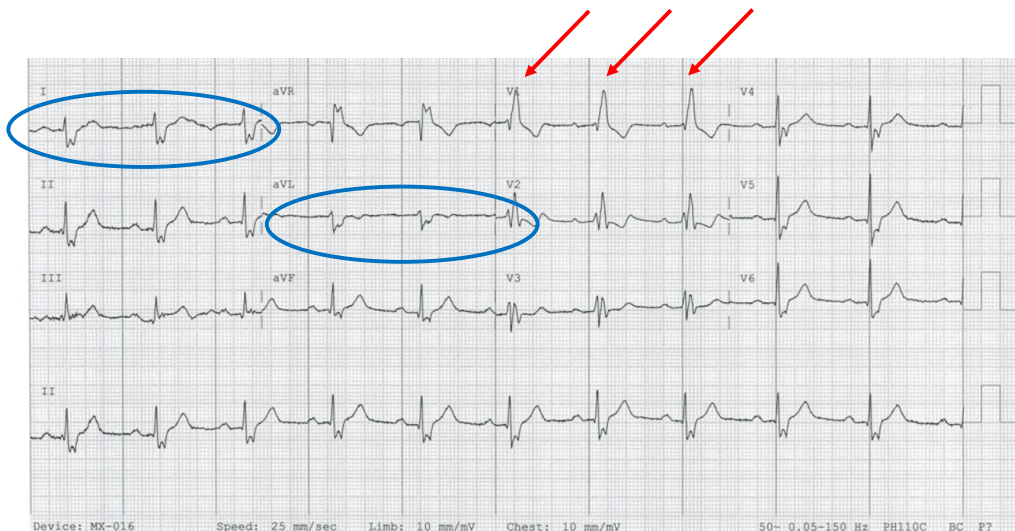
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 - LBBB,
 - RBBB
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 - **Trifascicular block**



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Bifascicular Blocks



RBBB with LPFB and RAD

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Bifascicular Blocks: Clinical Significance

- Structural heart disease present (50-80%) and extensive fibrosis of the conducting system.
- ** **risk of progression to complete heart block**
(with damage to the 3rd fascicle)

➤ Clinical context is important:

- Rate of progression to complete heart block is 1-4% per year
- In symptom free patients, these figures are ~1% per year
- *Patients with syncope have a 17% annual risk of progression*
- **Syncope/presyncope in the context of a bifascicular block is an indication for admission and monitoring!!!!**

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Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, Mobitz 2 and 3rd Degree A-V Block*

- **Bradycardias, conduction abnormalities (blocks)**
 - Sinus node dysfunction (“sick sinus syndrome”)
(includes tachycardia-bradycardia syndrome)
 - **LBBB: no issue** (except if new + CP = AMI)*
 - **RBBB: no issue** (except if ST elev V1 = Brugada)*
 - OR**
 - **Bifascicular blocks = RBBB + L ant. fascicular block***
RBBB + L post. fascicular block*
 - **Trifascicular blocks**

* Admit/Cardiology

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Syncope and EKG Findings

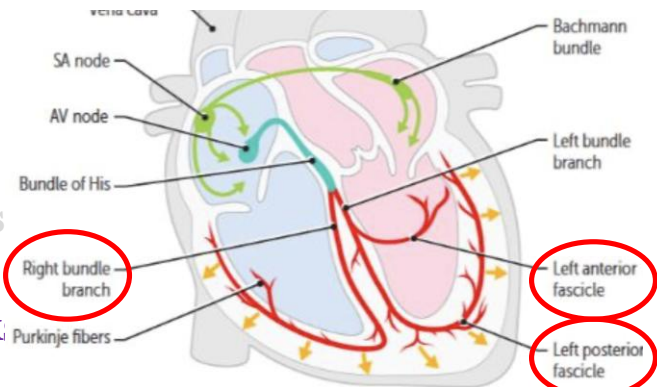
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 - Sinus node dysfunction (“sick sinus syndrome”) (includes tachycardia-bradycardia syndrome)

– LBBB

– RBBB

– Bifascicular blocks

– **Trifascicular block**



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Trifascicular Blocks:

1) "True" Trifascicular: RBBB + LAFB + 3rd AV Block



True Trifascicular Block:

- Right bundle branch block
- Left axis deviation (Left anterior fascicular block)
- Third degree heart block

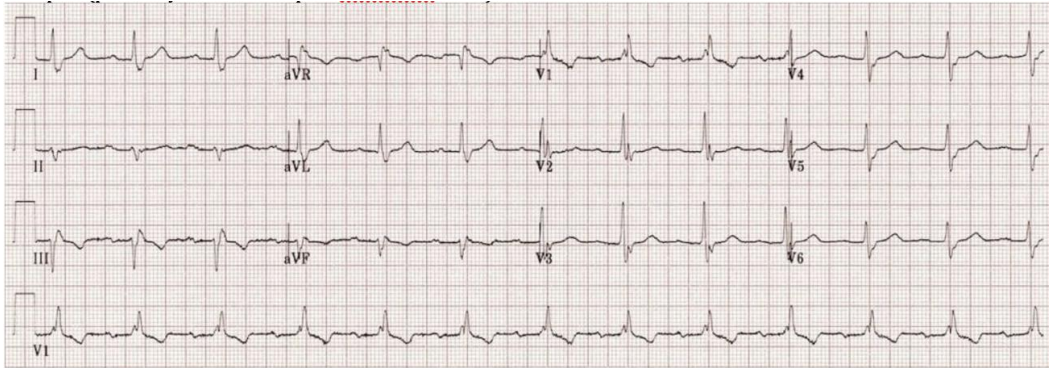
*** Admit/Cardiology**

68

Trifascicular Blocks:

1) "True" Trifascicular: RBBB + LAFB + 3rd AV Block

2) "Incomplete" Trifascicular: RBBB + LAFB + 1st AV Block



Bifascicular block + first degree AV block

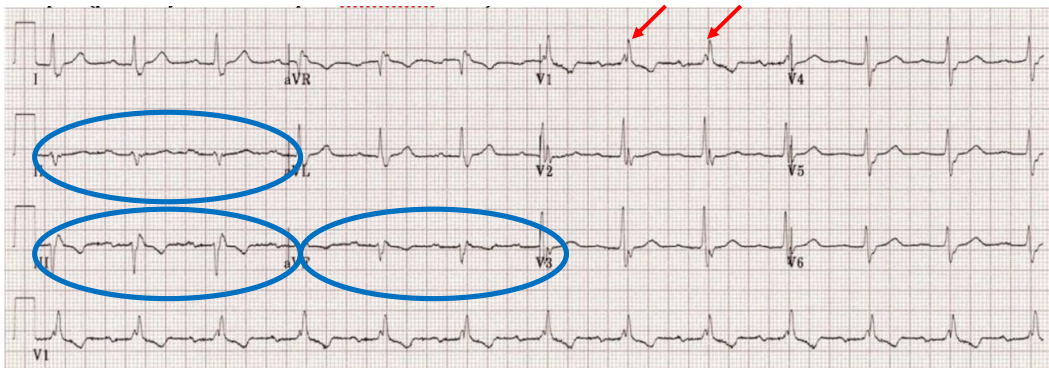
- Right bundle branch block
- Left axis deviation (= left anterior fascicular block)
- First degree AV block

69

Trifascicular Blocks:

1) "True" Trifascicular: RBBB + LAFB + 3rd AV Block

2) "Incomplete" Trifascicular: RBBB + LAFB + 1st AV Block



Bifascicular block + first degree AV block

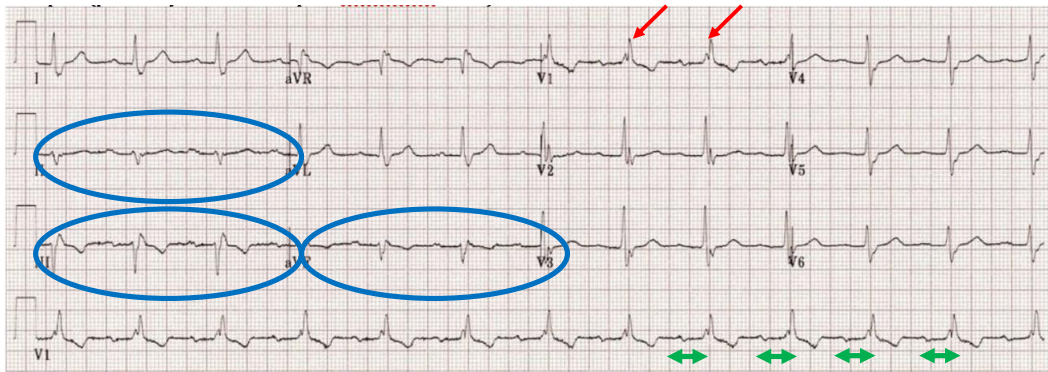
- Right bundle branch block
- Left axis deviation (= left anterior fascicular block)
- First degree AV block

70

Trifascicular Blocks:

1) "True" Trifascicular: RBBB + LAFB + 3rd AV Block

2) "Incomplete" Trifascicular: RBBB + LAFB + 1st AV Block



Bifascicular block + first degree AV block

- Right bundle branch block
- Left axis deviation (= left anterior fascicular block)
- First degree AV block

71

Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, Mobitz 2 and 3rd Degree A-V Block*

- **Bradycardias, conduction abnormalities (blocks)**
 - Sinus node dysfunction ("sick sinus syndrome")*
(includes tachycardia-bradycardia syndrome)*
 - LBBB: **no issue** (except if new + CP = AMI)*
 - RBBB: **no issue** (except if ST elev V1 = Brugada)*
 - OR**
 - **Bifascicular blocks = RBBB + L ant. fascicular block***
RBBB + L post. fascicular block*
 - **Trifascicular blocks***

* Admit/Cardiology

72

Syncope and EKG Findings

- **Bradycardias, conduction abnormalities (blocks)**
- Atrial and ventricular tachycardias
- Wolff-Parkinson-White (WPW)
- Prolonged QTc/Long QT syndrome
- Brugada Syndrome
- Hypertrophic Cardiomyopathy
- Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

* Admit/Cardiology

73

Syncope and EKG Findings

- Bradycardias, conduction abnormalities (blocks)
- Atrial and ventricular tachycardias
- Wolff-Parkinson-White (WPW)
- Prolonged QTc/Long QT syndrome
- Brugada Syndrome
- Hypertrophic Cardiomyopathy
- Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

* Admit/Cardiology

74

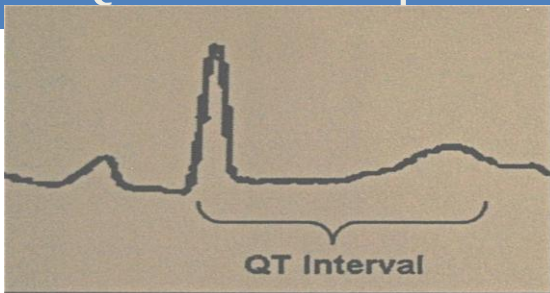
Long QT Syndrome (LQTS)

- **Acquired**
 - Meds, toxins, electrolyte disturbances, ACS, CNS events, HIV
- **Congenital: Autosomal dominant**
 - Associated with 11 genes, 600 mutations
 - Prevalence: estimates 1/2000-7000
 - Median age of sudden death = 32 years
 - Mortality = 20% in first year after syncope
 - 50% mortality within 5 years

***In series of 31 pts. with LQTS, 64% presented with syncope
40% of the patients were not identified at first presentation!
MacCormick JM, et al. Ann Emerg Med, July 2009***

75

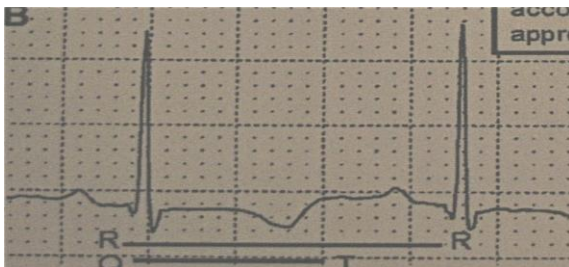
LQTS: Men > 460msec, Women > 440msec



Bazett Formula

$$QT_c = \frac{QT}{[RR]^{0.5}}$$

* In leads II, V5



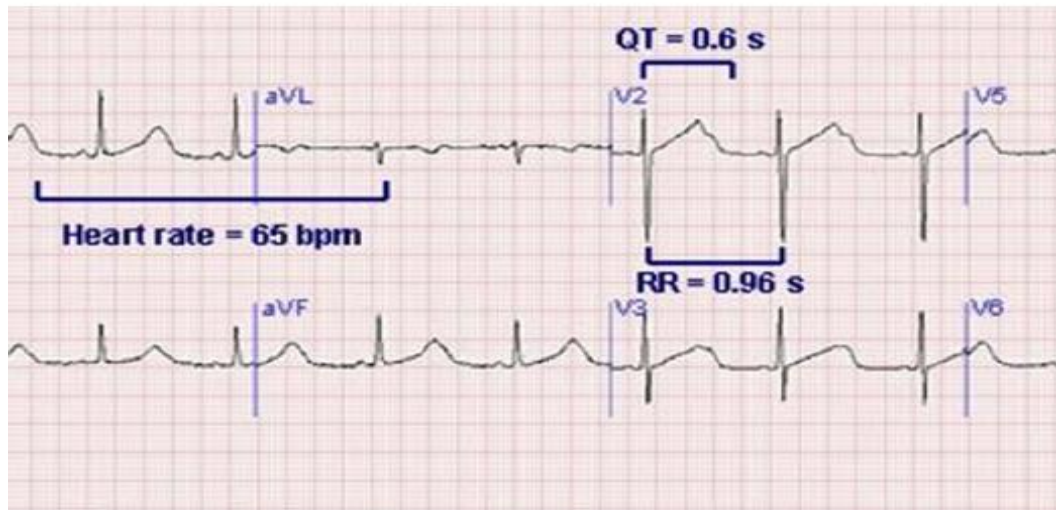
Rapid Bedside

If QT is < 1/2 of
R-R interval is OK*

*only for HR 60-100

76

Long QT Syndrome



77

Brugada Syndrome

- First described in 1992
- Originally thought to be a disease of men of Southeast Asian descent
 - In Philippines: **“Bangungot”**
“scream followed by sudden death during sleep”
 - In Japan: **“Pokkuri”**
“unexpected sudden death at night”
 - In Thailand: **“Lai Tai”** - “death during sleep”
 - In Laos - one death per 1000 inhabitants!!!

78

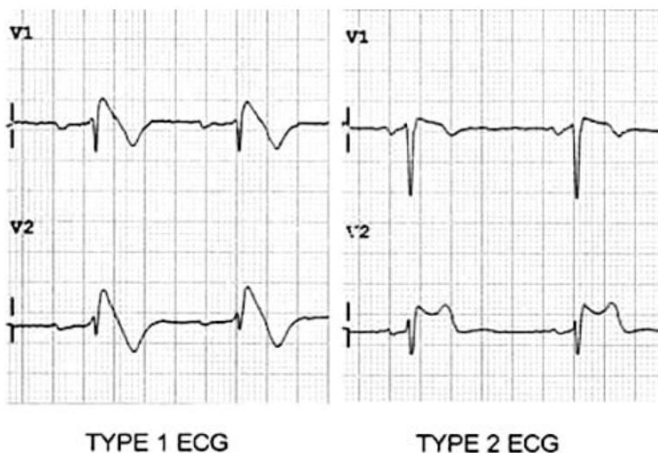
Brugada Syndrome

- Now believed to be responsible for 40-60% of patients with “idiopathic V fib”
 - Second only to MVA cause of death in young adults in some countries
- Mutation of SCN5A cardiac Na⁺ channel on Chromosome 3 (*“channelopathy”*)
- Recurrent episodes of polymorphic V tach

79

Brugada Syndrome: EKG

- **RBBB** with ST elevation V1-3

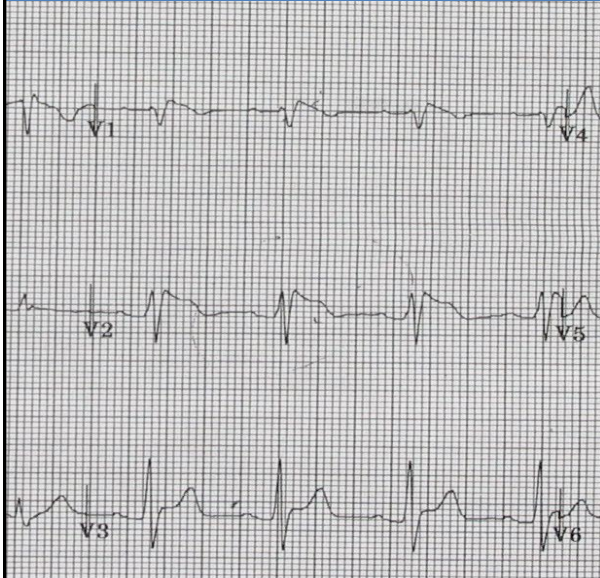


Type I -
“coved-shaped” ST
*is diagnostic

Type II -
“concave-shaped” ST
*is non-diagnostic,
But suggests dx in
appropriate patient

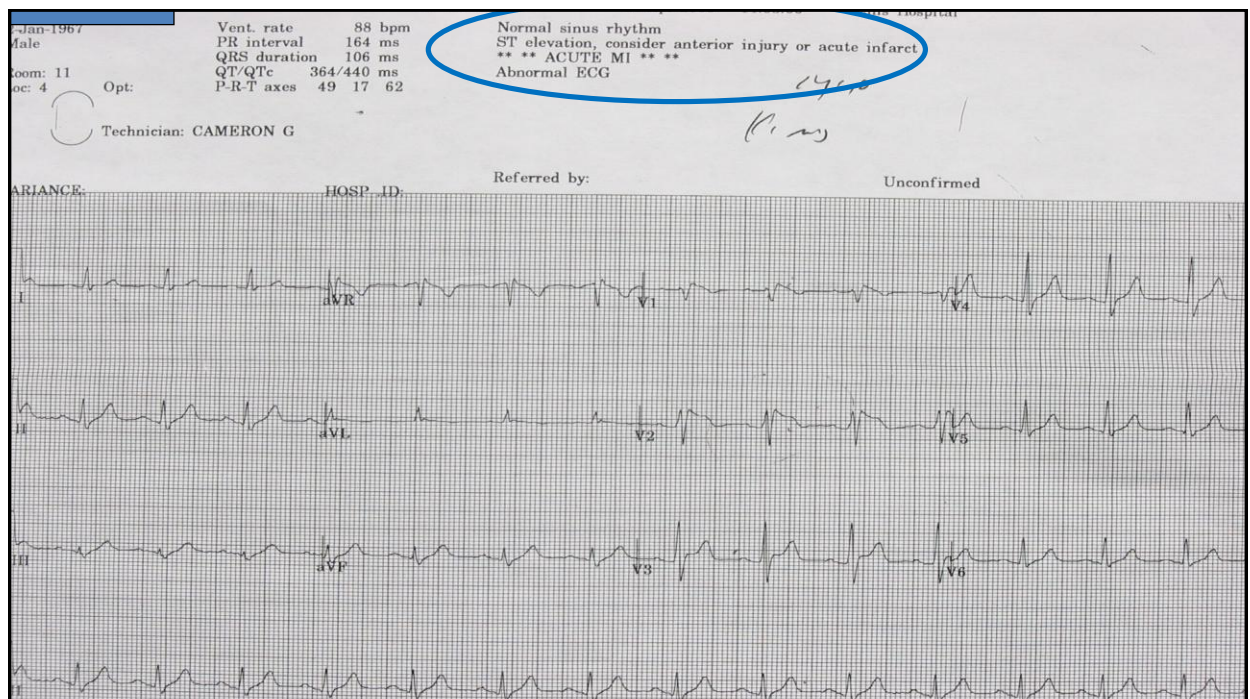
80

Brugada Syndrome: EKG

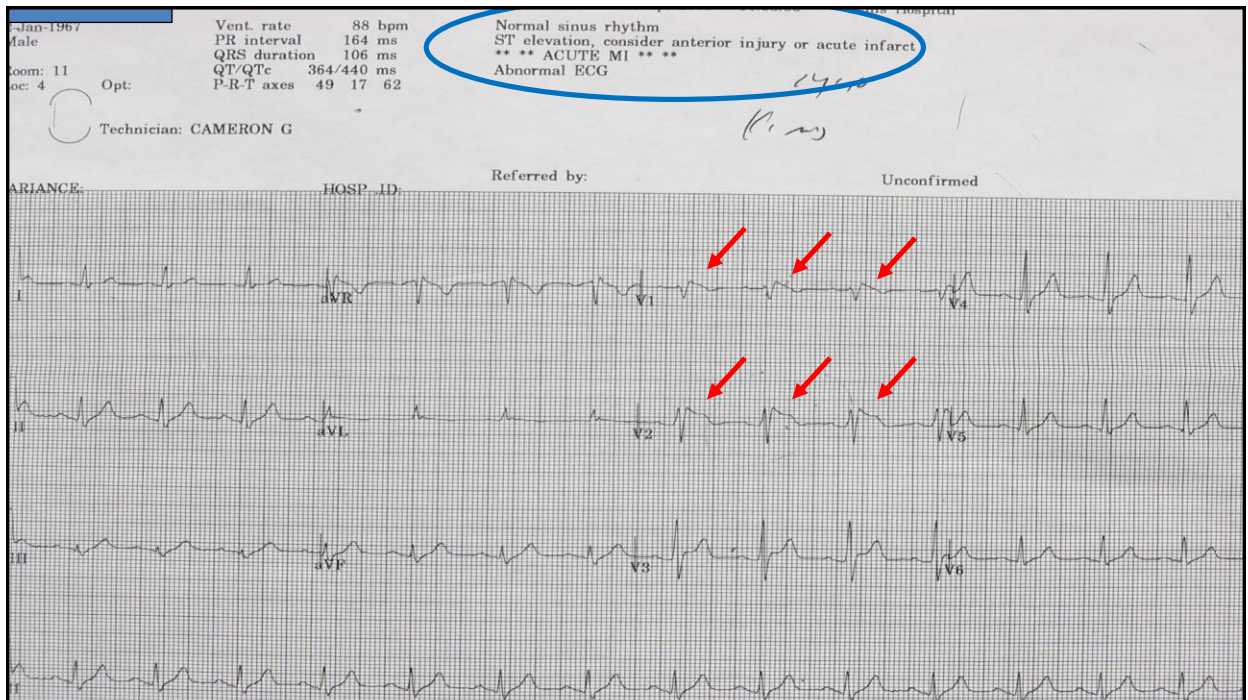


- Look at V1- 3
- Type I

81



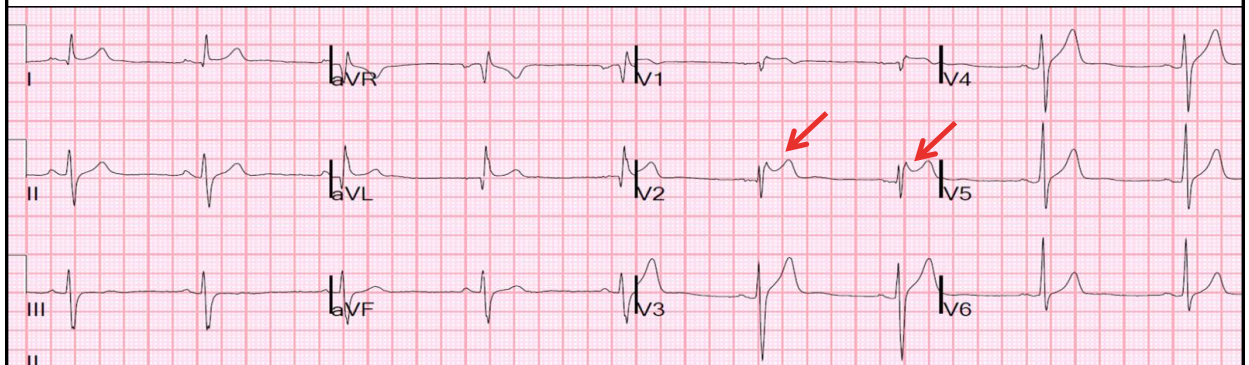
82



83

Brugada Syndrome: EKG

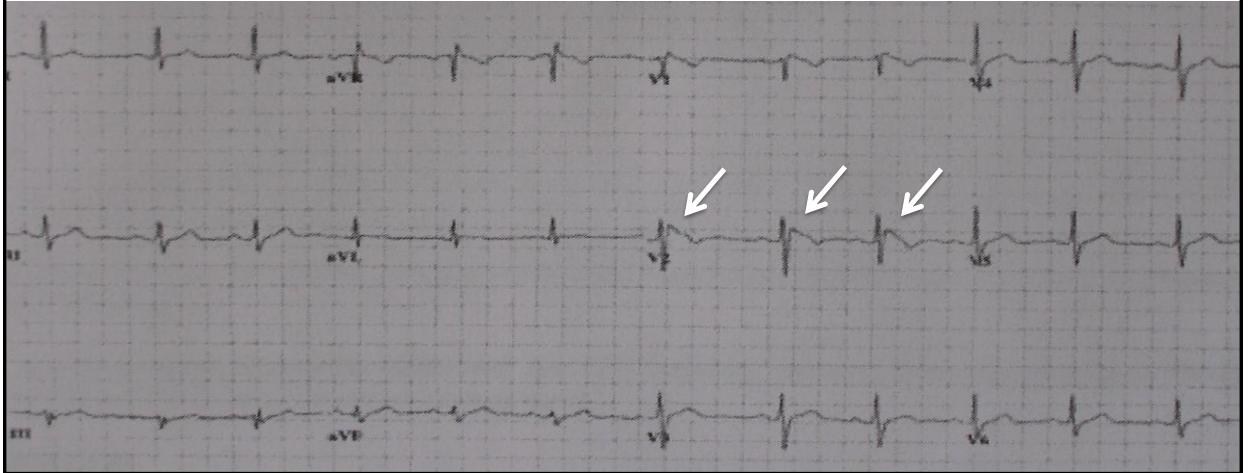
- Type II - incomplete RBBB; “concave-shaped” ST, V1-3



Note “saddleback” in V2

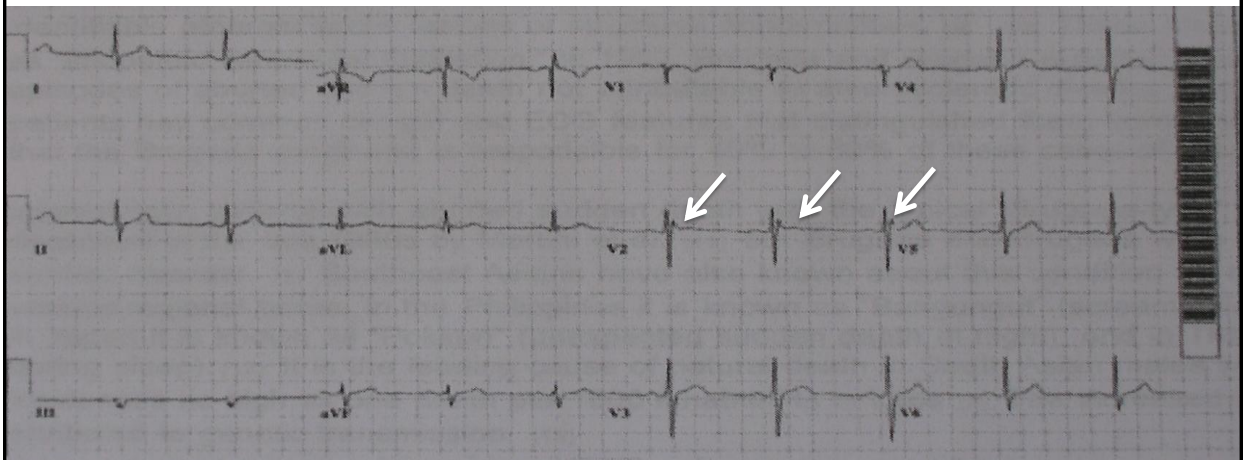
84

30 y/o AA Female, Chest Tightness, Palpitations,
Near-syncope. Previous Hx of Previous Syncopal Episodes



85

9 Years Earlier...



Courtesy A. Mattu, Univ Maryland.

86

Hypertrophic Cardiomyopathy (HCM)

- First described in mid-19th century
- Previous names:
 - Hypertrophic obstructive cardiomyopathy (HOCM)
 - Idiopathic hypertrophic subaortic stenosis (IHSS)
- Characteristics:
 - Thickened myocardium
 - without ventricular dilation
 - absence of conditions that result in hypertrophy (HTN, Aortic stenosis)
- Incidence: approx. 1 in 500 persons
- Annual mortality rate =
=1-2% in unselected pts with HCM

87

Hypertrophic Cardiomyopathy (HCM)

- Genetics: Autosomal dominant with variable penetrance
 - 11 mutant genes, > 500 mutations
- Variable manifestations
 - ==> asymmetric ventricular hypertrophy
 - Most pronounced in anterior ventricular septum
 - Increase occurs most commonly during periods of growth (ie. adolescence)
- Presentation:
 - May be asymptomatic
 - Chest pain
 - Symptoms of LVOT obstruction (SOB, DOE, syncope)
 - Sudden death

Most common cause of sudden death in athletic endeavors!

88

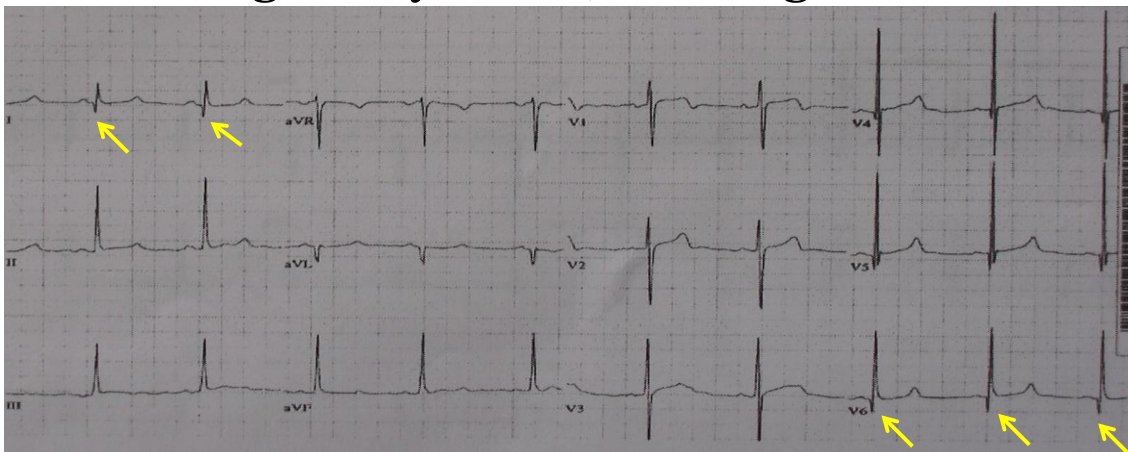
Hypertrophic Cardiomyopathy (HCM)

- PE: murmur noted in 30-40% only
 - Increases with Valsalva
- Chest x-ray: heart with normal size
- EKG: most are abnormal!!!!
 - 1) Large amplitude QRS complex (c/w LVH) ==> **Most common**
 - 2) Deep, narrow Q waves in:
 - Inferior leads (II, III, AVF) and/or
 - Lateral leads (I, aVL, V5-6) ==> **Most specific**
 - 3) ST changes are common → **But non-specific**

89

HCM - LVH, Q Waves in I, aVL, V5-6

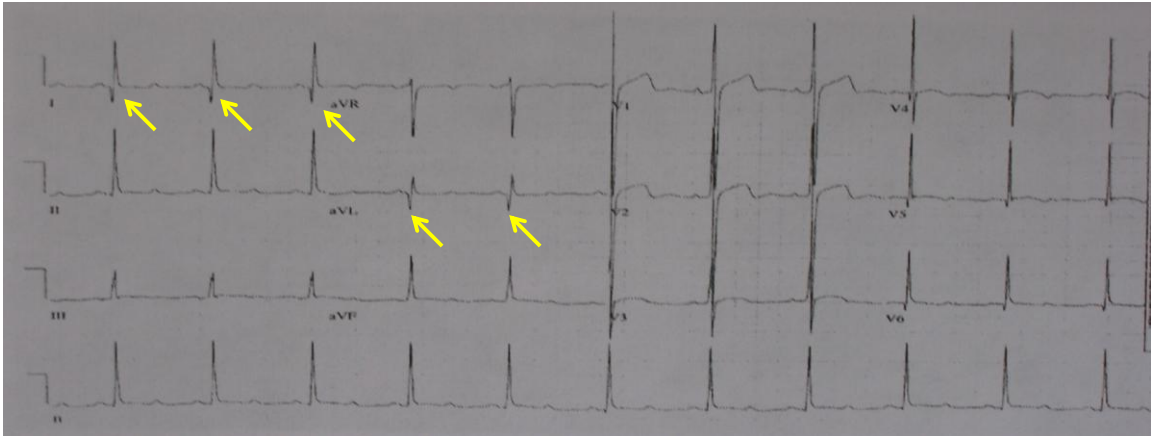
- 30 y/o male lightheaded, palpitations after running. 2 days later, running => SCD.



90

HCM - LVH, Q Waves in I, aVL, V5-6

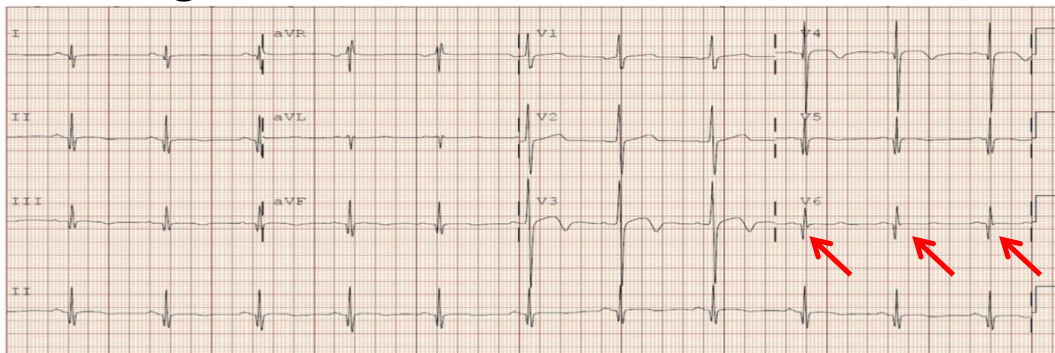
- 29 y/o male 3rd ED visit for lightheaded, palpitations with exertion.



91

HCM - LVH, Q Waves in I, aVL, V5-6

- 34 y/o male with CP and lightheadedness after biking 1-2 hours.



Q waves are defined as 25% of the height of the ensuing R wave

Beck S et al. Ann Emerg Med. May 2017

92

Hypertrophic Cardiomyopathy (HCM)

- Treatment: often medical (B-blockers)
- Complications:
 - 10-40% develop atrial fibrillation
 - Increase incidence of WPW

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Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

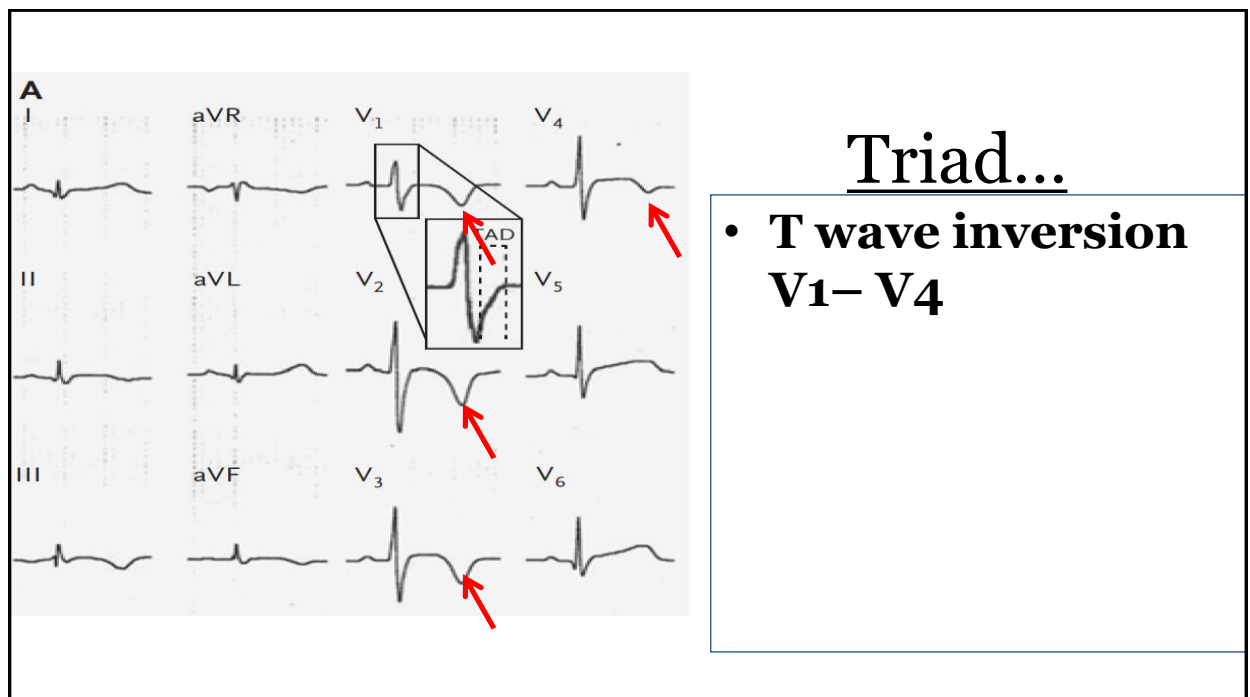
- Described in 1982 (*Marcus et al*)
- Progressive loss of R ventricle myocardium (***replaced with fibrofatty tissue → thinning AND.. Ventricular arrhythmias***)
- Autosomal dominant (*but incomplete penetrance*)
- Prevalence: 1/ 2000-5000 (Italy, Germany)
 - Veneto region: 20% of deaths in young people

94

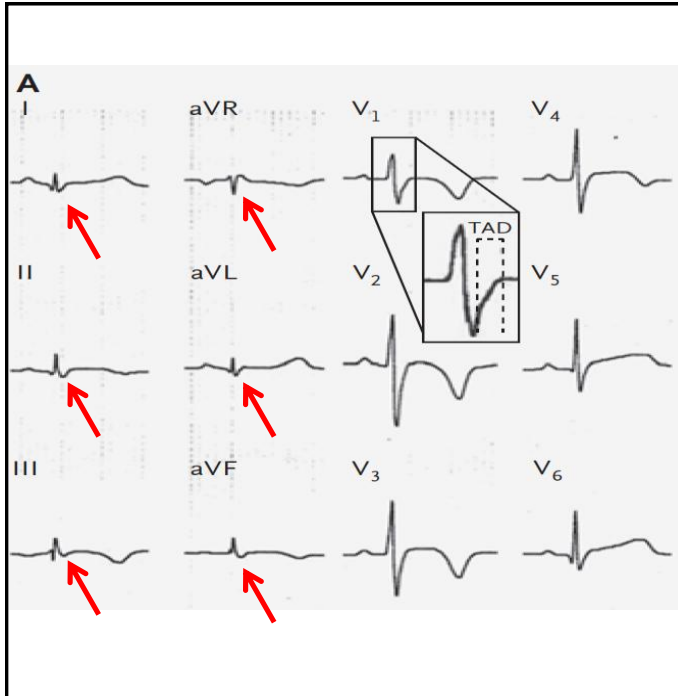
Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

- Presentation: palpitations or effort-induced syncope
- When: 2nd – 4th decade of life.

95



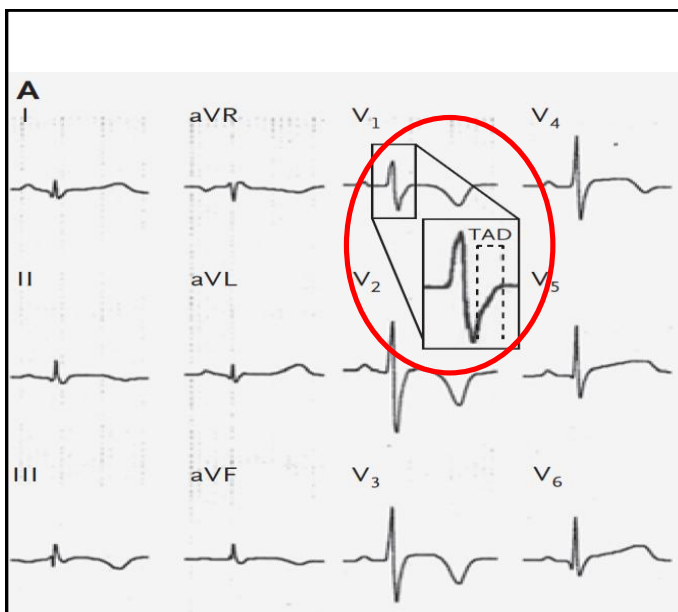
96



Triad...

- **T wave inversion V1– V4**
- **Low voltage limb leads**

97



Triad...

- **T wave inversion V1– V4**
- **Low voltage limb leads**
- **Terminal activation duration (TAD)* is prolonged (normal < 55msec)**

TAD – nadir of S wave to the end of depolarization

98

Syncope and EKG Findings

1st Degree A-V Block, Mobitz 1, Mobitz 2 and 3rd Degree A-V Block*

- SSS/Tachy-brady, conduction abnormalities (blocks)
- Atrial and ventricular tachycardias
- Wolff-Parkinson-White (WPW)
- Prolonged QTc/Long QT syndrome
- Brugada Syndrome
- Hypertrophic Cardiomyopathy
- Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)

* Admit/Cardiology

99

Syncope: The Evaluation

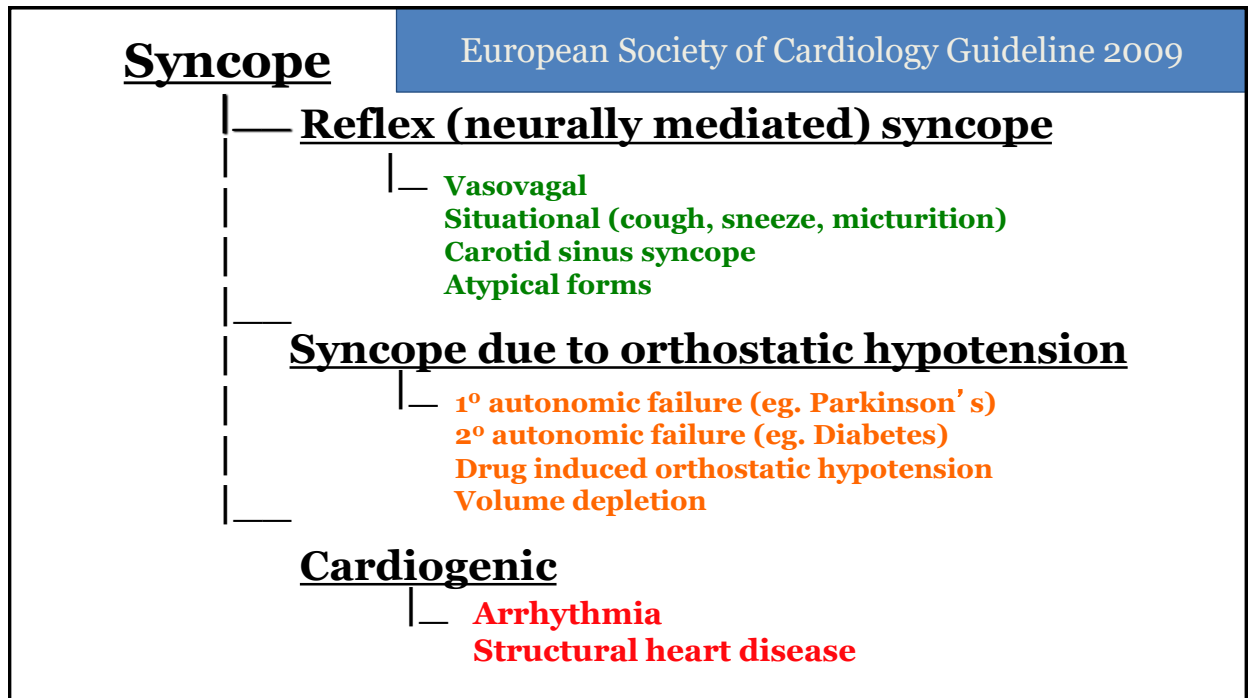
- I. The History-
 - Step 1: Risk Factors
 - Step 2: The episode
 - Step 3: What are the meds?
 - Step 4: "How are you feeling"?

II. The Physical Exam

III. The EKG (and "sniper" diagnostics)

Decision Time: Admit or discharge?

100



101

The Bottom Line - and Where Do We Go from Here?

- 650 ED patients with syncope prospectively evaluated
- Results: 495 (76%) diagnosed in ED
 - Vasovagal syncope 234/650 (47%)
 - Hypotension 156/650 (32%)
 - Neuro/psychiatric 41/650 (8%)
 - Cardiac (AMI, PE, AS) 29/650 (6%)
 - Arrhythmia 26/650 (5%)
- 155(24%) unexplained syncope

Sarasin FP, et al. Heart 2002; 88: 363-67

102

Unexplained Syncope: Evaluation

In-patient

VS

Out-patient

Will a clinical decision rule (CDR) help?

Answer: Perhaps...risk stratify?

103

CDR's for Syncope

- SF Syncope Rule
- Boston
- OESIL - Italy
- EGSYS - Italy
- ROSE – Scotland
- STePs (2008)
- Syncope Risk score
- Canadian Syncope Risk score**

Predisposition to vasovagal symptoms Triggered by being in a warm crowded place, prolonged standing, fear, emotion, or pain	No 0	Yes -1
Heart disease history CAD, atrial fibrillation or flutter, CHF, valvular disease	No 0	Yes +1
sBP <90 or >180 mmHg On any reading	No 0	Yes +2
Elevated troponin >99th percentile of normal population	No 0	Yes +2
Abnormal QRS axis <-30° or >100°	No 0	Yes +1
QRS duration >130 ms	No 0	Yes +1
Corrected QT interval >480 ms	No 0	Yes +2
ED diagnosis Based on ED evaluation	Vasovagal syncope -2 Cardiac syncope +2 Neither 0	
0 points Canadian Syncope Risk Score	Low risk 1.9% risk of 30-day serious adverse event (death, arrhythmia, MI — full list in Evidence)	

104

Unexplained Syncope: Evaluation ACC/AHA 2017

A. ECHO → ...if structural heart dz is suspected (Class IIa)

B. EST → ...if associated with exertion (Class IIa)

C. Non-invasive ECG monitoring

- Holter, Event recorders
- Implantable loop recorders



...if you suspect
arrhythmia (Class IIa)

D. Tilt-table testing → ...useful for suspect vasovagal (Class IIa)

E. Carotid artery studies →

105

Anderson KL, et al. Ann Emerg Med
October 2012; 60:478-484.

323 ED pts admitted to “observation unit”

267 had “**normal**” EKG

235 underwent ECHO

Result: # of abnormal ECHO studies=

o, Zero, Zip, nada, etc....

106

Unexplained Syncope: Evaluation ACC/AHA 2017

A. ECHO → ...if structural heart dz is suspected (Class IIa)

B. EST → ...if associated with exertion (Class IIa)

C. Non-invasive ECG monitoring

- Holter, Event recorders
- Implantable loop recorders



...if you suspect
arrhythmia (Class IIa)

D. Tilt-table testing → ...useful for suspect vasovagal (Class IIa)

E. Carotid artery studies →

107



108

Unexplained Syncope: Evaluation ACC/AHA 2017

- A. ECHO → ...if structural heart dz is suspected (Class IIa)
- B. EST → ...if associated with exertion (Class IIa)
- C. Non-invasive ECG monitoring
 - Holter, Event recorders → ...if you suspect arrhythmia (Class IIa)
 - Implantable loop recorders
- D. Tilt-table testing → ...useful for suspect vasovagal (Class IIa)
- E. Carotid artery studies → No, no no!!! (Class III)

109

The Bottom Line - and Where Do We Go from Here?

- 650 ED patients with syncope prospectively evaluated
- Results: 495 (76%) diagnosed in ED
 - Vasovagal syncope 234/650 (47%)
 - Hypotension 156/650 (32%)
 - Neuro/psychiatric 41/650 (8%)
 - Cardiac (AMI, PE, AS) 29/650 (6%)
 - Arrhythmia 26/650 (5%)
- 155(24%) unexplained syncope

Sarasin FP, et al. Heart 2002; 88: 363-67

110

Reflex (Neurally Mediated) Syncope- Vasovagal Syncope – Most Common

- 20-40% of the population have an episode in lifetime^{1,2}
- Theory: Failure (abnormal) autonomic NS to maintain BP & HR
 - Starts with excessive peripheral pooling (500-800ml blood)
 - Subsequent paradoxical rise in vagal activity
 - Can be “situational” (eg. defecation, urination, cough)

¹Salari N, et al [Global Epidemiology](#) June 2024, 100136

²M Tajdini, et al. *European Heart J*, November 2023

111

Reflex (Neurally Mediated) Syncope- Vasovagal Syncope – Most Common

- 20-40% of the population have an episode in lifetime^{1,2}
- Theory: Failure (abnormal) autonomic NS to maintain BP & HR
 - Starts with excessive peripheral pooling
 - Subsequent paradoxical rise in vagal activity
 - Can be “situational” (eg. defecation, urination, cough)

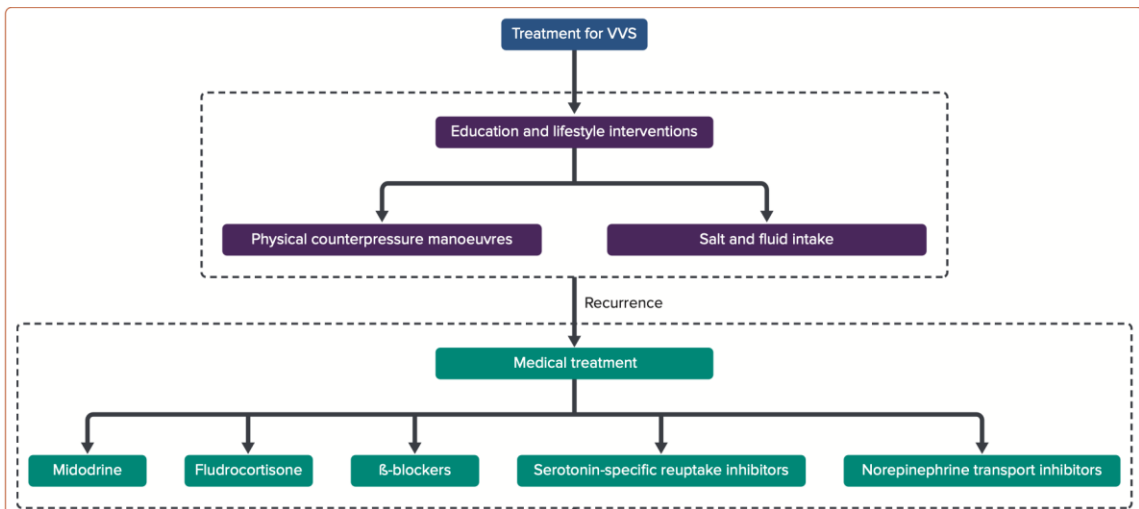
35% of patients will have syncope recurrences

- not associated with increased mortality,
- + associated with injury and reduced quality of life

Raj SR, et al. *JACC: Clinical Electrophysiology*, 2022

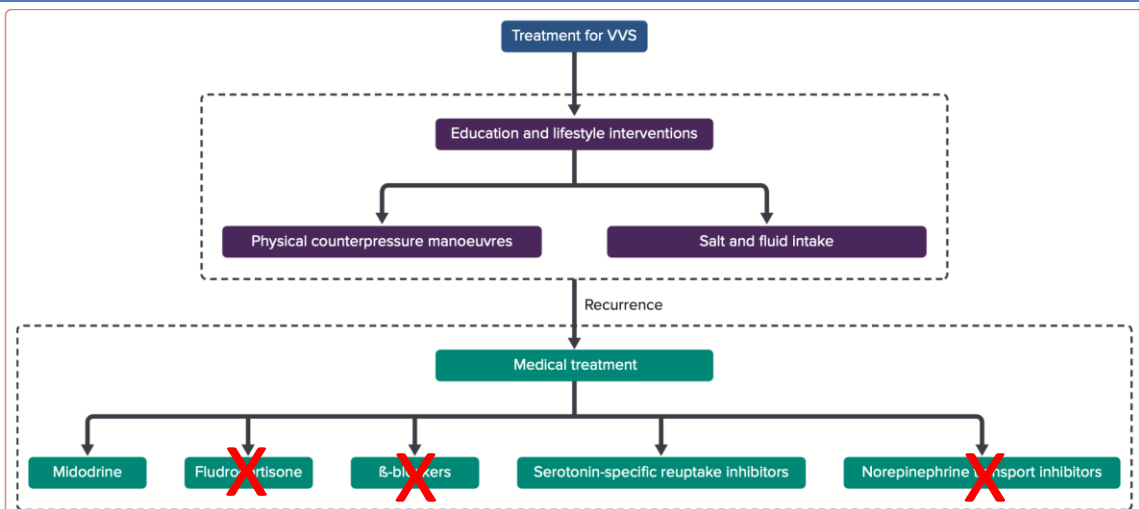
112

Recurrent Vasovagal Syncope: Management



113

Recurrent Vasovagal Syncope: Management



114

ARS Question 2: Which One of the Following Does **NOT** Decrease Recurrent Episodes of Vasovagal Syncope?

- A. Midodrine
- B. B-blockers
- C. SSRI
- D. Closed loop stimulation pacing



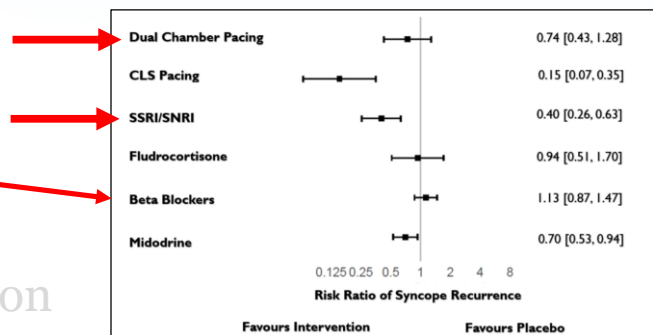
115

ARS Question 2: Which One of the Following Does **NOT** in Decrease Recurrent Episodes of Vasovagal Syncope?

A. Midodrine

B. B-blockers

C. SSRI

D. Closed loop stimulation
pacing

Kaza N, et al. Open Heart 2024; 11: e002669



116

Recurrent Vasovagal Syncope: Medical Management

- **Midodrine:** 2 RCT, placebo-controlled studies: 179 patients

Methods: 133 pts with aver. 6 syncope episodes in the prior year
(median age, 32 years; 73% female)

Results @ 1year:	<u>Midodrine</u>	<u>placebo</u>
(+) syncopal episode	28/66 (42%)	41/67 (61%)

$RRR = 0.69$ (95% CI, 0.49 to 0.97; $P = 0.035$)

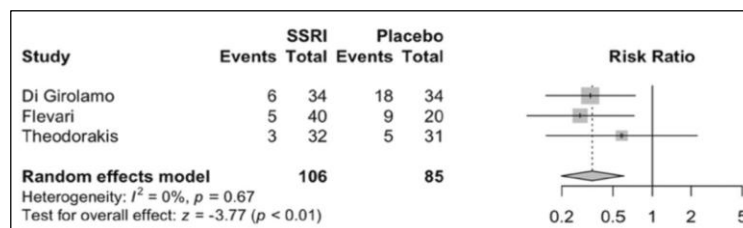
NNT= 5

Sheldon R, et al . Ann Intern Med 2021; 174: <https://doi.org/10.7326/M20-5>

117

Recurrent Vasovagal Syncope: Medical Management

- **Midodrine:** 2 RCT, placebo-controlled studies: 179 patients
- **SSRI/SNRI:** 3 RCT, placebo-controlled studies: 204 patients
Paroxetine and fluoxetine

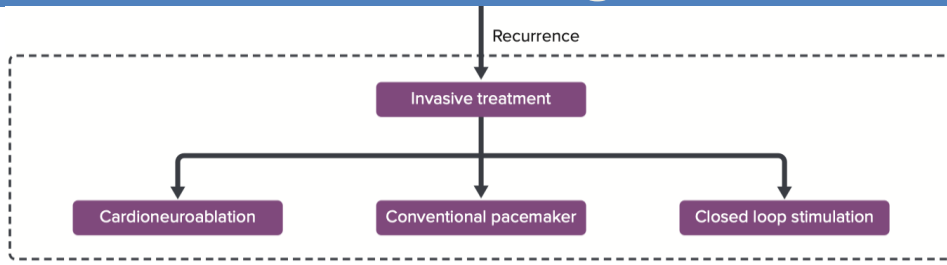


13% Vs. 37%; NNT= 4

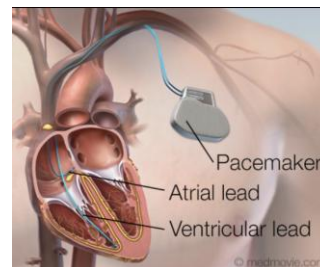
Raj P et al. Clinical Autonomic Research (2023) 33:811–819

118

Recurrent Vasovagal Syncope: When Medical Management Fails....

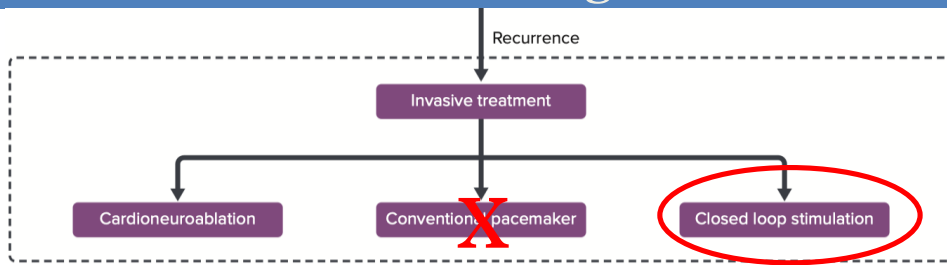


Sensor in right atrium identifies reduced right ventricular filling, triggering the CLS algorithm → ...increase the pacing rate

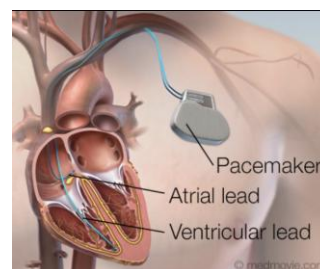


119

Recurrent Vasovagal Syncope: When Medical Management Fails....

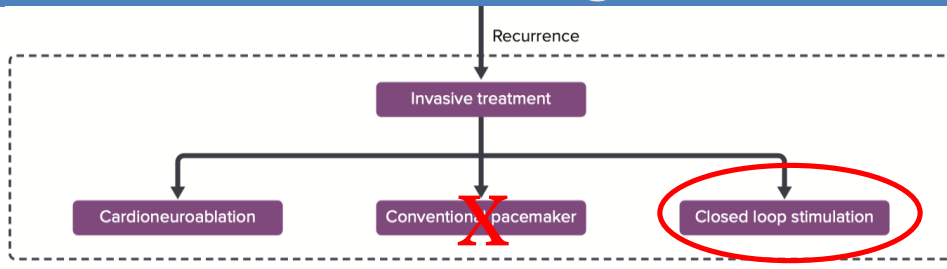


Sensor in right atrium identifies reduced right ventricular filling, triggering the CLS algorithm → ...increase the pacing rate



120

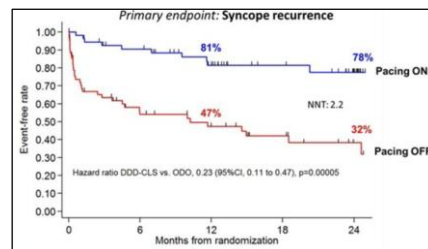
Recurrent Vasovagal Syncope: When Medical Management Fails....



4 RCT trials: 345 patients¹

¹Kaza N, et al. Open Heart 2024; 11: e002669

²Brignole M et al. European Heart Journal, 2021; 42: 508–516



2

121

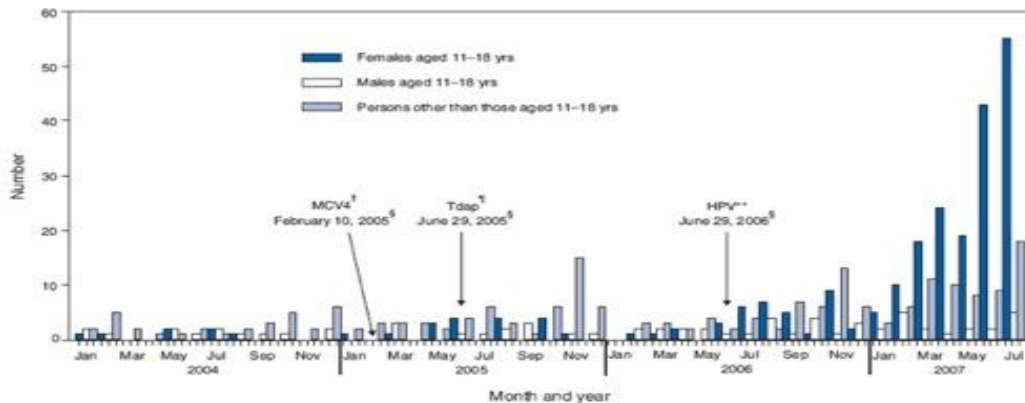
Last Word on Vasovagal Syncope: In the Office... Be Prepared

- At risk: young > old, women > men (1.5x)
Note: average of age of 1st episode = 14
- Painful procedures
 - blood draws
 - Vaccination
 - Tdap, meningococcal, **HPV**

122

Watch Out for HPV Vaccine Syncope

FIGURE. Number of postvaccination syncope* episodes reported to the Vaccine Adverse Event Reporting System, by month and year of report — United States, January 1, 2004–July 31, 2007



* Includes persons aged ≥ 5 years who had syncope onset after vaccination on the same date.

† Meningococcal conjugate vaccine.

‡ Date on which the Advisory Committee on Immunization Practices decided to add this newly licensed adolescent vaccine to the Vaccines for Children Program.

§ Tetanus toxoid, reduced diphtheria toxoid, and acellular pertussis vaccine.

** Quadrivalent human papillomavirus recombinant vaccine. HPV is licensed only for females.

123

Syncope

- Definition - *very useful*
- Pathophysiology - *very useful*
- H & P elements - *that are useful*
- Tests/
Clinical Policies - *use EKG + “sniper diagnostics”*
- EKG's - *that are worrisome!*
- The outpatient evaluation

124

Thank You for Your Time and Consideration!!

Post Lecture ARS Questions.....

125

ACEP (2007) and AHA/ACC (2017) Policies States Which of the Following Tests Should Be Performed in All Patients with Syncope?

- A. EKG
- B. EKG + CBC
- C. EKG + CBC + troponin
- D. EKG + CBC + troponin + head CT
- E. Head CT

126

Which One of the Following Does NOT Decrease Recurrent Episodes of Vasovagal Syncope?

- A. Midodrine
- B. B-blockers
- C. SSRI's
- D. Closed loop stimulation pacing



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Supplemental Slides

128

The Role of Echo in the Evaluation of Syncope: A Prospective Study.

Sarasin FP, et al. Heart 2002; 88: 363-67.

- 155 patients with unexplained syncope
 - All get ECHO
- Result: # of abnormal ECHO studies=

o, Zero, Zip, nada, etc...

129

Echocardiography in the Evaluation of Patients with Syncope.

Recchia D, et al. J Gen Intern Med 1995; 10: 649

- 128 patients **admitted patients** with unexplained syncope
 - All get ECHO
- Result: # of abnormal ECHO studies=

o, Zero, Zip, nada, etc...

130

Holter Monitoring for Syncope: Diagnostic Yield in Different Patient Groups.

Kuhne M, et al. QJ Med 2007; 100: 771-7.

• Age group (yrs)	# pts. without structural HD	Diagnostic Holter (%)
<40	39	0 (0%)
40 - 50	29	0 (0%)
51 - 60	53	2 (4%)
61 - 70	59	2 (3%)
71 - 80	153	12 (8%)
81 - 90	100	7 (7%)
> 90	18	5 (28%)

131

“Old” ACC/AHA 2006 Statement.. On Tilt-table Testing

- **Sensitivity 26%-80%**
- **Specificity is approximately 90%.**

“In patients with a negative evaluation, ie, no evidence of ischemia and a structurally normal heart, the pretest probability that the diagnosis is neurocardiogenic syncope is high, so head-up tilt-table testing contributes little to establishing the diagnosis.”

In other words... in a patient with a normal evaluation who has a negative tilt table test, the most likely diagnosis is still neurocardiogenic syncope

132

Recurrent Vasovagal Syncope: Management

- Physical counterpressure maneuvers
 - PC Trial: 223 pts - decreased syncope episodes (2006)
- Increased fluid and salt intake
- Midodrine 5mg tid

These do NOT work...

- Beta-blockers (see POST trial, Circulation, 2006)
- fludrocortisone (POST 2 trial, J Am Coll Cardiol. 2016)
- pacemaker (*providing not prolonged asystole*)
(see Vasovagal Pacemaker Study II, JAMA, 2003)

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Note on Pacemakers:

1) Useful in (Spontaneous) Brady/Asystole

- Methods: 77 pts with either:
 - >3 sec asystole + syncope or
 - > 6 sec without syncope
 - All receive pacer, randomly assigned to:
- Results:

	<u>Pacer "on"</u>	<u>Pacer "off"</u>
syncope in 2 yrs	25%	57%

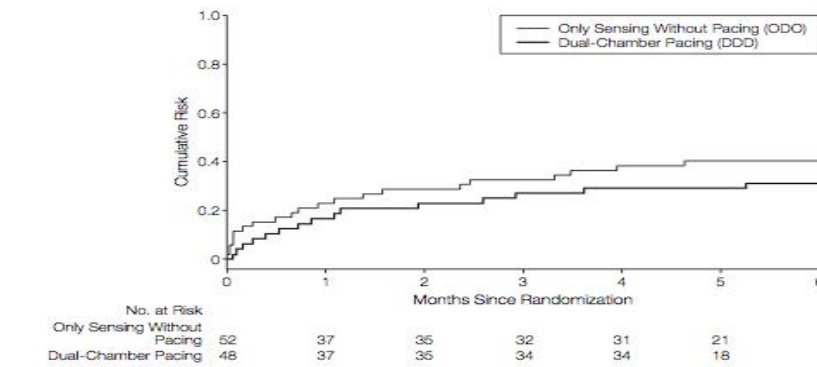
ISSUE-3 trial, Circulation, published ahead of print, 5/7/12

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Note on Pacemakers:

- 1) Useful in (Spontaneous) Brady/Asystole
- 2) Not Useful in Vasovagal Syncope

Figure 2. Time to First Recurrence of Syncope



*VPS II trial
JAMA 2003
100 patients*

Relative risk reduction of 30.2% (95% confidence interval, -33.2% to 63.4%; log-rank $P = .14$).

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Excellent Reference



Europace (2013) **15**, 1389–1406
doi:10.1093/europace/eut272

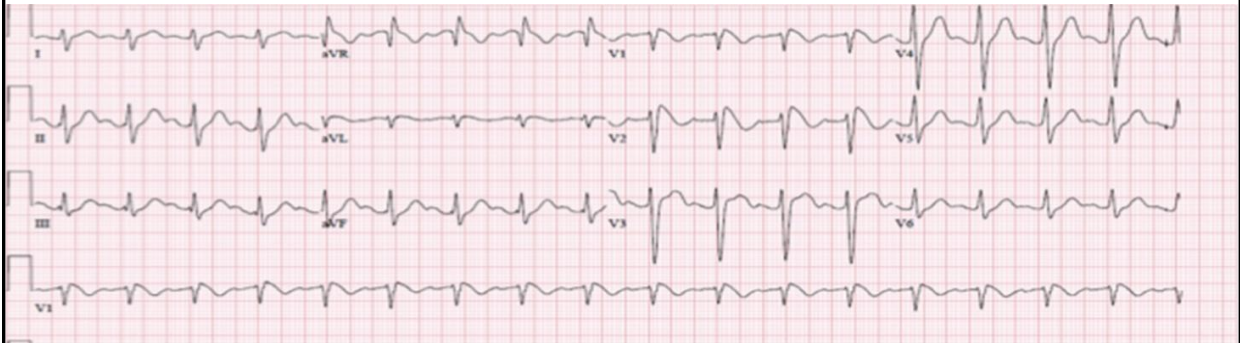
HRS/EHRA/APHRS CONSENSUS STATEMENT

Executive summary: HRS/EHRA/APHRS expert consensus statement on the diagnosis and management of patients with inherited primary arrhythmia syndromes

Silvia G. Priori, (HRS Chairperson)¹, Arthur A. Wilde, (EHRA Chairperson)², Minoru Horie, (APHRS Chairperson)³, Yongkeun Cho, (APHRS Chairperson)⁴, Elijah R. Behr⁵, Charles Berul⁶, Nico Blom^{7*}, Josep Brugada⁸, Chern-En Chiang⁹, Heikki Huikuri¹⁰, Prince Kannankeril^{11‡}, Andrew Krahn¹², Antoine Leenhardt¹³, Arthur Moss¹⁴, Peter J. Schwartz¹⁵, Wataru Shimizu¹⁶, Gordon Tomaselli^{17†}, Cynthia Tracy¹⁸

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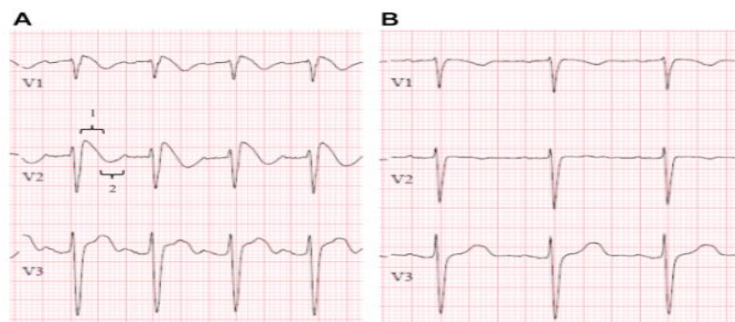
47-year-old Male Unresponsive, Found with Empty Bottles of Amitriptyline and Cyclobenzaprine



Kim HS, et al Ann Emerg Med, May 2017

137

47-year-old Male Unresponsive, Found with Empty Bottles of Amitriptyline and Cyclobenzaprine



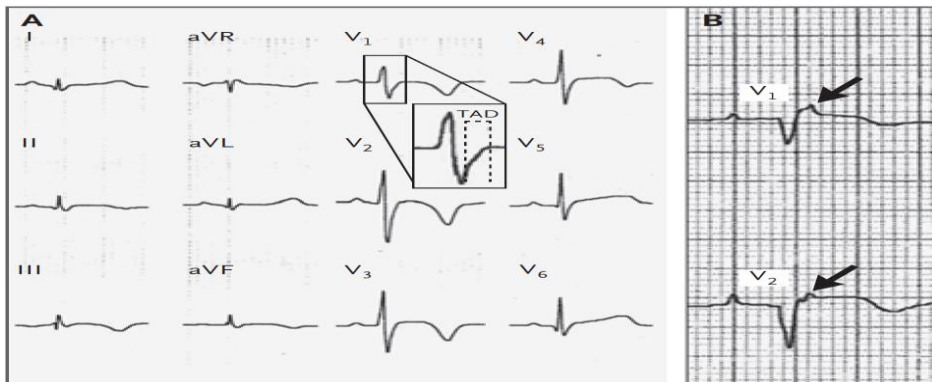
A. Before treatment

B. After Na bicarb

Kim HS, et al Ann Emerg Med, May 2017

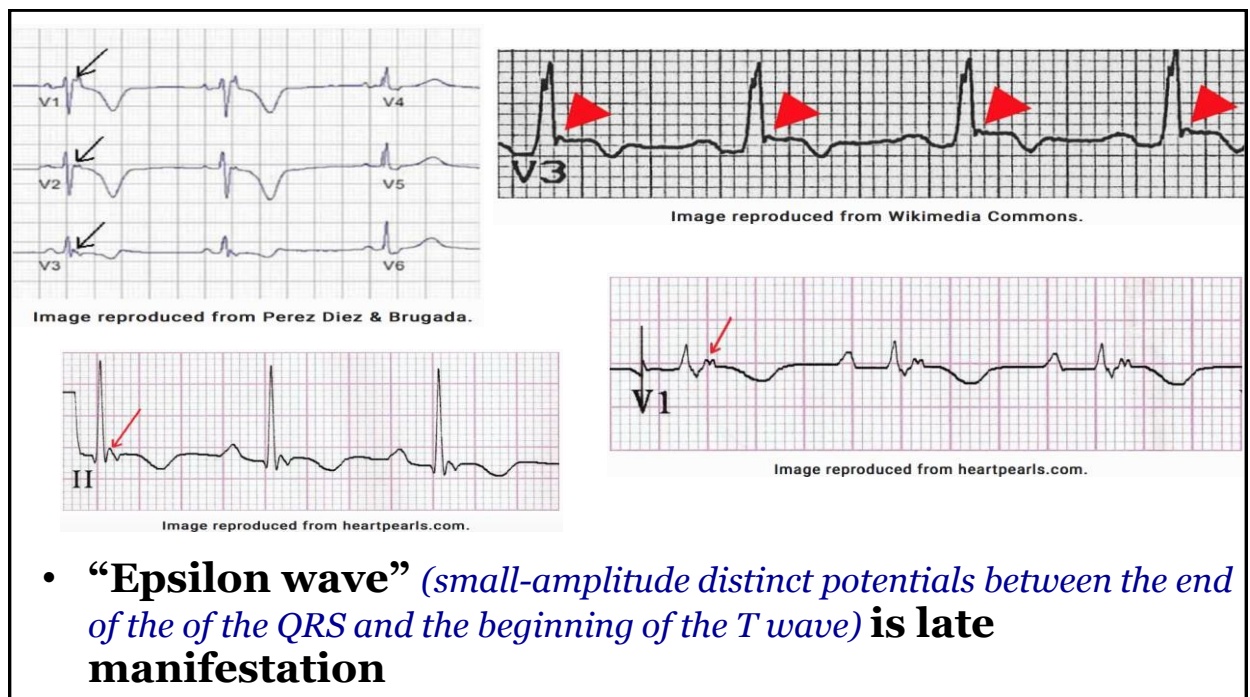
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Arrhythmogenic Right Ventricular Cardiomyopathy (ARVC)



- **“Epsilon wave”** (small-amplitude distinct potentials between the end of the of the QRS and the beginning of the T wave) **is late manifestation**

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- **“Epsilon wave”** (small-amplitude distinct potentials between the end of the of the QRS and the beginning of the T wave) **is late manifestation**

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Antipsychotics and QTc

