

QLSC 600

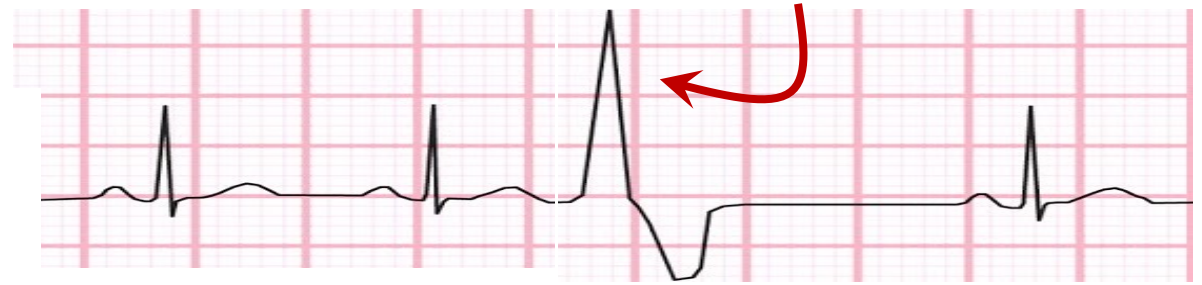
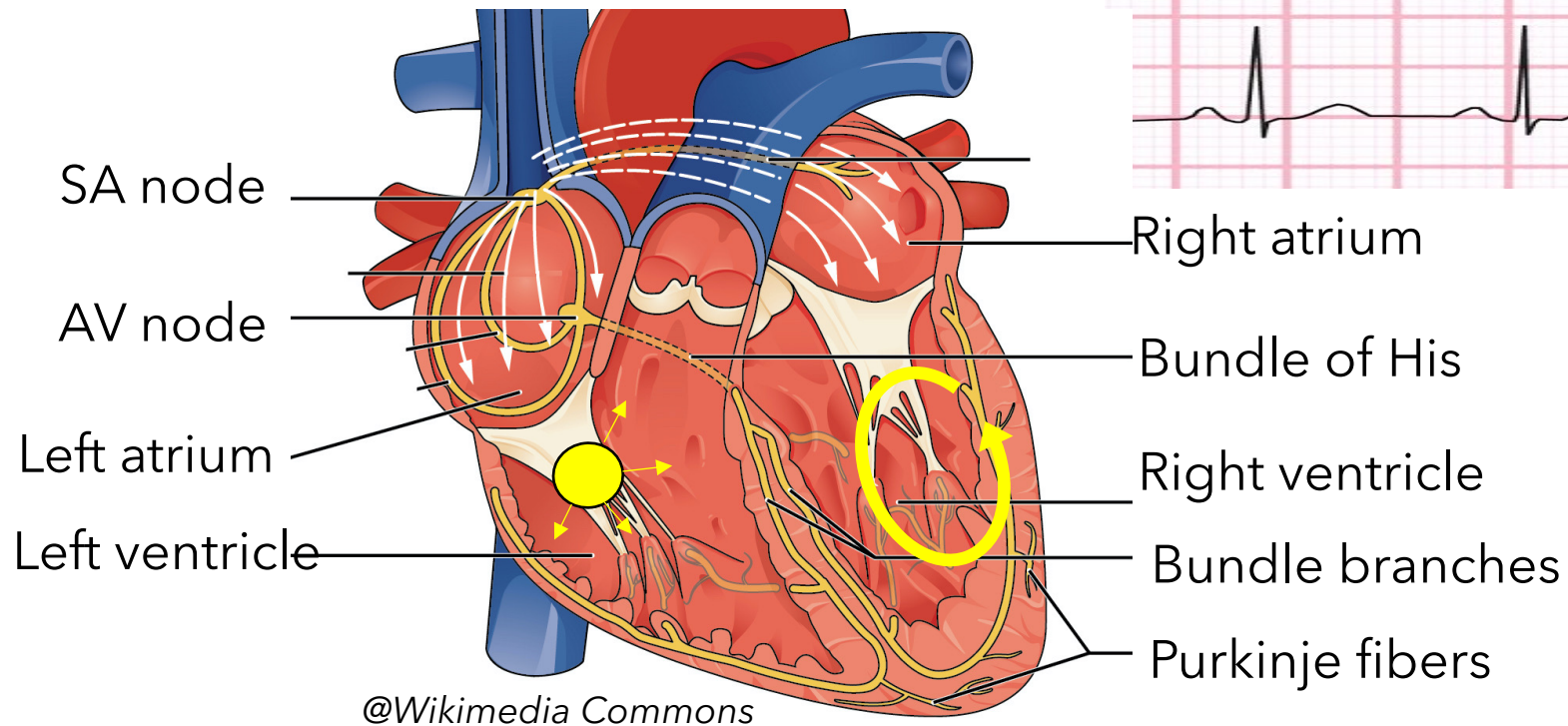
# Cardiac arrhythmia

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Instructors: Thomas Bury, Leon Glass

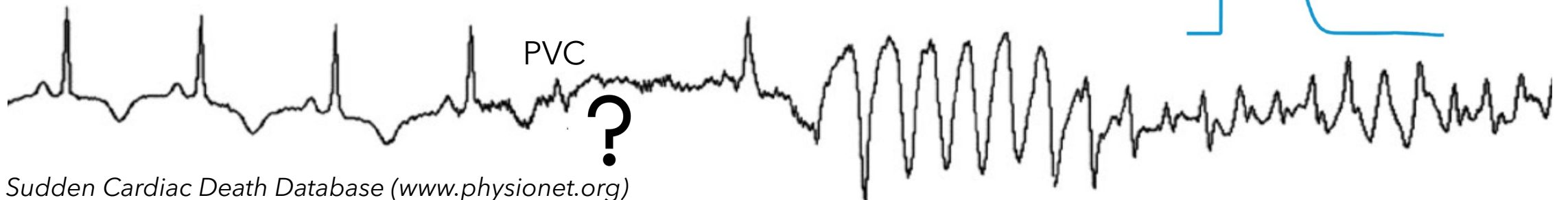
# Ventricular arrhythmia

PVC = Premature Ventricular Complex



## Initiation mechanisms

1. Ectopic pacemaker
2. Reentry
3. Triggered activity



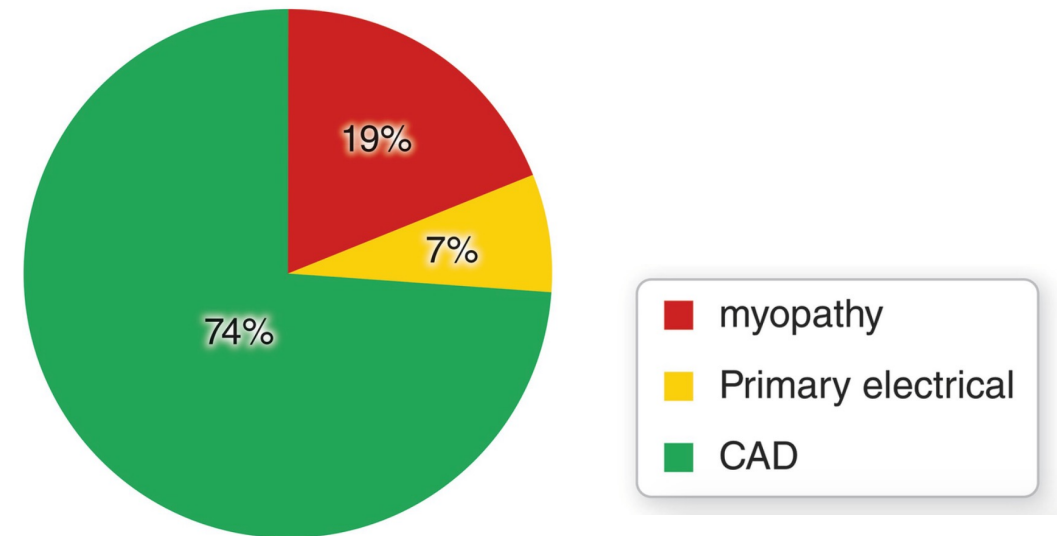
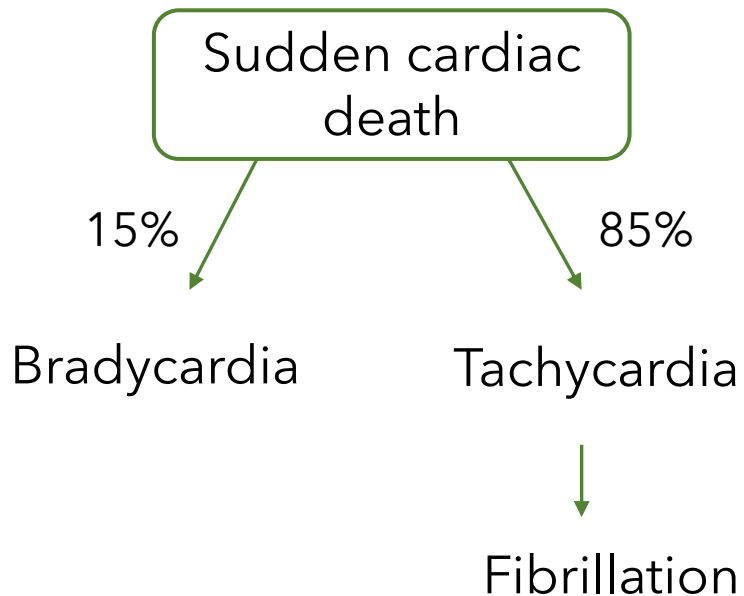
Sudden Cardiac Death Database ([www.physionet.org](http://www.physionet.org))

# Sudden cardiac death - epidemiology

- Cardiovascular disease is a leading cause of death (31% of all global deaths)
- 50% are cases of 'sudden cardiac death' - death within 1 hour after the onset of symptoms

The World Health Organisation

**Gerber et al.** Secular trends in deaths from cardiovascular diseases: a 25-year community study. *Circulation*.



**J. Anthony Gomes.** Heart Rhythm disorders: History, Mechanisms, and Management Perspectives, Springer.

# Why analyse PVC dynamics?

- Frequent PVCs show increased risk of cardiac death

**Dukes et al.** Ventricular ectopy as a predictor of heart failure and death. JACC (2015)

- Ventricular fibrillation is often preceded by frequent PVCs

**Deyell et al.** Sudden Cardiac Death Risk Stratification. Circulation research. (2015)

- PVC dynamics in patients are rich but poorly understood – still cannot reliably infer mechanism from pattern
- Problem of risk stratification for sudden cardiac death

How can we analyse PVC dynamics?

# Holter recording

1947 - Norman Holter



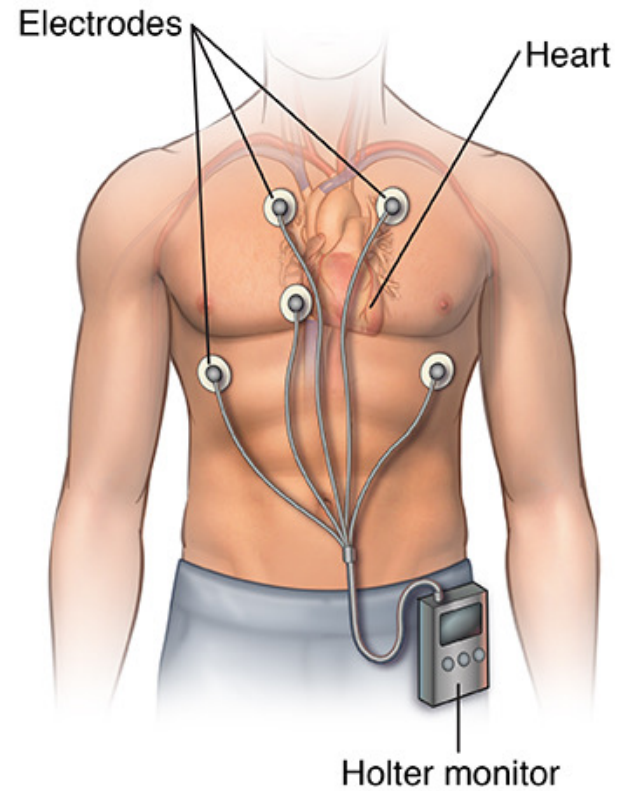
*Wikipedia.* The original Holter biotelemetry apparatus in 1947 weighing 85 lb

1954 - Briefcase recorder



*Image courtesy of National Museum of American History*

1960 - Tape recorder



2010 onwards:



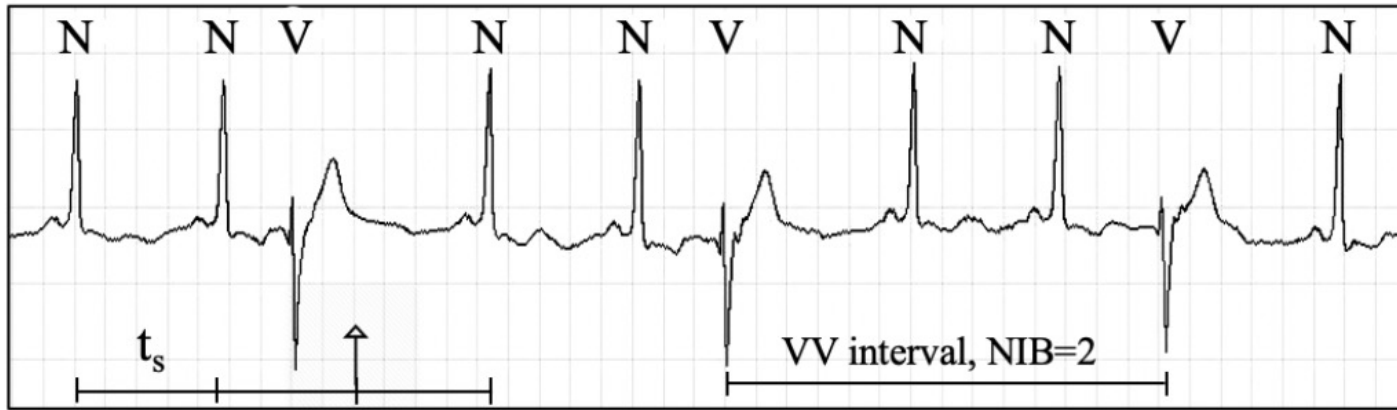
*Icentia cardioSTAT*



*Apple watch*

**H L Kennedy.** The history, science and innovation of Holter technology. *Annals of Noninvasive Electrocard.* (2006)

# ECG data processing

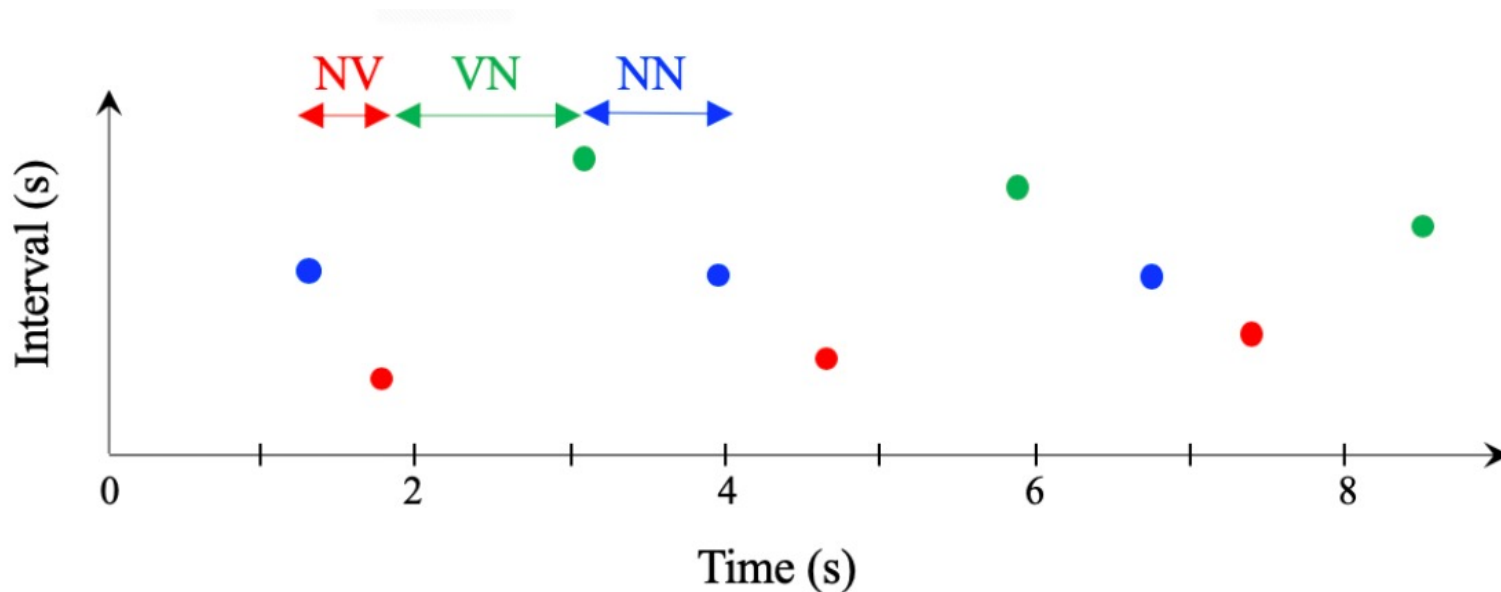


## Beat detection algorithm (Icentia)

N - normal (sinus) beat

V - premature ventricular complex (PVC)

Timing - precise to 250Hz



## Beat-to-beat interval plot

NV - coupling interval

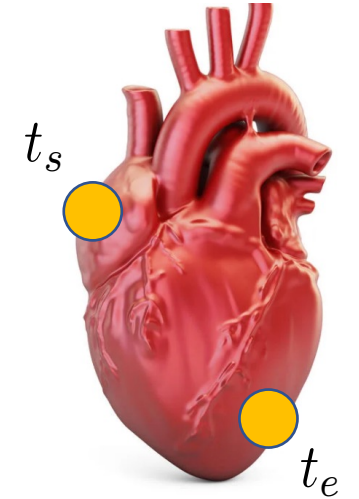
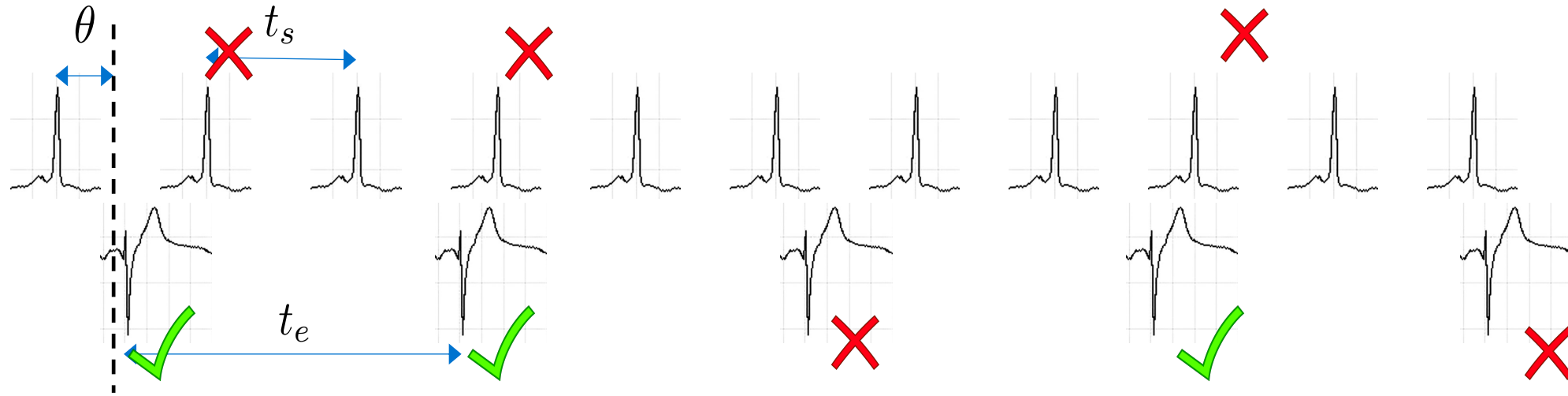
VN - compensatory pause

NN - sinus interval

## App demonstration

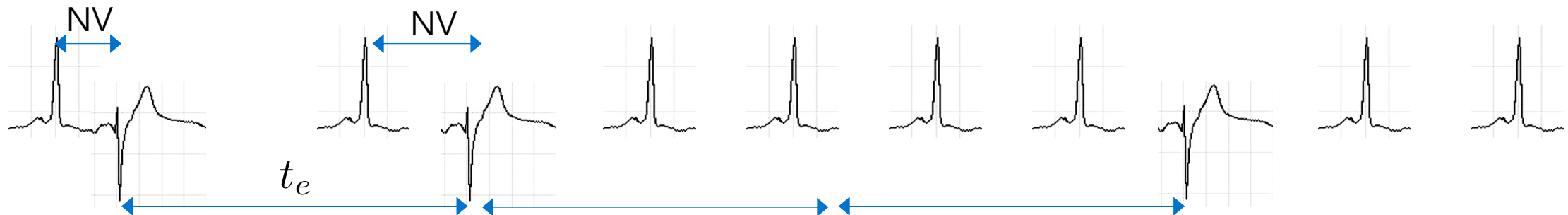
# Pure parasystole

- Ectopic pacemaker independent of sinus pacemaker



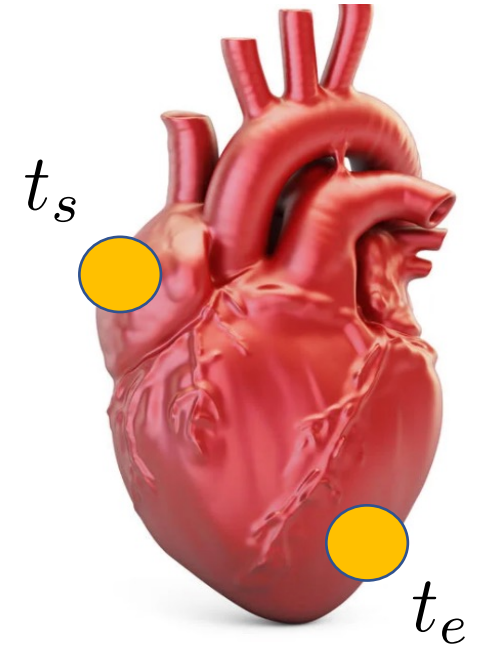
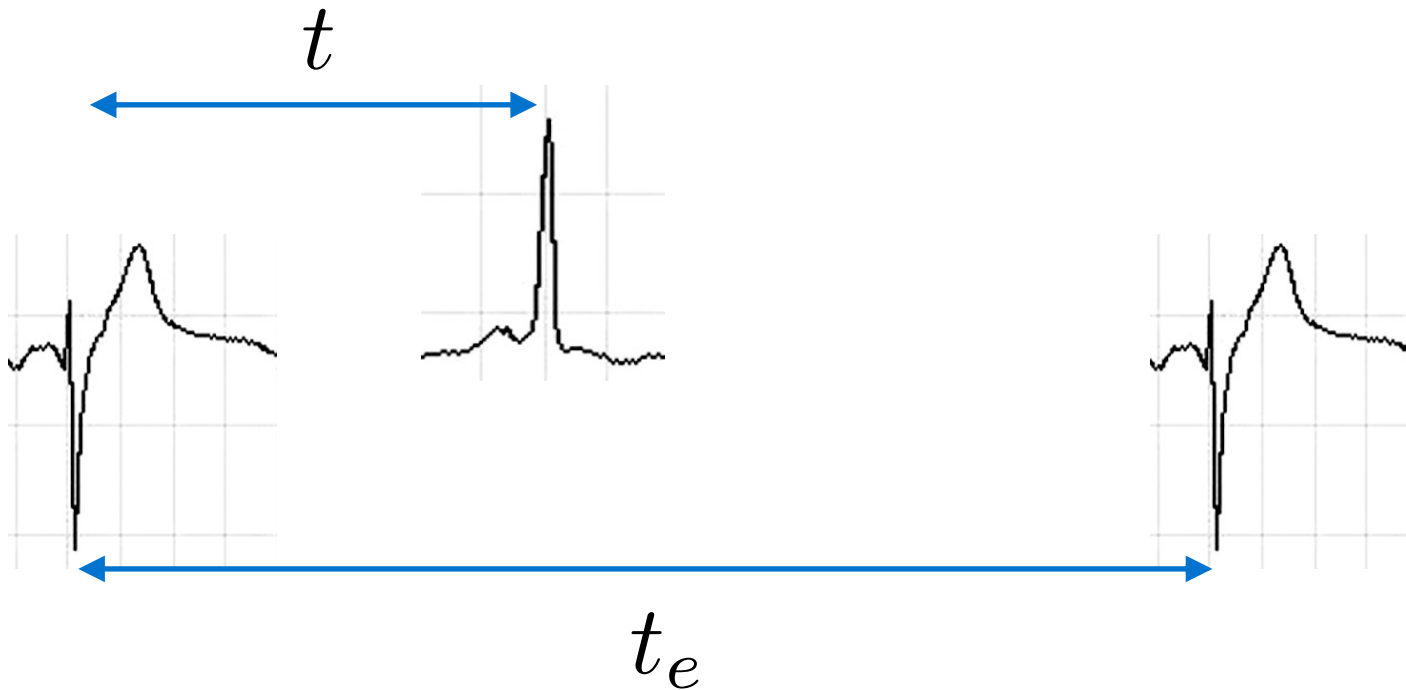
## Assumptions

- Ectopic beat blocked if it falls inside refractory period,  $\theta$
- Ectopic beat expressed if it falls outside of refractory period,  $\theta$
- Sinus beat following an expressed ectopic beat is blocked (compensatory pause)



# Difference equation for phase

$\phi_i$  Phase of the  $i^{\text{th}}$  sinus beat (expressed or not) in the ectopic cycle



$$\phi = \frac{t}{t_e}$$

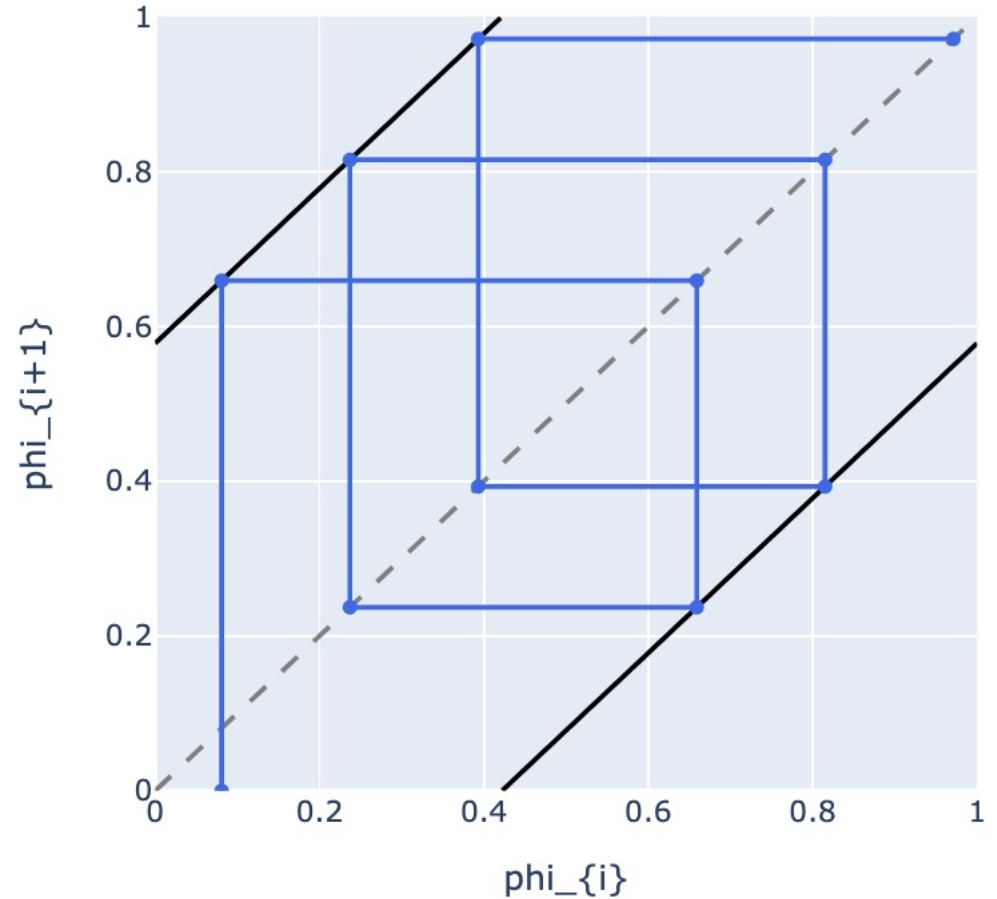
Find the expression for  $\phi_{i+1}$  in terms of  $\phi_i$  and make a cobweb plot.



$$\phi_{i+1} = \begin{cases} \phi_i + \frac{t_s}{t_e} & 0 \leq \phi_i < 1 - \frac{t_s}{t_e} \\ \phi_i + \frac{t_s}{t_e} - 1 & 1 - \frac{t_s}{t_e} \leq \phi_i < 1 \end{cases}$$

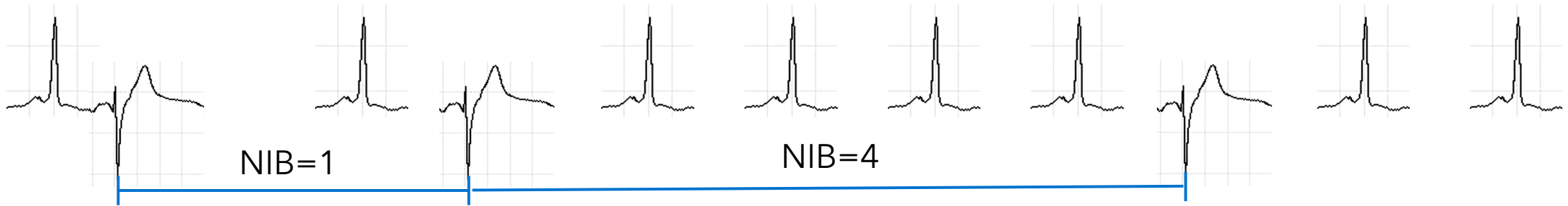
Or put more simply...

$$\phi_{i+1} = \phi_i + \frac{t_s}{t_e} \pmod{1}$$



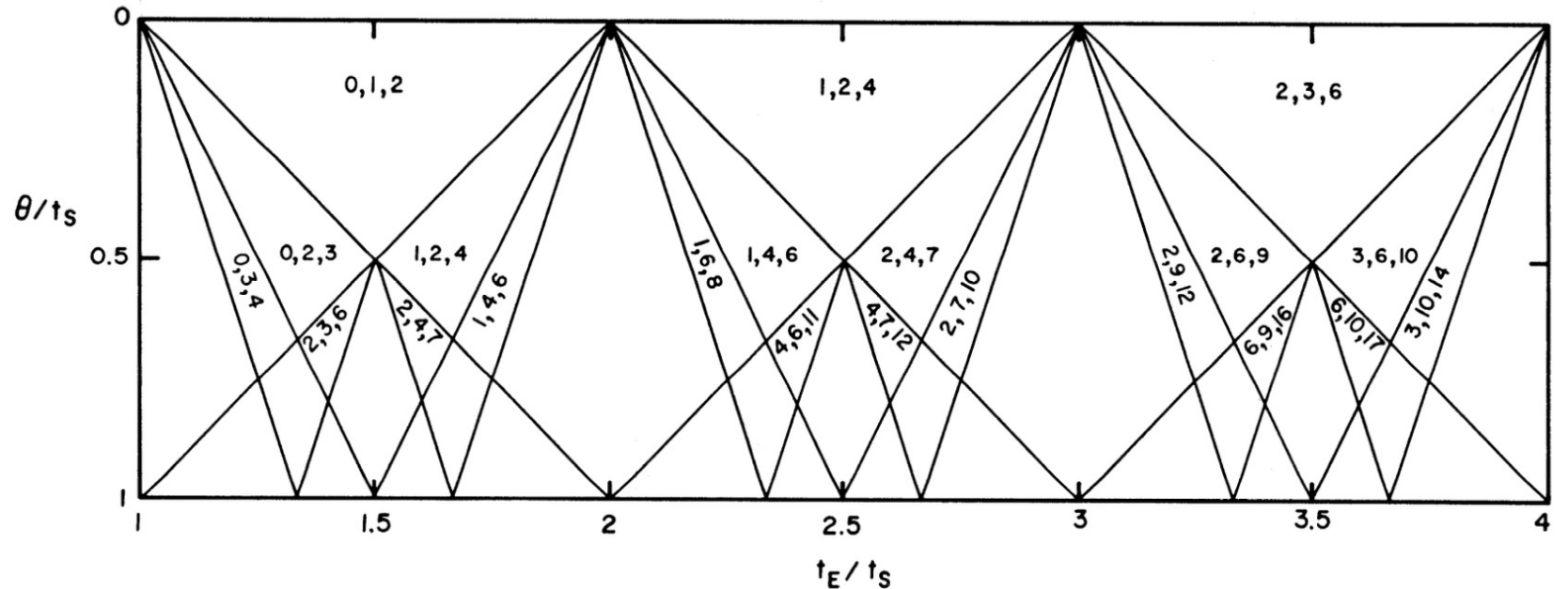
# Pure parasystole

NIB: Number of intervening sinus beats between two ectopic beats



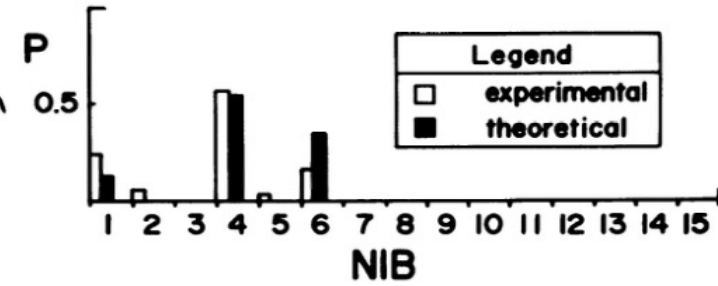
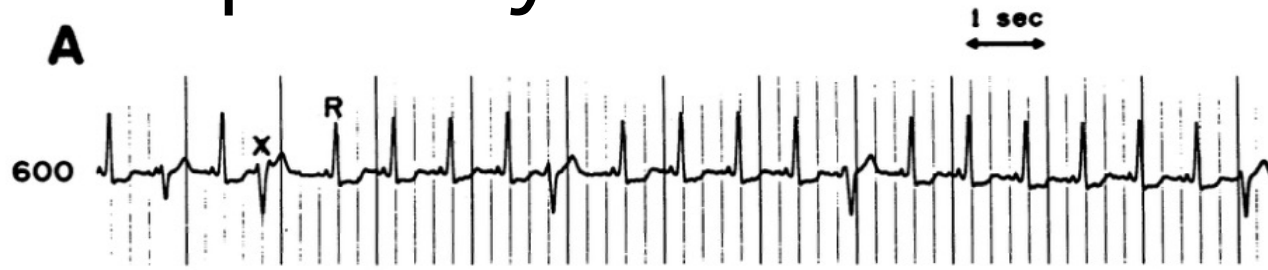
## Rules

1. Three different values
2. One of which is odd
3. Sum to two smaller values  $<$  largest value



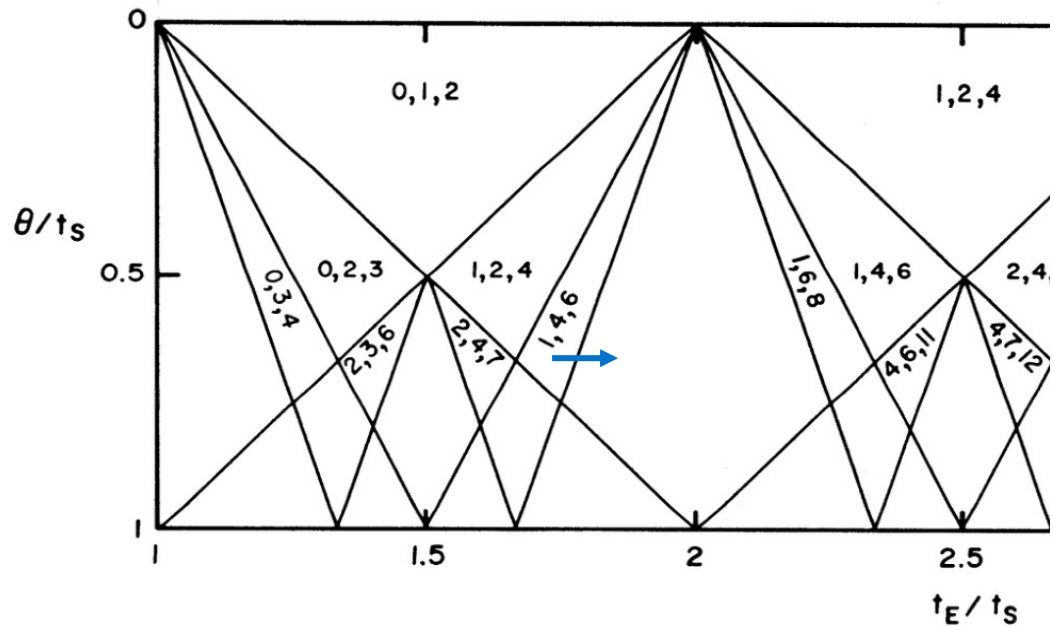
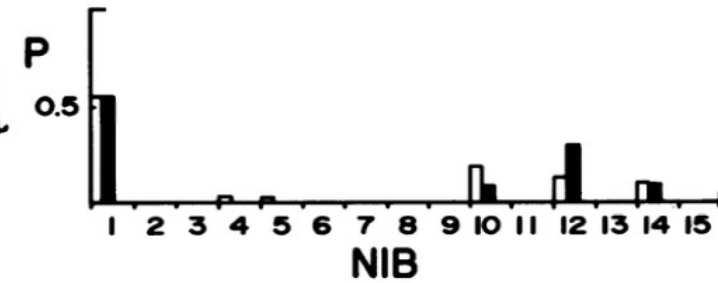
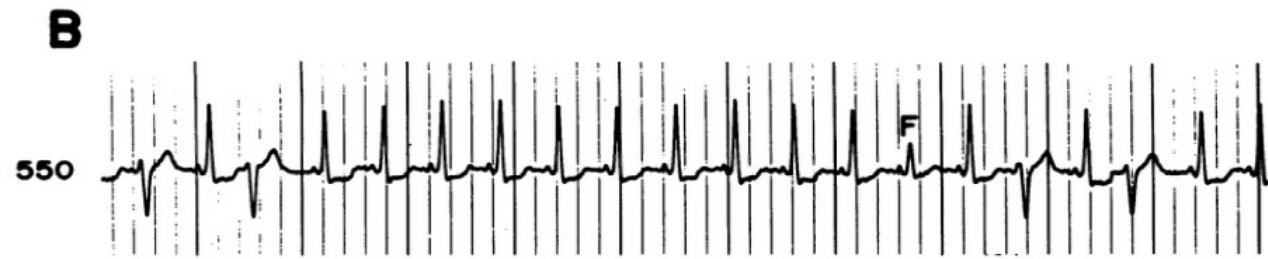
**Glass et al.**, Dynamics of pure parasystole. American Journal of Physiology (1986).

# Pure parasystole



$$\theta/t_s = 0.6$$

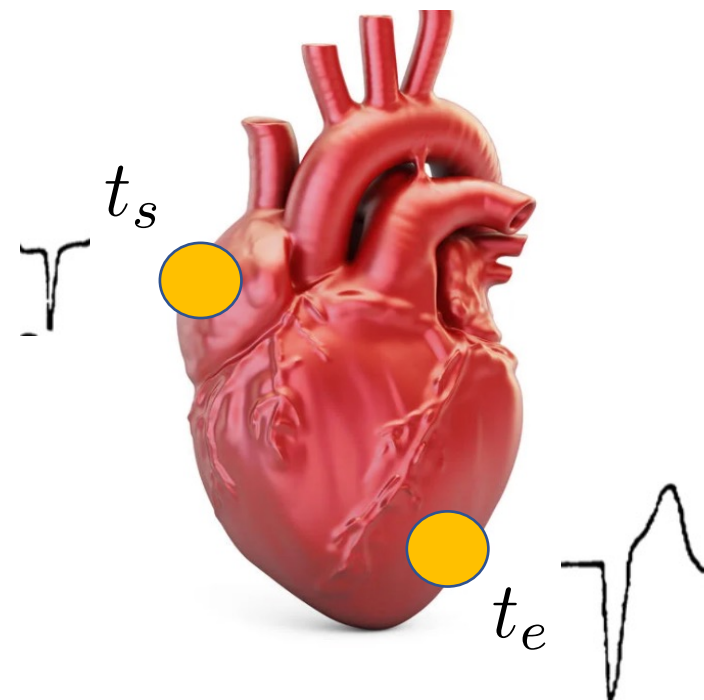
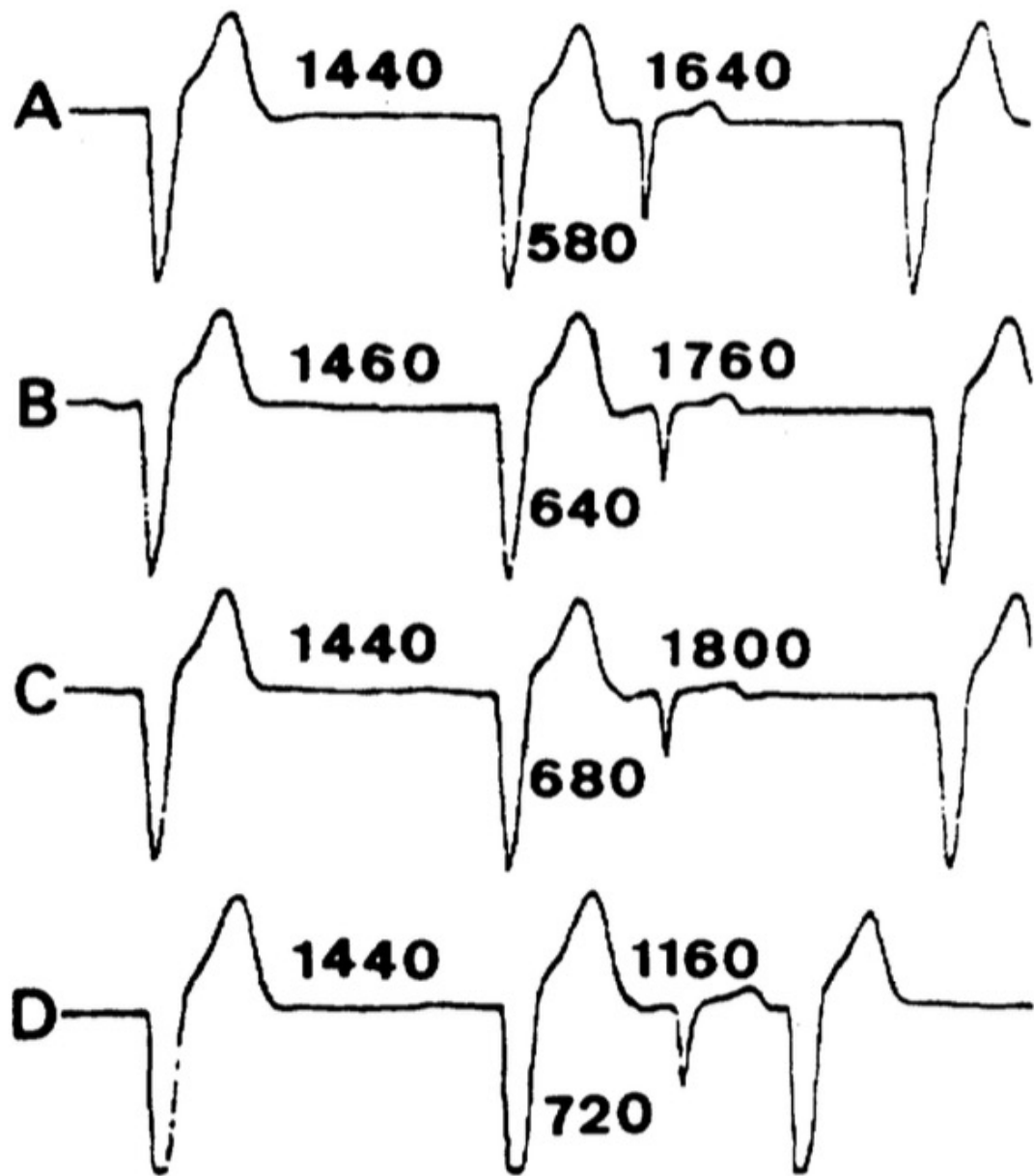
$$t_e = 1050\text{ms}$$



**Courtemanche et al.,** Beyond pure parasystole: promises and problems in modeling complex arrhythmias. American Journal of Physiology (1989)

## Factors not included

1. Modulation of ectopic focus
2. Conduction time to and from ectopic focus
3. Variable refractory period
4. Stochasticity in parameters



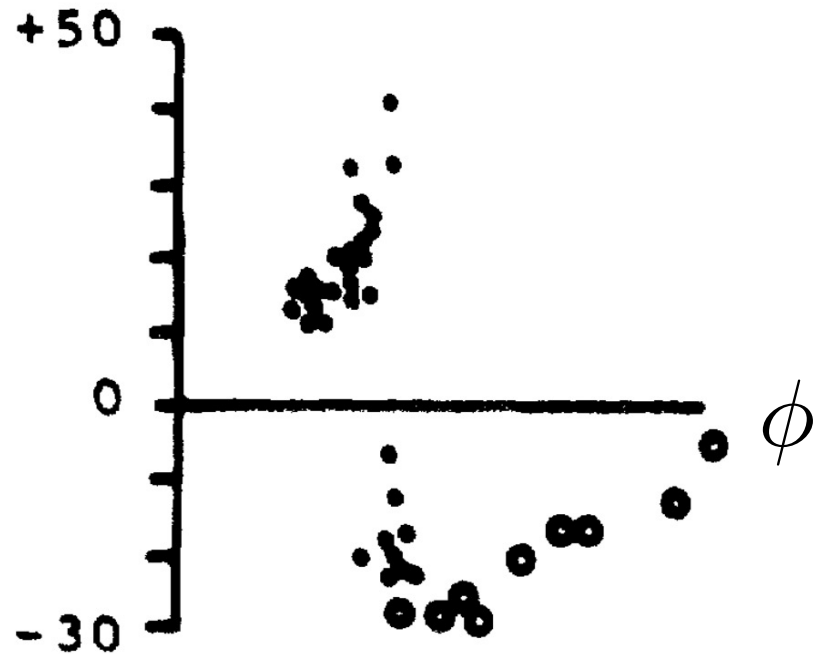
**Nau et al.,** Modulation of Parasystolic Activity by Nonparasystolic Beats. *Circulation* (1982)

# Modulated parasystole

**Pacemakers are influenced by external stimuli**

$\frac{T - t_e}{t_e}$  Normalized change in ectopic cycle length

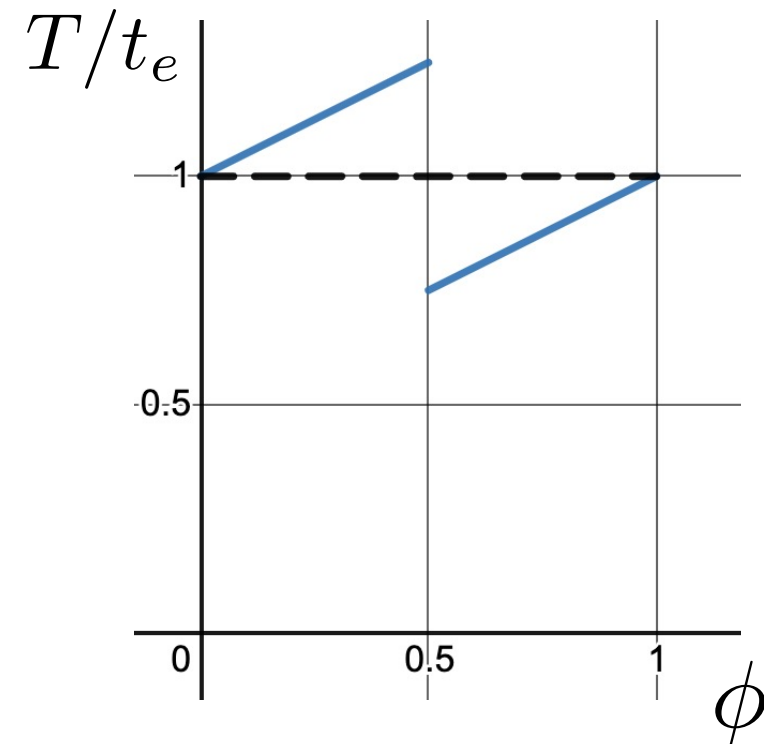
**X-R-X**



**Nau et al.**, Modulation of Parasystolic Activity by Nonparasystolic Beats. *Circulation* (1982)

$$\frac{T}{t_e} = \begin{cases} k\phi + 1 & 0 \leq \phi < 0.5 \\ k(\phi - 1) + 1 & 0.5 \leq \phi < 1 \end{cases}$$

$k$  is the strength of resetting

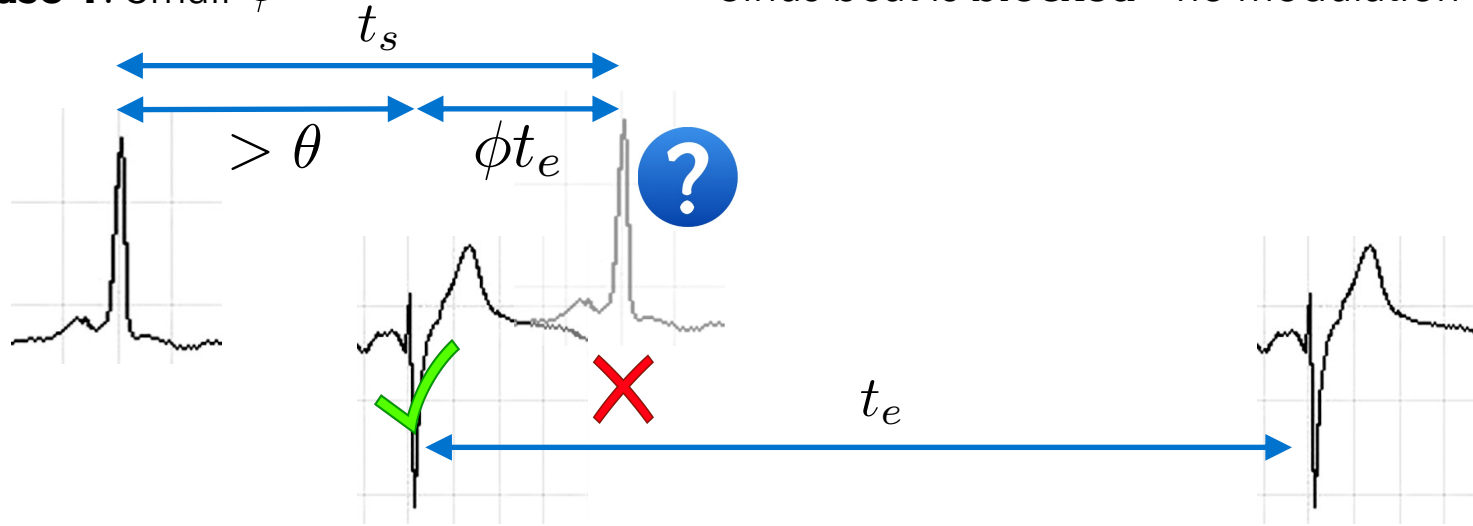


# Difference equation

Courtemanche et al., American Journal of Physiology (1989)

**Case 1:** Small  $\phi$

Sinus beat is **blocked** - no modulation of ectopic focus

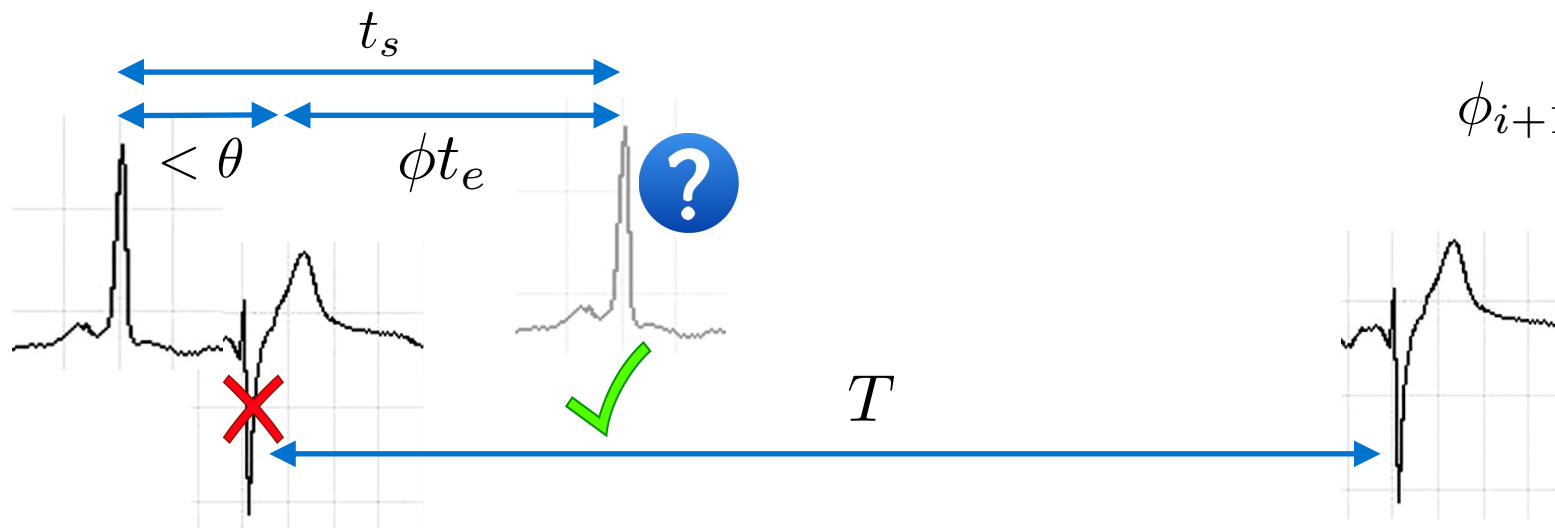


$$\phi_{i+1} = \phi_i + \frac{t_s}{t_e} \text{ mod } 1$$

$$0 \leq \phi_i < (t_s - \theta)/t_e$$

**Case 2:** Large  $\phi$

Sinus beat is **expressed** - modulation of ectopic focus



**Shift in phase due to modulation**

$$\phi_{i+1} = \phi_i + \frac{t_s}{t_e} + \boxed{1 - \frac{T(\phi_i)}{t_e}} \text{ mod } 1$$

$$(t_s - \theta)/t_e \leq \phi_i < 1$$

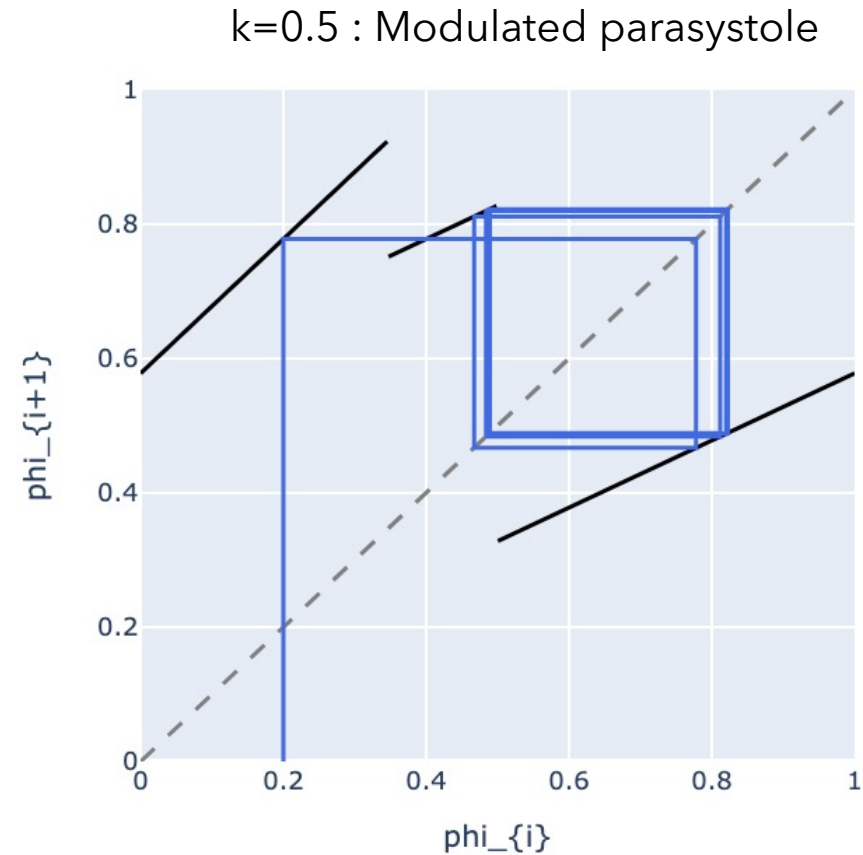
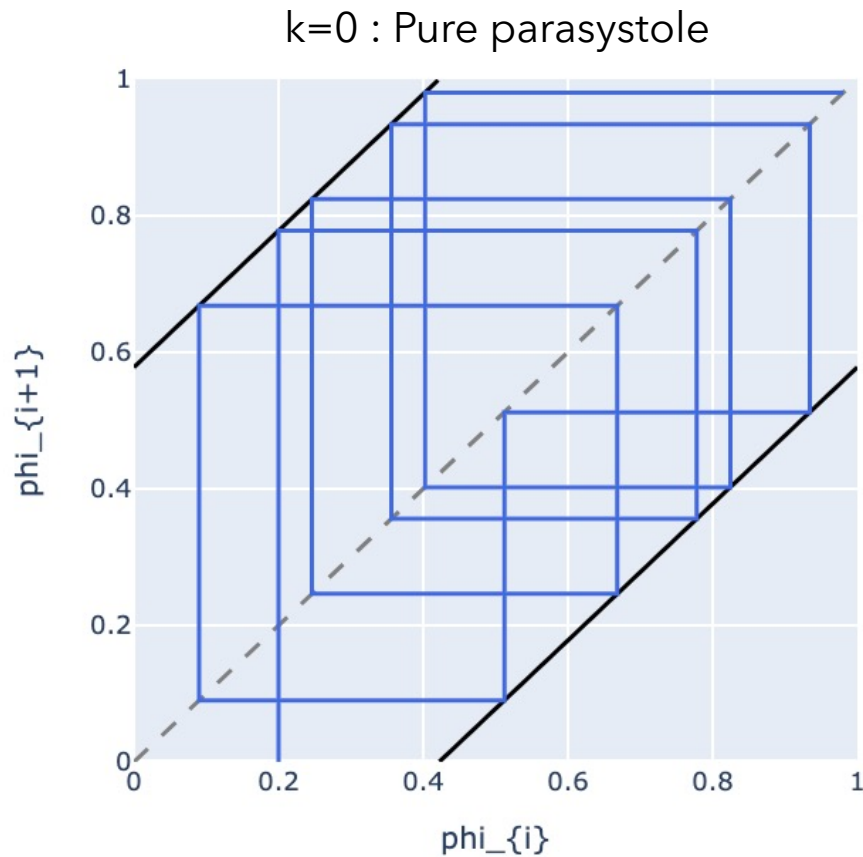
# Difference equation

Courtemanche et al., American Journal of Physiology (1989)

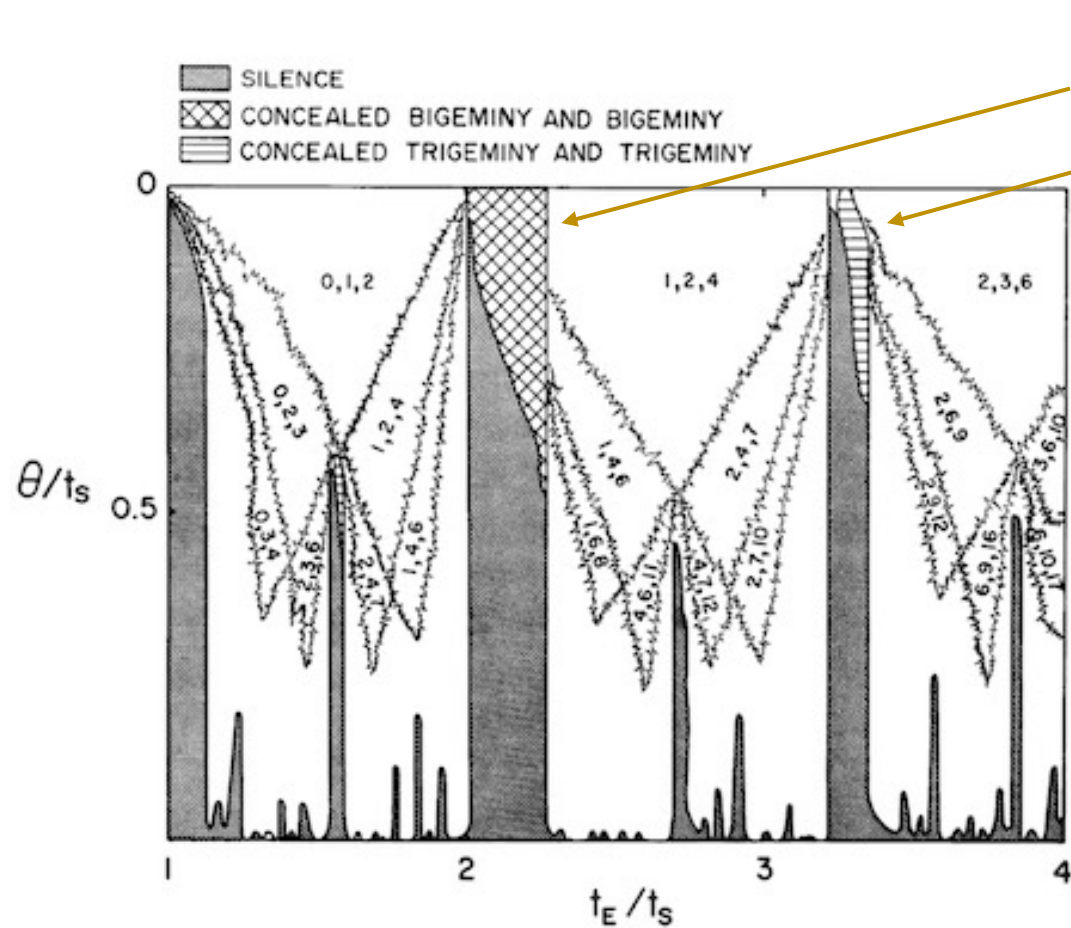
Investigate dynamics of modulated parasystole using online app  
<https://modulated-parasystole-cobweb.herokuapp.com/>

How does this differ qualitatively from pure parasystole (k=0)?

$$\phi_{i+1} = \begin{cases} \phi_i + \frac{t_s}{t_e} & 0 \leq \phi_i < \frac{t_s - \theta}{t_e}, \\ \phi_i + \frac{t_s}{t_e} + 1 - f(\phi_i) & \frac{t_s - \theta}{t_e} \leq \phi_i < 1, \end{cases}$$



# Entrainment



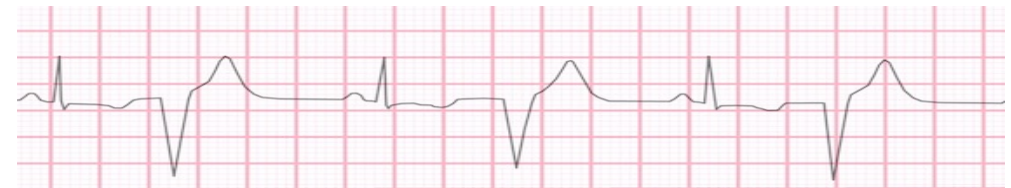
2:1 entrainment zone

3:1 entrainment zone

Modulated ectopic period is an integer multiple of sinus period

$$\phi_{i+1} = \begin{cases} \phi_i + \frac{t_s}{t_e} & 0 \leq \phi_i < \frac{t_s - \theta}{t_e}, \\ \phi_i + \frac{t_s}{t_e} + 1 - f(\phi_i) & \frac{t_s - \theta}{t_e} \leq \phi < 1, \end{cases}$$

## 2:1 entrainment



## 3:1 entrainment



**Courtemanche et al.**, American Journal of Physiology (1989)