

Practical Aspects of BP Assessment

Lawrence Satin, MD, FACC

Chief Medical Officer

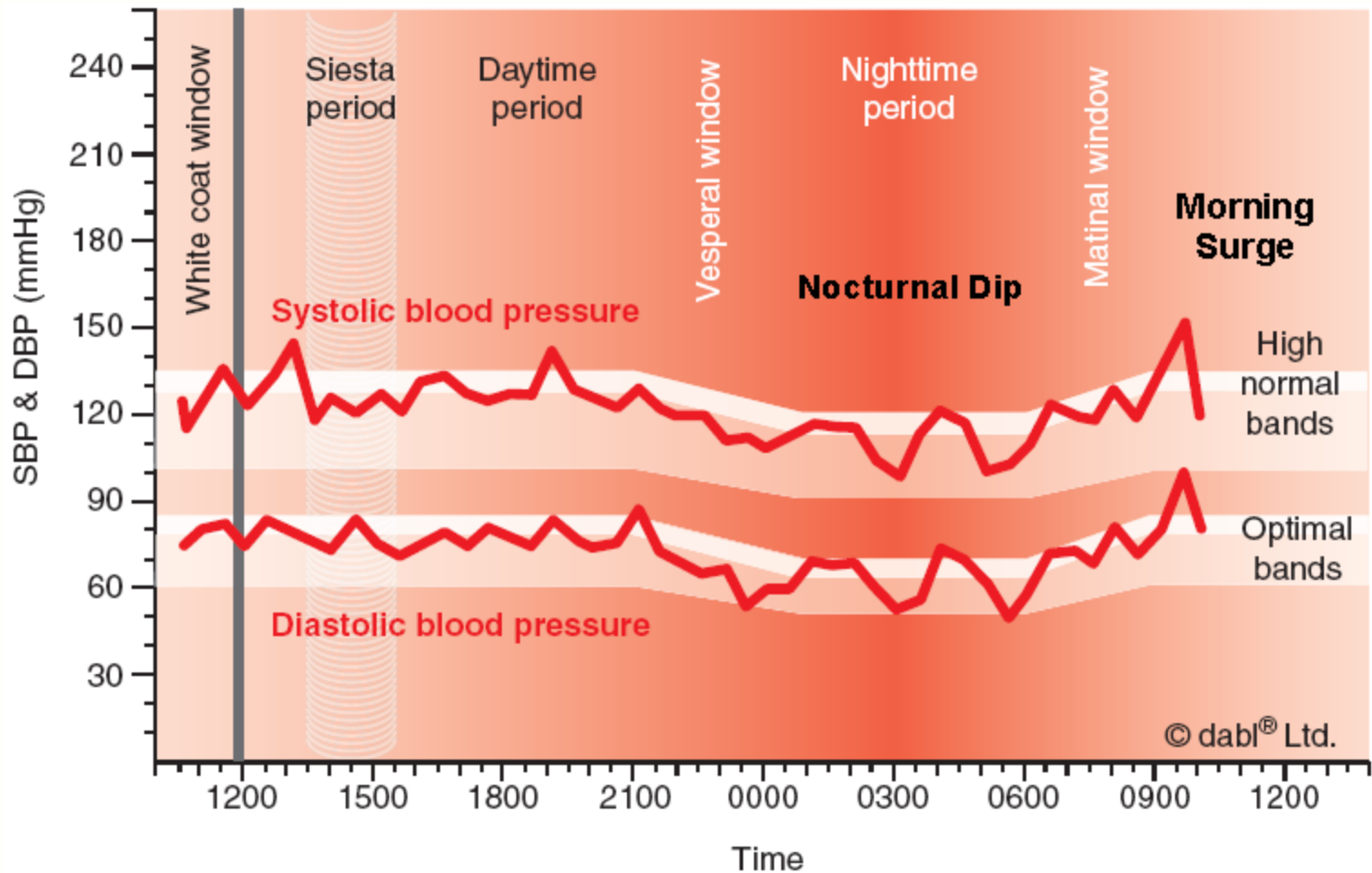
CSRC Annual Meeting - 2011

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What We Already Know

- ❑ **ABPM is better than Manual BP Cuff**
- ❑ **Small increases in BP can increase population CV risk**
 - BUT a shift in the mean population risk ...
 - Not the same risk for each individual
- ❑ **Circadian BP rhythm change and worse CV outcomes**
 - Excessive morning surge
 - Lack normal nocturnal fall in BP (non-dippers)
- ❑ **Drugs can induce different effects in different phases of the BP circadian rhythm**
 - ACE inhibitors super-dipping effect when dosed at night
 - Calcium channel blockers transform non-dippers to dippers
- ❑ **Central Aortic Pressure vs Peripheral Brachial Pressure**
 - Beta blocker less effective ↓ CV events vs calcium channel blocker

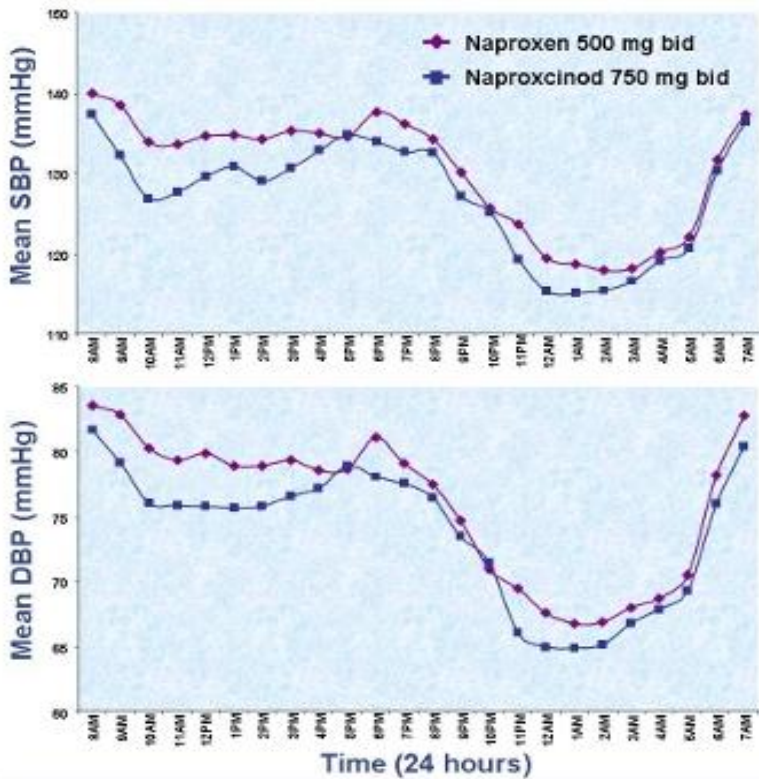
The Value-Add of ABPM



Can ABPM Ever Make a Difference vs Manual BP?

Naproxen vs Naproxcinod

(NSAID vs Cyclo-oxygenase Nitric Oxide Donor)

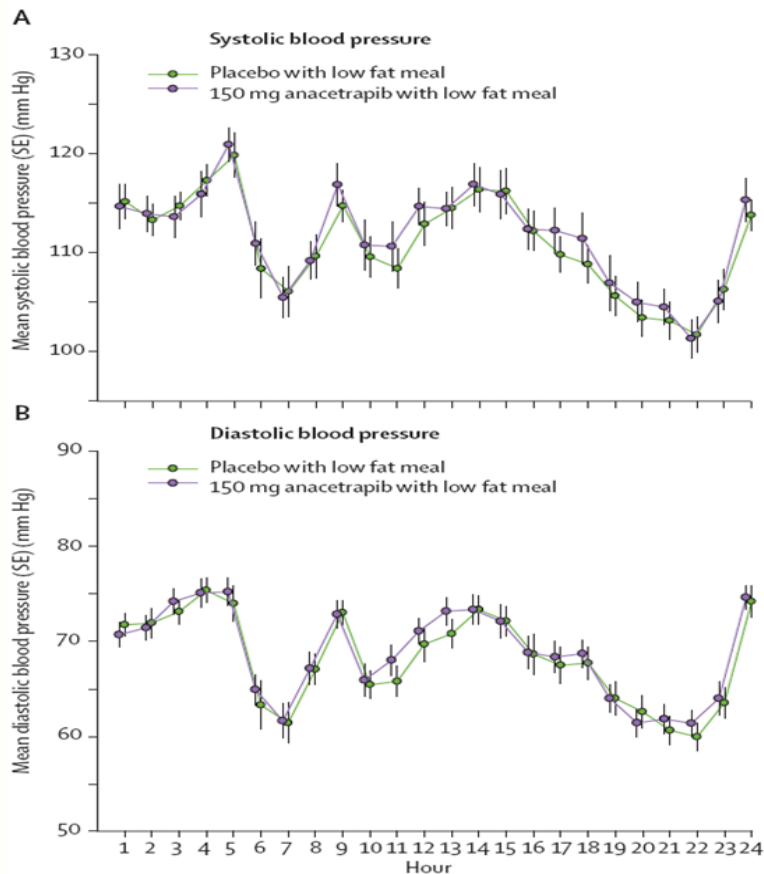


Mean systolic and diastolic blood pressure over 24 h

Naproxcinod approved by EMA 2010, FDA awaiting further studies.

Torcetrapib vs Anacetrapib

(Cholesteryl Ester Transfer Protein Inhibition)



Torcetrapib trials stopped 2006 with 60% increase in deaths, only 1 of 6 trials showed increase in BP. Anacetrapib 2009 DEFINE no increase in BP or MACE. Above, Krishna, et al. Lancet, Dec 2007, v370, p 1907-14. Merck 2 yr follow-up due Dec 2012.

What We DO NOT Know

- ❑ Do all drugs that increase BP do so in a concentration dependent manner?
- ❑ What is the magnitude and duration from onset of dosing for increased BP to constitute a CV risk?
- ❑ Can SHORT TERM studies evaluate BP risk in NHVs?
- ❑ How helpful is pre-clinical evaluation of BP risk?
- ❑ Should we also be looking at physiologic changes?
 - Cardiac output
 - Peripheral vascular resistance
 - Central vs brachial peripheral blood pressure curves