Biomarker Based Strategies for Exclusion of Acute MI: A U.S. Perspective

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Populations Undergoing Troponin Testing Differ around the World

How does patient selection for testing affect diagnostic performance of hs-cTn?

What might happen with 99th percentile ULN

| MI probability | Positive tests with standard assay (per 1000 patients) | Positive tests with hs assay | Additional positive tests meeting MI definition | Additional positive tests not meeting MI definition |
|----------------|--|------------------------------|---|---|
| 17% | 199 | 328 | 21 | 108 |
| 10% | 146 | 275 | 12 | 117 |
| 5% | 108 | 237 | 8 | 121 |
| 3% | 93 | 222 | 3 | 126 |

de Lemos, J.A., D.A. Morrow, and C.R. deFilippi, *Highly sensitive troponin assays and the cardiology community: a love/hate relationship?* Clin Chem, 2011. **57**(6): p. 826-9.

Implementation Strategy

Leadership Team

Laboratory Medicine Emergency Medicine Hospitalists Cardiology Administration

Protocol Development

Develop hs-cTn algorithm
Start with published data
Consider risk stratification
Develop processes for
Comprehensive education
Secondary testing
Plan for ↑cTn not from MI
Consultation guidelines

Careful implementation

Beta testing of algorithm Created order sets Provided clinical support

Monitor outcomes

Safety
Efficacy
Secondary Testing
Cost





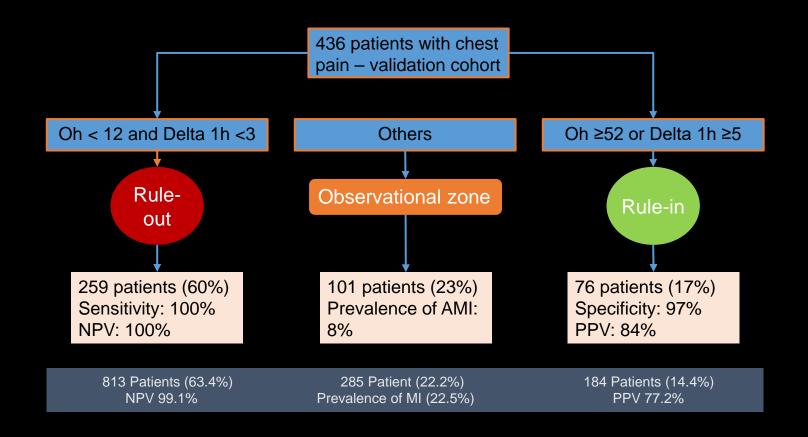
0 hour Rule Out

| Author, year | hs-cTnT threshold | N | % < LLD | Sensitivity (%) | NPV (%) |
|-----------------------------------|----------------------|------|---------|-----------------|------------|
| Body R. JACC 2011 | <3 ng/L | 703 | 28% | 100 | 100 |
| McRae A. Ann Emerg Med 2017 | <6 ng/L | 7130 | 42% | 99.8 | 99.9 |

Pros: Immediate R/O in low risk patients, validated

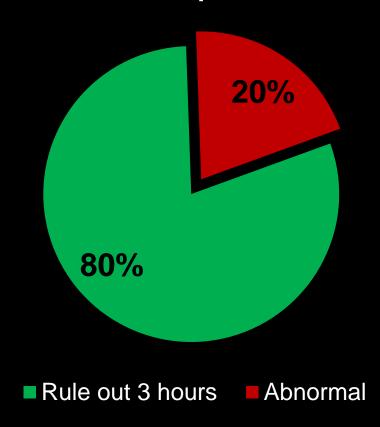
Cons: Applies to only 25-30% of patients, Not suitable for early presenters

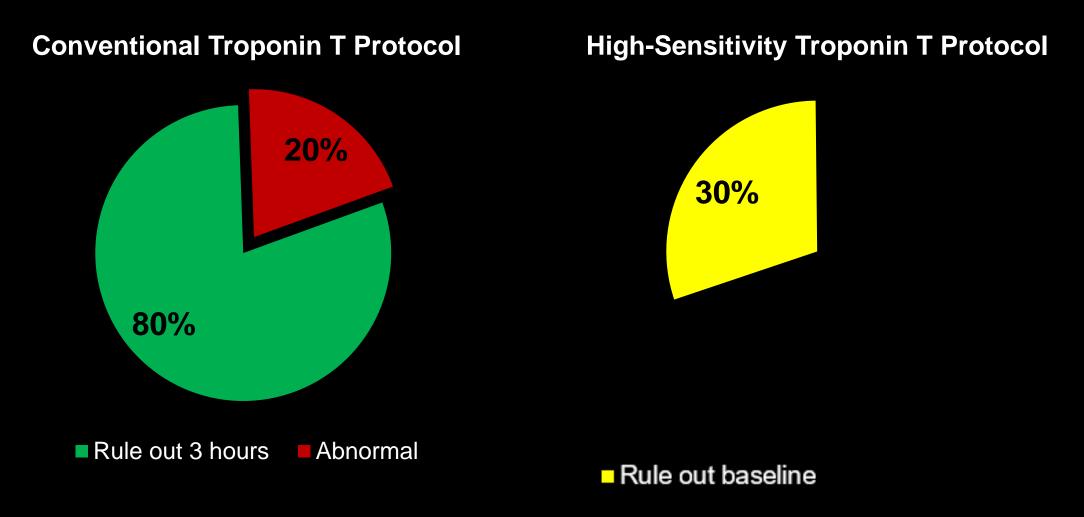
1-hour rule out with hs-cTnT

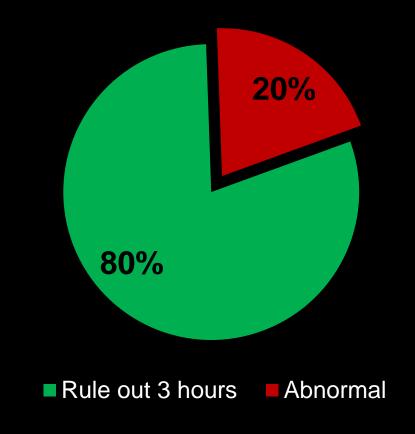


Trop T-hs >=52 ng/L at any time is abnormal

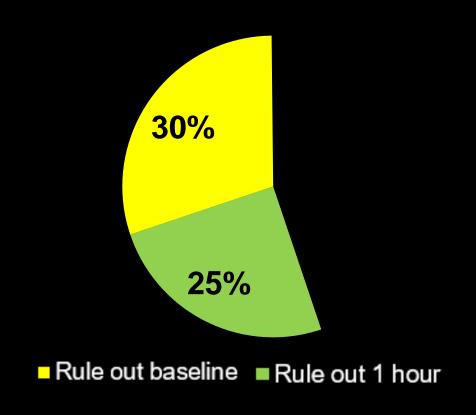
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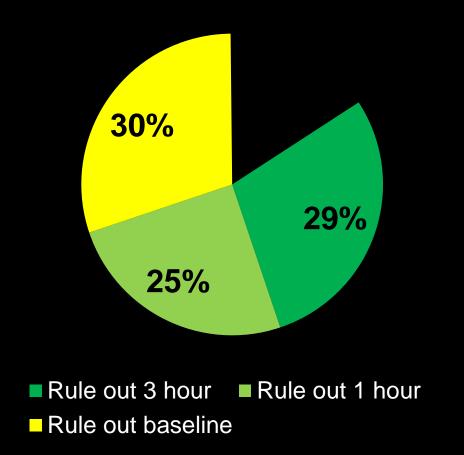


High-Sensitivity Troponin T Protocol



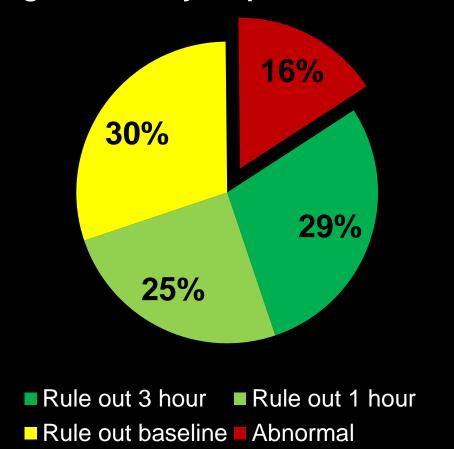
80% ■ Rule out 3 hours Abnormal

High-Sensitivity Troponin T Protocol

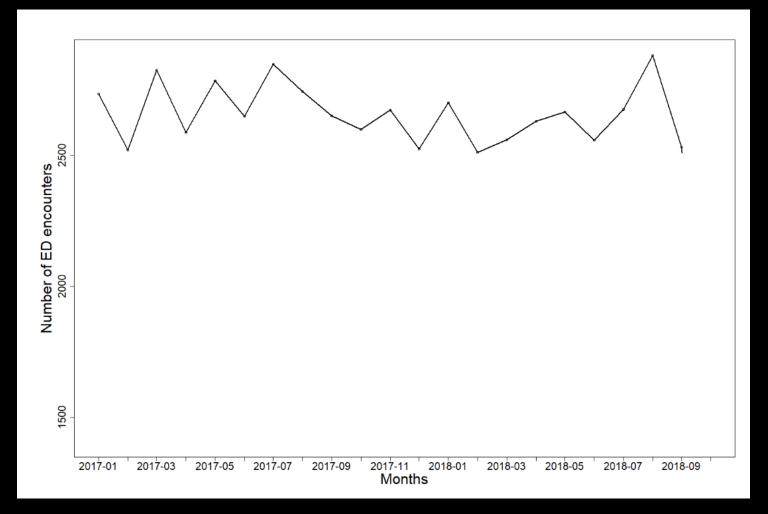


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High-Sensitivity Troponin T Protocol

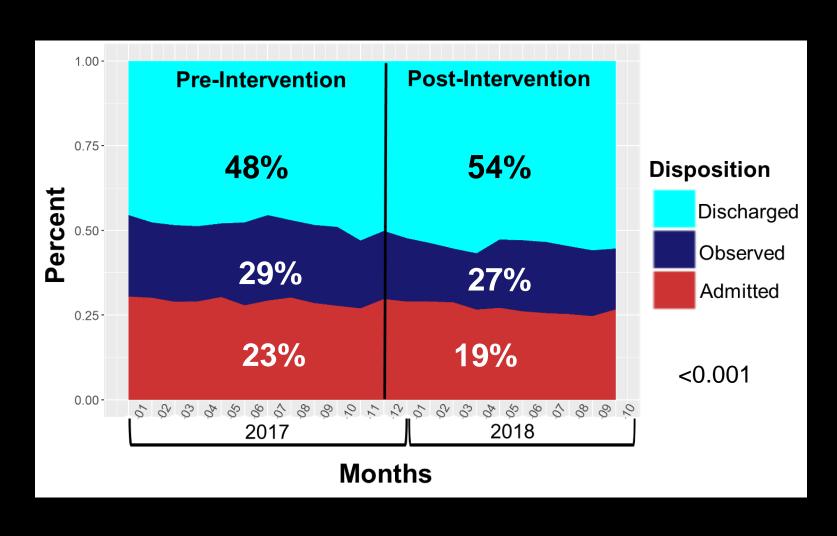


High rates of Cardiac Testing in Urban Safety Net Emergency Department

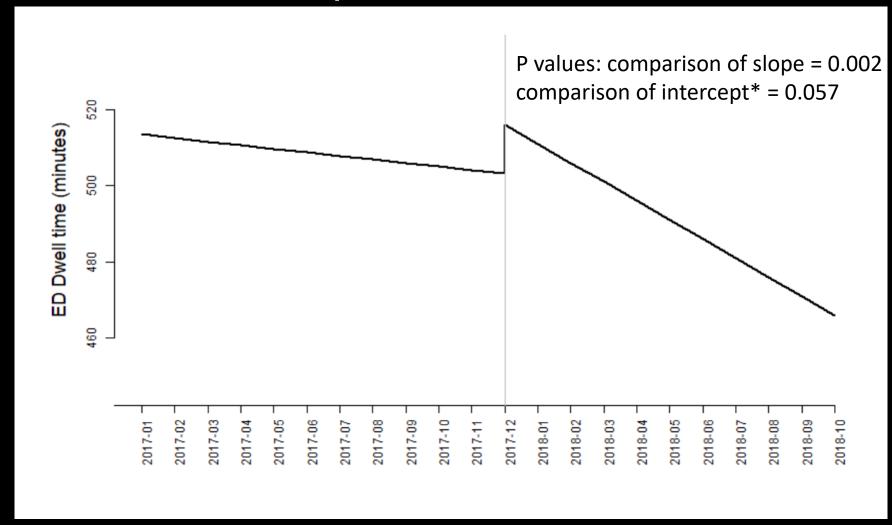


Mean of 2,603.5 encounters/month With ECG and troponin obtained

Proportion of encounters resulting in ED discharge, vs. observation vs. admission



ED Dwell Times Declined More Rapidly After Implementation



^{*} P value for comparison of intercept testing compared the end of the pre and beginning of the post implementation periods

Lessons Learned

- 99th URL will lower the PPV for MI in the US population
 - Potential for over-utilization of cardiac testing and costs
- Rapid 0/1/3 hour can potentially mitigate this issue by ruling out and risk stratifying a large portion
 - Additional safety data are needed
- Effects on resource utilization will vary based on the strategy implemented and the characteristics of the health system