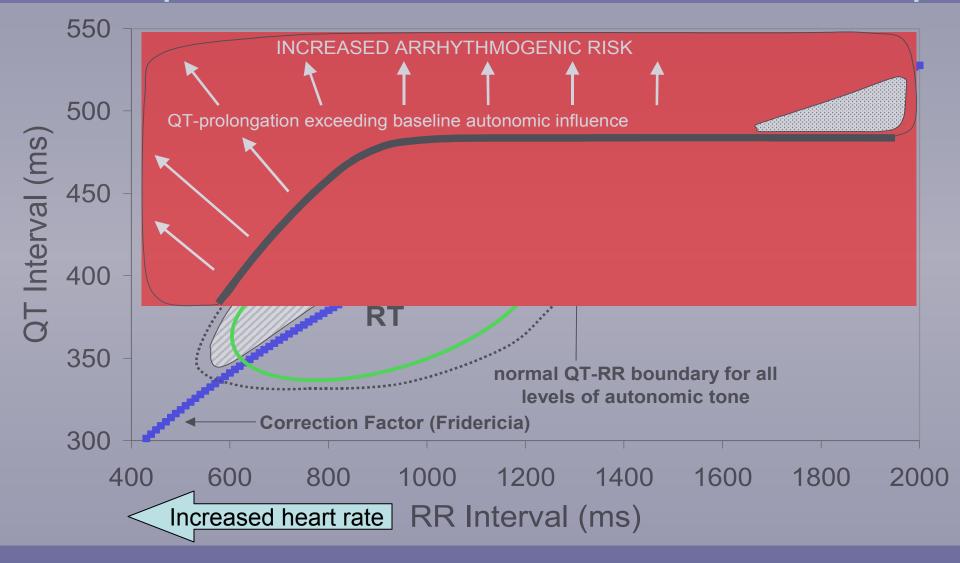
ECG restitution of the beat-to-beat QT-TQ intervals in humans. Can we begin assess arrhythmia vulnerability?

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Conceptualized view of QT-RR relationship

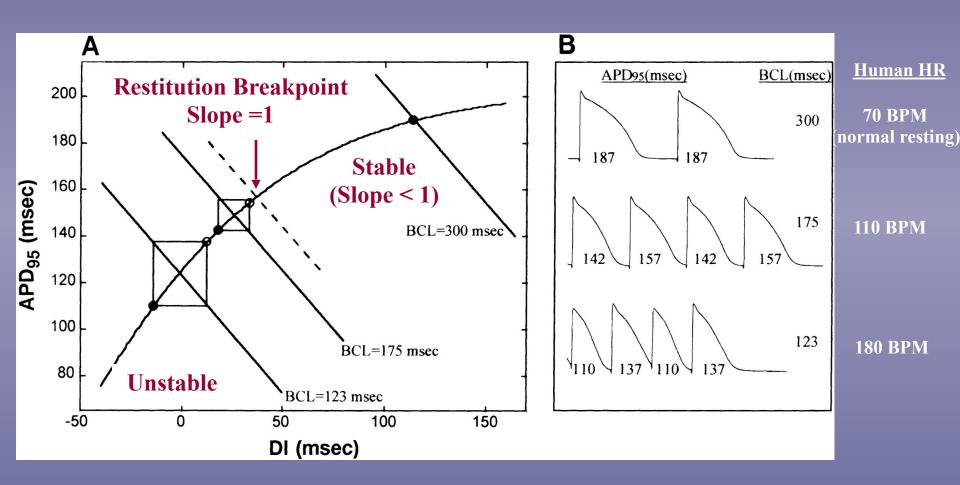


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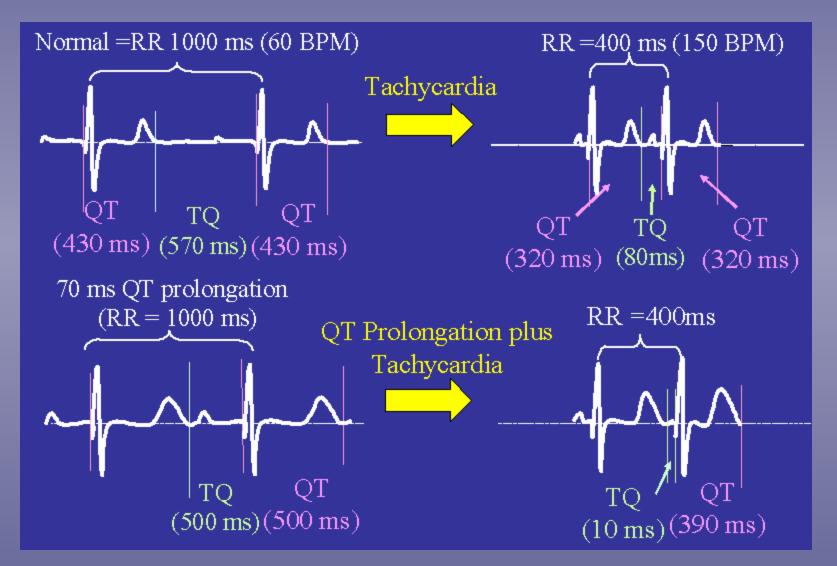
ECG Restitution

- Ability of the heart to recover from one beat to the next.
- Examines the relationship between action potential duration (QT interval) and diastolic interval (TQ interval) through the ECG.
- Replaces the traditional invasive electrophysiology procedure
- As QT/TQ ratio increases between beats this may be associated with increase arrhythmia vulnerability due to re-entry

Effects of heart rate on restitution and alternans



QT prolongation during tachycardia dramatically impairs restitution (increase QT/TQ ratio >1)



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Methods in normals and TdP

- Sequential beat-to-beat analyses of QT, TQ, and RR
- 22.5-hour Holter on 3 successive days
- Baseline, 160 (n= 38) or 320 (n=19) mg oral Sotalol

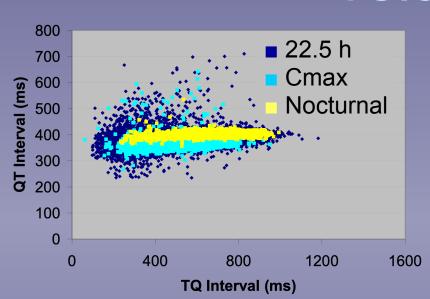
TdP Case Study

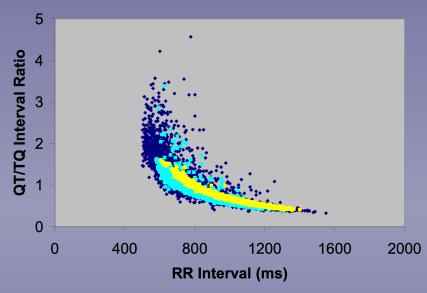
66 yo female CAD: Holter after sotalol (2 mg/kg, iv) prior to TdP

Restitution Parameters

- Lower TQ 5th quantile: boundary of lower 5% of beats
- % Beats with QT/TQ > 1: Reflects relative time spent in on steep portion of restitution relationship
- Upper 98% quantile of QT/TQ ratio: reflects degree and magnitude of steepness in restitution relationship for beats that may pose the greatest risk

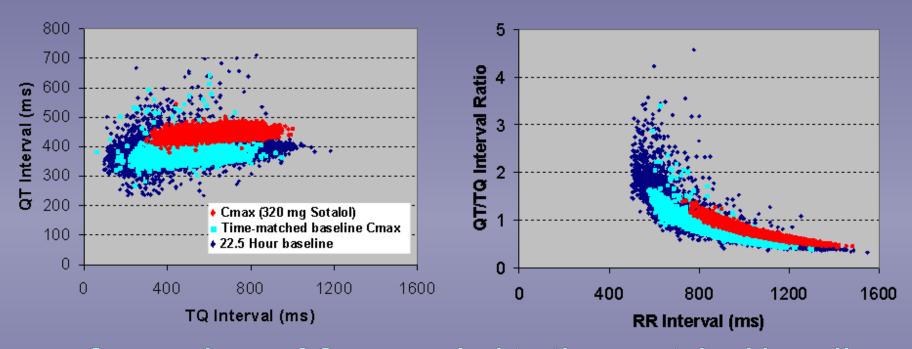
Baseline in normals volunteers





| Parameters | Baseline Assessment Periods (98% Confidence Bounds) | | |
|------------------------------------|---|------------------|-------------------|
| | 22.5 Hours | Cmax | Nocturnal |
| | | (Hours 2-4) | (Hours 19-21) |
| RR interval (ms) | 828 (794-858) | 827 (794-866) | 994 (947-1050)# |
| QT interval (ms) | 375 (366-383) | 373 (364-382) | 411 (401-420)# |
| QTc interval (Bazett) | 412 (405-420) | 409 (401-418) | 413 (405-422) |
| TQ interval (ms) | 456 (422-481) | 456 (425-488) | 586 (538-629)# |
| TQmin _{5th quantile} (ms) | 270 (252-287) | 298 (279-321) | 404 (369-442)# |
| %(QT/TQ ratio) > 1 | 25 (18-34) | 20 (13-32) | 6 (4-9)# |
| (QT/TQ ratio) _{max 98%} | 1.52 (1.43-1.62) | 1.41 (1.31-1.50) | 1.23 (1.13-1.34)# |
| quantile | | | |

Sotalol improves restitution despite QT prolongation



Comparison of Cmax period to time-matched baseline

RR: ↑ 235 ms

QT: 101 ms

TQ: 134 ms

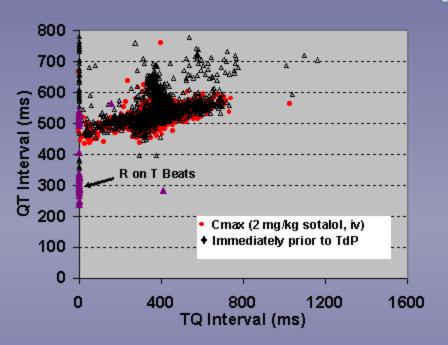
QTc: ↑ 52 ms

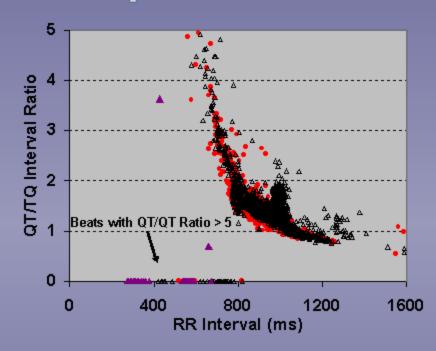
TQ 5th Quantile: QT/TQ 98th Quant: ↑ 115 ms ↓ 15%

%QT/TQ beats >1:

30%

Restitution is impaired prior to TdP





Comparison to Cmax period from normals

RR: ↓ 172 ms

QT: 1 53 ms

TQ: ↓ 225 ms

QTc: ↑ 95 ms

TQ 5th Quantile: 126 ms

QT/TQ 98th **Quant**: ↑ 58%

%QT/TQ beats >1: ↑ 722%

ECG Restitution Summary

- Can be obtained through digitized Holter recordings
- May be used to assess changes in autonomic state in conjunction with QT prolongation
- If boundaries can be defined, could delineate normal from abnormal repolarization status.

Other Contributors

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