

Cardiac Mechanics, Calcium Overload and Arrhythmogenesis

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Abstract— It is well-known that Ca^{2+} overload may cause cardiac arrhythmia. The results obtained in the model suggest that ectopic activity may emerge in a sub-critical myocardial region, e.g. comprising cardiomyocytes with moderately depressed $\text{N}^{+}\text{-K}^{+}$ pump.

Keywords— calcium overload, rhythm disturbances, cardiac mechanics

I. MOTIVATION AND AIM

A. Motivation

It is well-known that Ca^{2+} overload may cause cardiac arrhythmia. However, possible contribution of the mechanical factors to the arrhythmia development in Ca^{2+} -overloaded cardiomyocytes has been insufficiently addressed.

B. Aim.

Earlier we have developed a mathematical model of cardiomyocyte electro-mechanical function [1] that predicted a significant role of the intra- and extracellular mechanical factors in arrhythmogenesis. Model prediction was verified in experiments on papillary muscles from the right ventricle of guinea pigs overloaded with calcium [2].

II. METHODS

We utilized the cellular model to study effects of the electromechanical coupling between cardiomyocytes in a 1D heterogeneous muscle strand formed of 90% of normal (N) cardiomyocytes and 10% of sub-critical (SC) cardiomyocytes with decreased $\text{Na}^{+}\text{-K}^{+}$ pump activity. Single SC cardiomyocytes did not demonstrate spontaneous activity during isometric contractions at a reference length.

Regular fiber twitches at the reference initial cell length were induced by 1 bps electrical stimulation applied at an edge of the strand. Excitation spread along the tissue via electro-diffusional cell coupling followed by cell contractions and force development in the fiber.

III. RESULTS

Mechanical interactions between N- and SC-cells in the tissue resulted in the spontaneous activity emerged in the SC-zone between the regular stimuli. If the excitation wave spread from SC- to N-region, the SC-cells developed delayed after-depolarizations (DAD) that caused a slowly developing beat-to-beat decrease in the force of fiber contraction. If the excitation spread in opposite direction, DAD in the SC-cells induced reflected downward excitation waves capturing the normal region and followed by extrasystoles in the whole fiber

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