

# Transmural ventricular heterogeneities play a major role in determining T-wave morphology at different serum potassium levels

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**Background and aim:** End-stage renal disease (ESRD) affects more than 10% of the population. ESRD patients present impaired potassium homeostasis, which increases the risk for ventricular arrhythmias and sudden cardiac death. Preventive, noninvasive estimation of serum potassium,  $[K^+]$ , is therefore of major importance. We investigated three T-wave morphological descriptors, assessed their relationship with  $[K^+]$  and ascertained mechanisms responsible for their large inter-individual differences.

**Methods:** Electrocardiograms (ECGs) of twelve ESRD patients undergoing hemodialysis and ECGs calculated from simulated ventricular fibers were processed to quantify the T-wave width ( $T_w$ ), slope-to-amplitude ratio ( $T_{SA}$ ) and temporal morphological variability ( $d_w$ ).  $[K^+]$  was measured at different time points during hemodialysis and simulated from 2 to 8 mmol/l in the modeled ventricular fibers.

**Results:** In ESRD patients,  $T_w$ ,  $T_{SA}$  and  $d_w$  were closely related with  $[K^+]$  during hemodialysis, with median Spearman correlation coefficients of  $-0.5$ ,  $0.8$  and  $0.65$ , respectively. However, the pattern of such relationships depended highly on the characteristics of each patient. This variability, more manifest at high  $[K^+]$ , was reproduced in the simulations. Simulated descriptors were highly sensitive to the proportion of mid-myocardial cells, with 10% variations in this proportion leading to more than 15% changes in the T-wave descriptors.

**Conclusions:** Changes in  $[K^+]$  have remarkable effects on  $T_w$ ,  $T_{SA}$  and  $d_w$ , but the pattern of the relationship is highly patient-dependent, particularly under elevated  $[K^+]$ . Differences in the proportion of mid-myocardial cells may play a role in explaining such inter-individual variability.