

# Coupling of membrane nanodomain formation and enhanced electroporation near phase transition

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Biomembranes pass through phase transitions with a change in the environmental temperature. It was experimentally demonstrated that a transition not only involves membrane structural changes, but also a change in ion permeability [1,2]. At the main phase transition temperature  $T_m$ , the permeability displays a peak and current fluctuations of quantized amplitude can be measured.

Here, the temperature-dependent structural features of a one-component dipalmitoylphosphatidylcholine bilayer and its stability against an applied electric field was investigated at and close to  $T_m$  by means of atomistic molecular dynamics simulations [3]. The simulations demonstrate the dynamic appearance and disappearance of thin, interdigitating and thick, ordered lipid domains in fluid-like bilayers close to  $T_m$ , which vanished at higher temperatures. The structures were spatially related and likely represent precursors of the ripple phase forming below  $T_m$ . Similarly, a metastable two-phase bilayer consisting of a gel and a fluid domain adopted a thickness minimum at the phase interface formed by interdigitating and splaying lipids. Close to  $T_m$ , electroporation was enhanced with pores preferentially forming in the thin nanodomains. Together, these findings provide a link between the increased permeability and structural heterogeneity close to phase transition.

[1] D. Papahadjopoulos et al., BBA 311, 330-348 (1973).

[2] V. F. Antonov et al., Europ. Biophys. J. 34, 155-162 (2005).

[3] S.A. Kirsch and R.A. Böckmann, Biophys. J. 116, 2131-2148 (2019).