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Cross-correlation based methods for estimating the functional connectivity in cortical networks

Alessandro Ide*1,4, Michela Chiappalone^{1,2}, Luca Berdondini³, Vittorio Sanguineti⁴ and Sergio Martinoia¹

Address: ¹Neuroengineering and Bio-nano Technology group – NBT, DIBE, University of Genova, Genova, Italy, ²Italian Institute of Technology – IIT, Unit of Neuroscience and Brain Technology Genova, Italy, ³Sensors, Actuators and Microsystems Laboratory, IMT, University of Neuchâtel, Neuchâtel, Switzerland and ⁴Neurolab, DIST, University of Genova, Genova, Italy

Email: Alessandro Ide* - noriaki@dist.unige.it

* Corresponding author

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Introduction

Identification of the causal relationships between pairs of neurons plays an important role in the study of synaptic interactions within the nervous system at the population level. The simplest approach uses the cross-correlation function between pairs of spike trains. However, cross-correlograms cannot tell whether the observed peaks or troughs in the correlation function derive from either direct or indirect connections, or result from a common input. This limitation can be partly overcome with the notion of partial correlation or conditional firing probability [1,2].

Methods

Dissociated cortical neurons were obtained from rat embryos and plated into high-density microelectrode arrays (MEAs). Functional connectivity is estimated using cross-correlation based techniques and partial-correlation. To determine a connection, and also its strength, between two electrodes we calculated the area under the highest peak near to zero. Its latency determines the direction in the transmission of information.

Results

Cross-correlation based methods measure the direction of a possible connection between a pair of electrodes meanwhile, partial correlation comes out not only with direction but also eliminates indirect connections and gives the real strength between two channels. However, partialization presents some limitations when the number of neurons and connectivity of the network increases. Depending on how big is the network, partial correlation can show unreliable results or even breakdown in the identification of synaptic connections. In this work, we analyzed data obtained from cortical cultures coupled to high-density MEAs and we compared cross-correlation based techniques with partial-correlation analysis.

Discussion

Cross-correlation based methods are useful tools to estimate functional connectivity at the population level. Standard cross-correlation is applied just between pairs of electrodes and does not consider the entire network. It is the simplest method to infer about functional connectivity and obtain a general overview of the network map. On the other hand, partial correlation gives more details about the connectivity. Both methods can be used to study the development of the network or changes in the network behavior after electrical and/or chemical stimulation.

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