

Biology

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Daniela Niemeyer



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Catalog of Copyright Entries. Third Series Library of Congress. Copyright Office, 1963 **The Neuronal Functions of EF-hand Ca(2+)-binding Proteins 2nd Edition** Michael R. Kreutz, José R. Naranjo, Karl-Wilhelm Koch, Beat Schwaller, 2016-11-01 Ca² signaling in neurons is characterized by highly restricted and dynamic gradients called Ca² waves spikes transients and puffs depending upon their corresponding spatial and temporal features Based on this strict segmentation the Ca² ion provides a versatile basis for complex signaling in neuronal subcompartments with a spatial resolution of micro and nanodomains The multitude of Ca² regulated processes requires specialized downstream processing machinery translating the Ca² signal into alterations of cellular processes The broad range of different Ca² triggered phenomena in neurons ranging from neurotransmission to gene expression is reflected by the existence of a multitude of different Ca² binding proteins CaBPs from which numerous belong to the EF hand super family EF hand proteins can be subdivided into Ca² buffer and Ca² sensor proteins Whereas the first group has a very high affinity for Ca² exhibits little conformational change in the Ca² bound state and is thought to mainly chelate Ca² the second group has a lower affinity for Ca² and shows considerable conformational changes upon Ca² binding which usually triggers a target interaction Neuronal calcium sensor NCS proteins and the related Caldendrin CaBP Calneuron nCaBPs proteins are members of this latter group They resemble the structure of their common ancestor Calmodulin CaM with four EF hand Ca² binding motifs of which not all are functional However despite their structural homology with CaM NCS as well as nCaBPs are quite diverse in amino acid sequence It is therefore surprising that relatively few binding partners have been identified that are not CaM targets and this raises the question of the specificity and function of these interactions In terms of function binding of NCS and

nCaBP has frequently different consequences than binding of CaM which substantially increases the versatility of the Ca²⁺ tool kit. The general idea of this special issue is to provide an overview on the function of neuronal EF hand calcium binding proteins in health and disease. But we will not just provide a mere collection of articles to stress the function of each protein. The issue will mainly deal with emerging concepts on Ca²⁺ signaling buffering mediated by EF hand Ca²⁺ binding proteins. This includes questions like features that define the functional role of a EF hand calcium sensor in neurons, the conditions that make physiological relevance of a given interaction of a CaBP with its target plausible, the emerging synaptic role of these proteins and mounting evidence for their role in the regulation of protein trafficking. Structural aspects and biophysical studies will be covered. Another aspect will be the role of CaBPs in brain disease states. This aspect includes studies showing that CaBPs are targets of drugs in clinical use, studies showing that expression levels of calcium binding proteins are frequently altered in brain disease states as well as reports on mutations in EF hand calcium sensors linked to human disease.

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