



Membrane Morphology: Implications for Cellular Regulation and Function

Richard Bozelli*

Department of Biochemistry and Biomedical Sciences, McMaster University, Ontario, Canada

DESCRIPTION

Biological membranes are dynamic structures that enclose cells and organelles, defining their boundaries and regulating molecular traffic. Beyond their role as physical barriers, membranes play an important role in regulating various biological processes through their shape and curvature. The curvature of membranes influences membrane protein function, intracellular signaling, membrane trafficking, and even cell division. This article explores how membrane shape impacts these biological processes and the mechanisms by which cells regulate membrane curvature.

Membrane curvature and protein function

Membrane proteins are integral to the function of biological membranes, performing tasks such as transport, signaling, and cell adhesion. The curvature of membranes can significantly impact the structure and function of these proteins. For example, proteins embedded in highly curved membranes, such as those found in vesicles or tubules, may adopt different conformations compared to proteins in flat membranes. This conformational change can alter protein activity, affecting processes such as ion transport across membranes or receptor-ligand interactions. Furthermore, membrane curvature can enable the clustering of proteins involved in signaling pathways. Curved membranes provide spatial constraints that promote the assembly of protein complexes, thereby enhancing signal transduction efficiency. This clustering phenomenon is important for processes like receptor-mediated endocytosis, where curved membranes enable the formation of coated pits for receptor internalization and subsequent signal termination.

Membrane trafficking and vesicle formation

Membrane trafficking involves the transport of proteins and lipids between cellular compartments through vesicles. The formation of vesicles requires significant membrane curvature, which is regulated by specialized protein complexes. For instance, the formation of clathrin-coated vesicles, involved in

receptor-mediated endocytosis and intracellular transport, depends on the curvature-inducing properties of clathrin and adaptor proteins. These proteins assemble into a lattice structure on the membrane surface, inducing local curvature and enabling vesicle budding. Similarly, the biogenesis of other types of vesicles, such as exosomes and synaptic vesicles, involves proteins that can generate or stabilize membrane curvature. These proteins may include BAR (Bin/Amphiphysin/Rvs) domain proteins, which sense and induce membrane curvature through their unique structural motifs. The curvature-sensing ability of these proteins is crucial for vesicle formation and cargo sorting, ensuring precise targeting and delivery within the cell.

Membrane curvature in cellular signaling

Cellular signaling pathways are intricately regulated by membrane curvature, which influences the localization and activity of signaling molecules. For example, lipid rafts—microdomains enriched in cholesterol and sphingolipids, are often found in highly curved regions of the plasma membrane. These rafts serve as platforms for the assembly of signaling complexes, including receptor tyrosine kinases and G-protein-coupled receptors, which are important for initiating and propagating signals across the cell membrane. Moreover, membrane curvature can modulate the activity of enzymes involved in lipid metabolism and second messenger production. Changes in membrane curvature alter the accessibility of these enzymes to their substrates, thereby regulating the synthesis and turnover of lipid signaling molecules like phosphoinositides and diacylglycerol. These lipid species play essential roles in cellular processes such as membrane trafficking, cytoskeletal dynamics, and ion channel regulation, highlighting the importance of membrane curvature in signal transduction.

Membrane shape during cell division

During cell division, membranes undergo dramatic shape changes to accommodate processes such as chromosome segregation and cytokinesis. The formation of the mitotic spindle, a microtubule-based structure responsible for

Correspondence to: Richard Bozelli, Department of Biochemistry and Biomedical Sciences, McMaster University, Ontario, Canada, E-mail: rbozelli@mcmaster.ca

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chromosome movement, requires membranes to bend and deform. Membrane curvature is also essential for the formation of the contractile ring during cytokinesis, where actin and myosin assemble to divide the cell into two daughter cells. Furthermore, the maintenance of membrane curvature during cell division is regulated by a complex interaction of cytoskeletal components, membrane-associated proteins, and lipid composition. For example, the BAR domain protein family plays a vital role in shaping membranes during cytokinesis and membrane remodeling events associated with cell division. These proteins contribute to the formation of membrane tubules and vesicles required for intracellular transport and organelle biogenesis.

Regulation of membrane curvature

Cells employ various mechanisms to regulate membrane curvature and ensure proper function during physiological processes. One such mechanism involves the recruitment of curvature-sensing and curvature-inducing proteins to specific membrane regions. These proteins, such as BAR domain proteins and coat complexes like clathrin and COPII, sense local curvature cues and promote membrane remodeling events. Additionally, lipid composition plays a significant role in determining membrane curvature. Lipids with cone-shaped structures, such as phosphatidylethanolamine and

lysophosphatidylcholine, induce negative curvature, whereas lipids with inverted cone shapes, like diacylglycerol and phosphatidic acid, promote positive curvature. Cells regulate lipid composition through lipid synthesis pathways and lipid transfer proteins, ensuring dynamic control over membrane curvature in response to cellular needs.

CONCLUSION

Membrane shape plays an important role in regulating biological processes ranging from protein function and membrane trafficking to cellular signaling and cell division. The curvature of membranes influences the activity of membrane proteins, enables vesicle formation and trafficking, organizes signaling complexes, and shapes cellular membranes during division. Cells tightly regulate membrane curvature through a combination of protein-mediated mechanisms and lipid composition to ensure proper function and responsiveness to environmental cues. Understanding the impact of membrane shape on biological processes provides insights into cellular physiology and opens possibilities for therapeutic interventions targeting membrane-associated diseases. As research continues to unravel the complexities of membrane curvature regulation, the potential for innovative biomedical applications and treatments grows, potential advances in fields from drug delivery to regenerative medicine.