

Synthesis of BMP



Williams, MG.

European Bioinformatics Institute, New York University Langone Medical Center, Ontario Institute for Cancer Research, Oregon Health and Science University.

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This is just an excerpt of a full-length report for this pathway. To access the complete report, please download it at the <u>Reactome Textbook</u>.

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Introduction

Reactome is open-source, open access, manually curated and peer-reviewed pathway database. Pathway annotations are authored by expert biologists, in collaboration with Reactome editorial staff and cross-referenced to many bioinformatics databases. A system of evidence tracking ensures that all assertions are backed up by the primary literature. Reactome is used by clinicians, geneticists, genomics researchers, and molecular biologists to interpret the results of high-throughput experimental studies, by bioinformaticians seeking to develop novel algorithms for mining knowledge from genomic studies, and by systems biologists building predictive models of normal and disease variant pathways.

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Literature references

- Fabregat, A., Sidiropoulos, K., Viteri, G., Forner, O., Marin-Garcia, P., Arnau, V. et al. (2017). Reactome pathway analysis: a high-performance in-memory approach. *BMC bioinformatics, 18*, 142. 7
- Sidiropoulos, K., Viteri, G., Sevilla, C., Jupe, S., Webber, M., Orlic-Milacic, M. et al. (2017). Reactome enhanced pathway visualization. *Bioinformatics*, 33, 3461-3467. A
- Fabregat, A., Jupe, S., Matthews, L., Sidiropoulos, K., Gillespie, M., Garapati, P. et al. (2018). The Reactome Pathway Knowledgebase. *Nucleic Acids Res, 46*, D649-D655.
- Fabregat, A., Korninger, F., Viteri, G., Sidiropoulos, K., Marin-Garcia, P., Ping, P. et al. (2018). Reactome graph database: Efficient access to complex pathway data. *PLoS computational biology*, *14*, e1005968. *对*

This document contains 1 pathway and 2 reactions (see Table of Contents)

Synthesis of BMP 7

Stable identifier: R-HSA-1483171



Lysobisphosphatidic acid, also known as bis(monoacylglycerol) hydrogen phosphate (BMP), is enriched in late endosomes and not found in the endoplasmic reticulum (ER) or mitochondria where phosphatidylglycerol (PG) is synthesized. Late endosomes form membrane contact sites with the ER, providing a means for PG to enter the late endosome and be converted to BMP via hydrolysis by a phospholipase A2, followed by acylation, and a reorientation of the phosphoryl ester (Poorthuis & Hostetler 1978, Heravi & Waite 1999, Hullin-Matsuda et al. 2007, Gallala & Sandhoff 2010).

Literature references

- Gallala, HD., Sandhoff, K. (2010). Biological Function of the Cellular Lipid BMP-BMP as a Key Activator for Cholesterol Sorting and Membrane Digestion. *Neurochem Res.* 7
- Waite, M., Heravi, J. (1999). Transacylase formation of bis(monoacylglycerol)phosphate. *Biochim Biophys Acta, 1437,* 277-86. *↗*
- Hostetler, KY., Poorthuis, BJ. (1978). Conversion of diphosphatidylglycerol to bis(monoacylglyceryl)phosphate by lysosomes. J Lipid Res, 19, 309-15.

Lagarde, M., Schlame, M., Kobayashi, T., Xu, Y., Kawasaki, K., Hullin-Matsuda, F. et al. (2007). De novo biosynthesis of the late endosome lipid, bis(monoacylglycero)phosphate. *J Lipid Res, 48*, 1997-2008.

Editions

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PG transports from the ER membrane to the late endosome membrane 7

Location: Synthesis of BMP

Stable identifier: R-HSA-1483218

Type: transition

Compartments: endoplasmic reticulum membrane, late endosome membrane



Lysobisphosphatidic acid, also known as bis(monoacylglycerol) hydrogen phosphate (BMP), is enriched in late endosomes and not found in the endoplasmic reticulum (ER) or mitochondria where phosphatidylglycerol (PG) is synthesised. Late endosomes form membrane contact sites with the ER, providing a means for PG to enter the late endosome and be converted to BMP (Levine 2004, Eden et al. 2010, Kobayashi et al. 1998, Hullin-Matsuda et al. 2007, Kobayashi et al. 1999).

Followed by: PG is converted to BMP

Literature references

- Levine, T. (2004). Short-range intracellular trafficking of small molecules across endoplasmic reticulum junctions. *Trends Cell Biol, 14*, 483-90. 7
- White, IJ., Eden, ER., Futter, CE., Tsapara, A. (2010). Membrane contacts between endosomes and ER provide sites for PTP1B-epidermal growth factor receptor interaction. *Nat Cell Biol, 12*, 267-72.
- Lagarde, M., Schlame, M., Kobayashi, T., Xu, Y., Kawasaki, K., Hullin-Matsuda, F. et al. (2007). De novo biosynthesis of the late endosome lipid, bis(monoacylglycero)phosphate. *J Lipid Res, 48*, 1997-2008.
- Beuchat, MH., Gruenberg, J., Parton, RG., Lindsay, M., Kobayashi, T., Sakuraba, H. et al. (1999). Late endosomal membranes rich in lysobisphosphatidic acid regulate cholesterol transport. *Nat Cell Biol*, *1*, 113-8.
- Fang, KS., Gruenberg, J., Parton, RG., Kobayashi, T., de Moerloose, P., Stang, E. (1998). A lipid associated with the antiphospholipid syndrome regulates endosome structure and function. *Nature*, *392*, 193-7. *¬*

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PG is converted to BMP *对*

Location: Synthesis of BMP

Stable identifier: R-HSA-1483209

Type: omitted

Compartments: late endosome membrane



The biosynthetic pathway of lysobisphosphatidic acid, also known as bis(monoacylglycerol) hydrogen phosphate (BMP), is still not fully understood with the *in vivo* enzymes responsible yet to be fully identified. It appears to involve multiple steps including hydrolysis of phosphatidylglycerol (PG) by a phospholipase A2, acylation, and a reorientation of the phosphoryl ester (Poorthuis & Hostetler 1978, Heravi & Waite 1999, Hullin-Matsuda et al. 2007, Gallala & Sandhoff 2010).

Preceded by: PG transports from the ER membrane to the late endosome membrane

Literature references

- Gallala, HD., Sandhoff, K. (2010). Biological Function of the Cellular Lipid BMP-BMP as a Key Activator for Cholesterol Sorting and Membrane Digestion. *Neurochem Res.* 7
- Waite, M., Heravi, J. (1999). Transacylase formation of bis(monoacylglycerol)phosphate. *Biochim Biophys Acta, 1437,* 277-86. ↗
- Hostetler, KY., Poorthuis, BJ. (1978). Conversion of diphosphatidylglycerol to bis(monoacylglyceryl)phosphate by lysosomes. J Lipid Res, 19, 309-15.
- Lagarde, M., Schlame, M., Kobayashi, T., Xu, Y., Kawasaki, K., Hullin-Matsuda, F. et al. (2007). De novo biosynthesis of the late endosome lipid, bis(monoacylglycero)phosphate. *J Lipid Res, 48*, 1997-2008.

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Table of Contents

Introduction	1
🐐 Synthesis of BMP	2
PG transports from the ER membrane to the late endosome membrane	3
PG is converted to BMP	4
Table of Contents	5