

Bmal1:Clock,Npas2 heterodimer is phosphorylated and translocates to the nucleus

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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. [↗](#)

Sidiropoulos, K., Viteri, G., Sevilla, C., Jupe, S., Webber, M., Orlic-Milacic, M. et al. (2017). Reactome enhanced pathway visualization. *Bioinformatics*, 33, 3461-3467. [↗](#)

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. [↗](#)

Reactome database release: 88

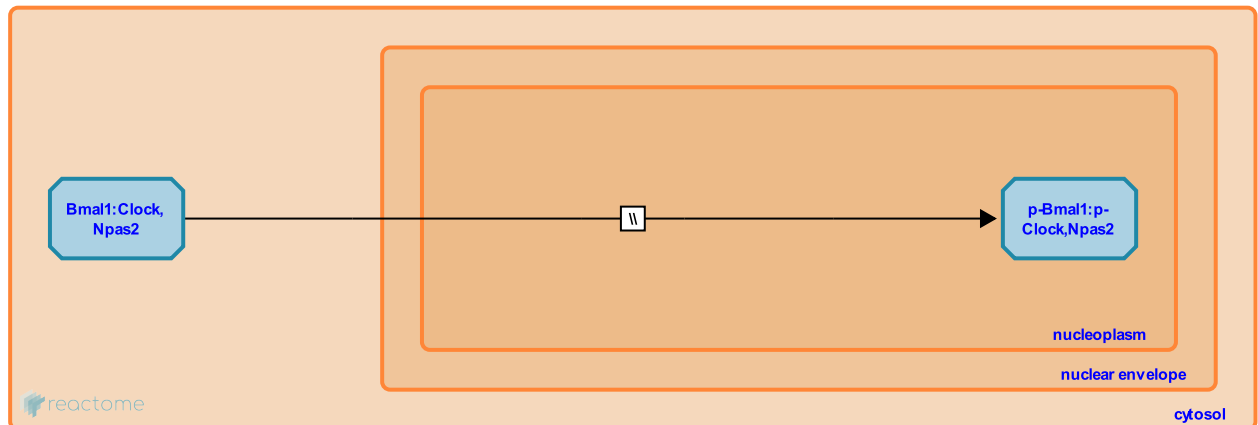
This document contains 1 reaction ([see Table of Contents](#))

Bmal1:Clock,Npas2 heterodimer is phosphorylated and translocates to the nucleus [↗](#)

Stable identifier: R-MMU-508656

Type: omitted

Compartments: nucleoplasm, cytosol



Bmal1 (Arntl), Clock, and Npas2 are phosphorylated by unknown kinases. The phosphorylation is dependent on the heterodimerization of Bmal1 with Clock or Npas2. Phosphorylated Bmal1:Clock/Npas2 is a much stronger transactivator of gene expression than is unphosphorylated Bmal1:Clock/Npas2.

Literature references

- Antoch, MP., Kondratova, AA., Gorbacheva, VY., Kondratov, RV., Gudkov, AV., Chernov, MV. (2003). BMAL1-dependent circadian oscillation of nuclear CLOCK: posttranslational events induced by dimerization of transcriptional activators of the mammalian clock system. *Genes Dev*, 17, 1921-32. [↗](#)
- Fukada, Y., Yoshitane, H., Takao, T., Du, NH., Satomi, Y., Okano, T. (2009). Roles of CLOCK phosphorylation in suppression of E-box-dependent transcription. *Mol Cell Biol*, 29, 3675-86. [↗](#)
- Eide, EJ., Vielhaber, EL., Virshup, DM., Hinz, WA. (2002). The circadian regulatory proteins BMAL1 and cryptochromes are substrates of casein kinase Iepsilon. *J Biol Chem*, 277, 17248-54. [↗](#)
- Hirayama, J., Nagai, K., Isojima, Y., Tamaru, T., Takamatsu, K., Sassone-Corsi, P. et al. (2009). CK2alpha phosphorylates BMAL1 to regulate the mammalian clock. *Nat Struct Mol Biol*, 16, 446-8. [↗](#)
- Jung, NC., Son, GH., Lee, BJ., Chang, SH., Kim, K., Kwon, I. et al. (2006). BMAL1 shuttling controls transactivation and degradation of the CLOCK/BMAL1 heterodimer. *Mol Cell Biol*, 26, 7318-30. [↗](#)

Editions

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