

UHRF1 binds chromatin with hemimethyl-

ated cytosine

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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. *オ*

This document contains 1 reaction (see Table of Contents)

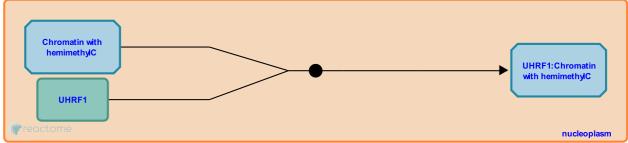
UHRF1 binds chromatin with hemimethylated cytosine **7**

Stable identifier: R-HSA-5334099

Type: binding

Compartments: nucleoplasm

Inferred from: Uhrf1 binds chromatin with hemimethylC (Mus musculus)



UHRF1 (also known as Np95) preferentially binds hemimethylated CG dinucleotides in DNA via its SRA domain (Avvakumov et al. 2008, Qian et al. 2008, and inferred from the mouse homolog). The UHRF1-bound unmethylated cytosine base is flipped out of the DNA helix and into a pocket of UHRF1 (Avvakumov et al. 2008). UHRF1 also binds dimethylated and trimethylated lysine-9 of histone H3 through its tandem Tudor domain (Nady et al. 2011, Rothbart et al. 2012, Rothbart et al. 2013, Cheng et al. 2013) and unmethylated histone H3 through its PHD domain (Hu et al. 2011, Wang et al. 2011, Rajakumara et al. 2011, Cheng et al. 2013).

Literature references

- Chédin, F., Wang, YX., Nady, N., Bronner, C., Lemak, A., Avvakumov, GV. et al. (2011). Recognition of multivalent histone states associated with heterochromatin by UHRF1 protein. J. Biol. Chem., 286, 24300-11. ↗
- Xue, S., Bronner, C., Avvakumov, GV., Li, Y., Walker, JR., Arrowsmith, CH. et al. (2008). Structural basis for recognition of hemi-methylated DNA by the SRA domain of human UHRF1. *Nature*, 455, 822-5.
- Wang, P., Hu, L., Xu, Y., Li, Z., Lin, Y. (2011). Crystal structure of PHD domain of UHRF1 and insights into recognition of unmodified histone H3 arginine residue 2. *Cell Res., 21*, 1374-8. 7
- Fang, J., Wang, P., Cheng, J., Zhu, T., Yang, H., Chen, F. et al. (2013). Structural insight into coordinated recognition of trimethylated histone H3 lysine 9 (H3K9me3) by the plant homeodomain (PHD) and tandem tudor domain (TTD) of UHRF1 (ubiquitin-like, containing PHD and RING finger domains, 1) protein. J. Biol. Chem., 288, 1329-39
- Hu, W., Wu, H., Wang, C., Cao, C., Li, G., Chen, P. et al. (2011). Structural basis for site-specific reading of unmodified R2 of histone H3 tail by UHRF1 PHD finger. *Cell Res.*, 21, 1379-82.

Editions

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