

DNMT3A:DNMT3L binds chromatin

Beekman, R., Martín-Subero, JI., May, B.

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

The contents of this document may be freely copied and distributed in any media, provided the authors, plus the institutions, are credited, as stated under the terms of [Creative Commons Attribution 4.0 International \(CC BY 4.0\) License](#). For more information see our [license](#).

02/05/2024

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.

The development of Reactome is supported by grants from the US National Institutes of Health (P41 HG003751), University of Toronto (CFREF Medicine by Design), European Union (EU STRP, EMI-CD), and the European Molecular Biology Laboratory (EBI Industry program).

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](#))

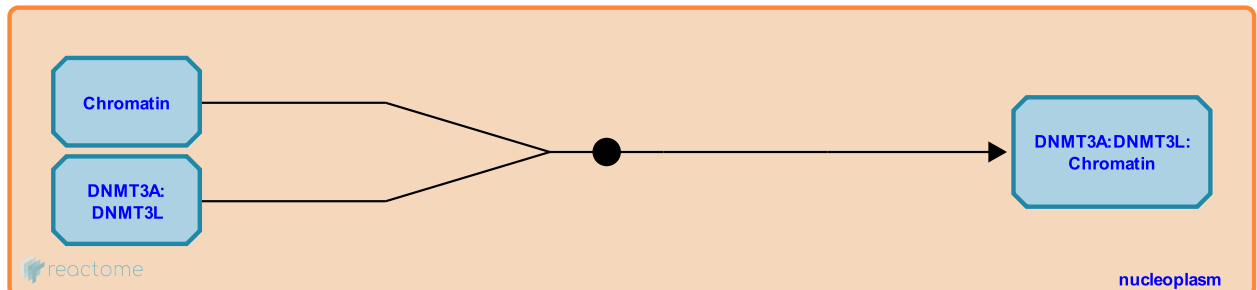
DNMT3A:DNMT3L binds chromatin [↗](#)

Stable identifier: R-HSA-5334179

Type: binding

Compartments: nucleoplasm

Inferred from: [Dnmt3a:Dnmt3l binds chromatin \(Mus musculus\)](#)



DNMT3L is a non-catalytic homologue of DNMT3A and DNMT3B which binds unmethylated lysine-4 of histone H3 (H3K4, Ooi et al. 2007) and recruits DNMT3A-isoform2 (germ cell specific) (Chen et al. 2005, Kareta et al. 2006, Ooi et al. 2007) and, to a lesser extent, other isoforms of DNMT3A via an interaction between the C-terminal regions of DNMT3L and DNMT3A (Chen et al. 2005). Furthermore, DNMT3A binds unmodified tails of histone H3 (Otani et al. 2009). DNMT3L and DNMT3A form tetramers (DNMT3L:DNMT3A:DNMT3A:DNMT3L) that bind DNA located in euchromatic regions (Holz-Schietinger et al. 2011, also inferred from mouse). DNMT3A without DNMT3L forms oligomers that are located in heterochromatin.

Literature references

- Chédin, F., Mann, JR., Hsieh, CL., Chen, ZX., Riggs, AD. (2005). Physical and functional interactions between the human DNMT3L protein and members of the de novo methyltransferase family. *J. Cell. Biochem.*, 95, 902-17. [↗](#)
- Inamoto, S., Shirakawa, M., Ariyoshi, M., Otani, J., Arita, K., Nankumo, T. (2009). Structural basis for recognition of H3K4 methylation status by the DNA methyltransferase 3A ATRX-DNMT3-DNMT3L domain. *EMBO Rep.*, 10, 1235-41. [↗](#)
- Chédin, F., Botello, ZM., Ennis, JJ., Kareta, MS., Chou, C. (2006). Reconstitution and mechanism of the stimulation of de novo methylation by human DNMT3L. *J. Biol. Chem.*, 281, 25893-902. [↗](#)
- Matje, DM., Reich, NO., Harrison, MF., Holz-Schietinger, C. (2011). Oligomerization of DNMT3A controls the mechanism of de novo DNA methylation. *J. Biol. Chem.*, 286, 41479-88. [↗](#)
- Lin, SP., Yang, Z., Li, K., Bernstein, E., Ooi, SK., Bestor, TH. et al. (2007). DNMT3L connects unmethylated lysine 4 of histone H3 to de novo methylation of DNA. *Nature*, 448, 714-7. [↗](#)

Editions

2014-02-21	Authored, Edited	May, B.
2014-07-24	Reviewed	Beekman, R., Martín-Subero, JI.