

# Dicer cleaves double-stranded RNA to yield double-stranded siRNA

May, B., Tomari, Y.

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](#).

05/05/2024

## 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

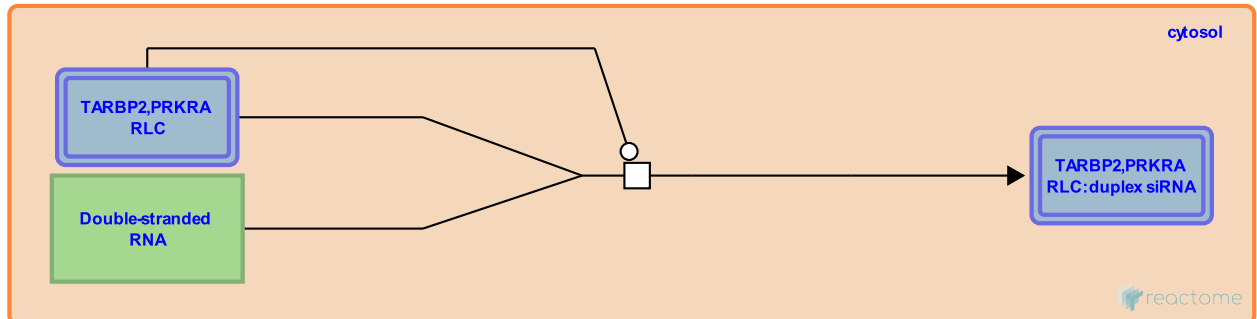
This document contains 1 reaction ([see Table of Contents](#))

## Dicer cleaves double-stranded RNA to yield double-stranded siRNA ↗

**Stable identifier:** R-HSA-426464

**Type:** transition

**Compartments:** cytosol



Double stranded RNA binds the RISC loading complex and DICER1, an RNase III component of the complex, cleaves double-stranded RNAs to yield short double-stranded RNAs of 21-25 nucleotides, duplex siRNAs (small interfering RNAs). SiRNAs are similar to microRNAs (miRNAs) in their final structure but differ from miRNAs in their source: siRNAs are produced from long double stranded RNAs that originate from viruses, transposable elements, centromeric repeats and other repetitive structures.

The RLC as originally characterized contains DICER1, AGO2, and TARBP2 (TRBP). Alternative RLCs appear to contain other Argonaute proteins (AGO1, AGO3, AGO4) rather than AGO2 and PRKRA rather than TARBP2. Diffusion activity of TARBP2 and PRKRA along duplex RNA may enhance processing by DICER1.

### Literature references

- Ching, YP., Kok, KH., Jin, DY., Ng, MH. (2007). Human TRBP and PACT directly interact with each other and associate with dicer to facilitate the production of small interfering RNA. *J. Biol. Chem.*, 282, 17649-57. ↗
- Mourelatos, Z., Maniataki, E. (2005). A human, ATP-independent, RISC assembly machine fueled by pre-miRNA. *Genes Dev*, 19, 2979-90. ↗
- Kim, YK., Park, SY., Hur, I., Lee, Y., Kim, VN., Suh, MR. (2006). The role of PACT in the RNA silencing pathway. *EMBO J.*, 25, 522-32. ↗
- Hung, JH., Weng, Z., Fukunaga, R., Zamore, PD., Han, BW., Xu, J. (2012). Dicer partner proteins tune the length of mature miRNAs in flies and mammals. *Cell*, 151, 533-46. ↗
- Gregory, RI., Shiekhattar, R., Chendrimada, TP., Norman, J., Nishikura, K., Cooch, N. et al. (2005). TRBP recruits the Dicer complex to Ago2 for microRNA processing and gene silencing. *Nature*, 436, 740-4. ↗

### Editions

2009-06-10	Authored, Edited	May, B.
2012-02-11	Reviewed	Tomari, Y.