

BRCA1 forms a heterodimer with BARD1

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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

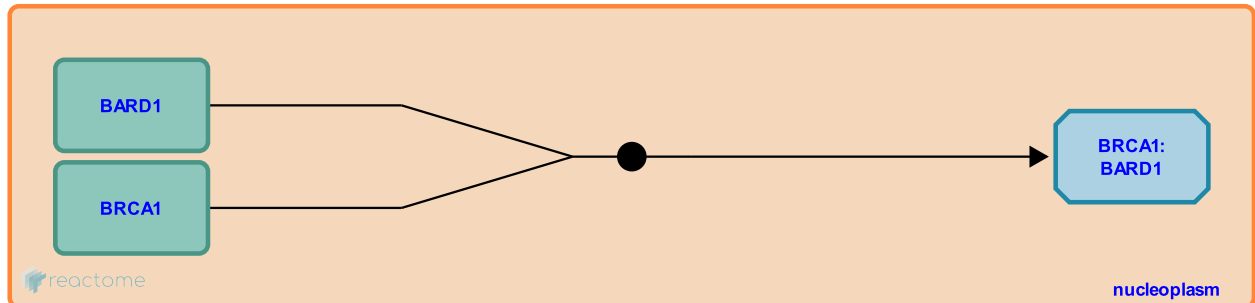
This document contains 1 reaction ([see Table of Contents](#))

BRCA1 forms a heterodimer with BARD1 [↗](#)

Stable identifier: R-HSA-5659781

Type: binding

Compartments: nucleoplasm



BRCA1 and BARD1 form a stable heterodimer through an interaction between sequences encompassing their N-terminal RING domains (Wu et al. 1996, Brzovic et al. 2001). In addition to the RING domains, BRCA1 and BARD1 both have tandem BRCT motifs at their C termini. The central region of BARD1 contains ankyrin repeats (Wu et al. 1996). Formation of BRCA1:BARD1 heterodimers is necessary for the repair of double-strand DNA breaks by homologous recombination (Westermarck et al. 2003, Laufer et al. 2007), BRCA1-mediated tumor suppression (Shakya et al. 2008) and normal development (McCarthy et al. 2003). Tumorigenic BRCA1 mutations that abolish the formation of BRCA1:BARD1 heterodimers have been reported (Wu et al. 1996, Brzovic et al. 2001).

Literature references

- Szabolcs, M., Baer, RJ., Ludwig, T., Shakya, R., Basso, K., Ospina, E. et al. (2008). The basal-like mammary carcinomas induced by Brca1 or Bard1 inactivation implicate the BRCA1/BARD1 heterodimer in tumor suppression. *Proc. Natl. Acad. Sci. U.S.A.*, 105, 7040-5. [↗](#)
- Murty, VV., Nandula, SV., Baer, RJ., Jasin, M., Ludwig, T., Modi, AP. et al. (2007). Structural requirements for the BARD1 tumor suppressor in chromosomal stability and homology-directed DNA repair. *J. Biol. Chem.*, 282, 34325-33. [↗](#)
- King, MC., Hoyt, DW., Rajagopal, P., Brzovic, PS., Kleit, RE. (2001). Structure of a BRCA1-BARD1 heterodimeric RING-RING complex. *Nat. Struct. Biol.*, 8, 833-7. [↗](#)
- Phung, A., Bowcock, AM., Wu, LC., Hwang, LY., Wang, ZW., Baer, R. et al. (1996). Identification of a RING protein that can interact in vivo with the BRCA1 gene product. *Nat. Genet.*, 14, 430-40. [↗](#)
- Baer, RJ., Jasin, M., Moynahan, ME., Westermarck, UK., Olshen, AB., Reyngold, M. (2003). BARD1 participates with BRCA1 in homology-directed repair of chromosome breaks. *Mol. Cell. Biol.*, 23, 7926-36. [↗](#)

Editions

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