

Type II receptor serine/threonine kinases in GtoPdb v.2025.3

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Abstract

Type II protein receptor serine/threonine kinases interact with transforming growth factor beta (TGF β), bone morphogenic protein (BMP) or MÄllerian inhibiting substrate (MIS). Type II RSTKs then phosphorylate the kinase domain of their [Type I RSTK](#) partner - sometimes referred to as the signal propagating unit. This causes displacement of protein partners, such as the FKBP12 FK506-binding protein [FKBP1A](#) ([P62942](#)) and allowing the binding and phosphorylation of particular members of the Smad family.

Contents

This is a citation summary for Type II receptor serine/threonine kinases in the [Guide to Pharmacology](#) database (GtoPdb). It exists purely as an adjunct to the database to facilitate the recognition of citations to and from the database by citation analyzers. Readers will almost certainly want to visit the relevant sections of the database which are given here under database links.

[GtoPdb](#) is an expert-driven guide to pharmacological targets and the substances that act on them. GtoPdb is a reference work which is most usefully represented as an on-line database. As in any publication this work should be appropriately cited, and the papers it cites should also be recognized. This document provides a citation for the relevant parts of the database, and also provides a reference list for the research cited by those parts. For further details see [\[4\]](#).

Please note that the database version for the citations given in GtoPdb are to the most recent preceding version in which the family or its subfamilies and targets were substantially changed. The links below are to the current version. If you need to consult the cited version, rather than the most recent version, please contact the GtoPdb curators.

Database links

Type II receptor serine/threonine kinases

<https://www.guidetopharmacology.org/GRAC/FamilyDisplayForward?familyId=319>

Receptors

[ActR2\(activin A receptor type 2A\)](#)

<https://www.guidetopharmacology.org/GRAC/ObjectDisplayForward?objectId=1791>

[ActR2B\(activin A receptor type 2B\)](#)

<https://www.guidetopharmacology.org/GRAC/ObjectDisplayForward?objectId=1792>

[MISR2\(anti-Müllerian hormone receptor type 2\)](#)

<https://www.guidetopharmacology.org/GRAC/ObjectDisplayForward?objectId=1793>

[BMPR2\(bone morphogenic protein receptor type 2\)](#)

<https://www.guidetopharmacology.org/GRAC/ObjectDisplayForward?objectId=1794>

[TGFB2\(transforming growth factor beta receptor 2\)](#)

<https://www.guidetopharmacology.org/GRAC/ObjectDisplayForward?objectId=1795>

References

1. Anastassiadis T, Deacon SW, Devarajan K, Ma H and Peterson JR. (2011) Comprehensive assay of kinase catalytic activity reveals features of kinase inhibitor selectivity. *Nat Biotechnol* **29**: 1039-45 [[PMID:22037377](#)]
2. Asai-Coakwell M, March L, Dai XH, Duval M, Lopez I, French CR, Famulski J, De Baere E, Francis PJ and Sundaresan P *et al.*. (2013) Contribution of growth differentiation factor 6-

dependent cell survival to early-onset retinal dystrophies. *Hum Mol Genet* **22**: 1432-42

[PMID:23307924]

3. Berger C, Herrmann T, Lu C, Sheppard K-A, Trifileff E and Urlinger S. (2013) Compositions and methods for increasing muscle growth. Patent number: [US8388968](#). Assignee: Novartis Ag. Priority date: 27/04/2009. Publication date: 05/03/2013.
4. Buneman P, Christie G, Davies JA, Dimitrellou R, Harding SD, Pawson AJ, Sharman JL and Wu Y. (2020) Why data citation isn't working, and what to do about it *Database* **2020** [PMID:32367113]
5. Davis MI, Hunt JP, Herrgard S, Ciceri P, Wodicka LM, Pallares G, Hocker M, Treiber DK and Zarrinkar PP. (2011) Comprehensive analysis of kinase inhibitor selectivity. *Nat Biotechnol* **29**: 1046-51 [PMID:22037378]
6. Deep S, Walker KP, Shu Z and Hinck AP. (2003) Solution structure and backbone dynamics of the TGFbeta type II receptor extracellular domain. *Biochemistry* **42**: 10126-39 [PMID:12939140]
7. Engers DW, Frist AY, Lindsley CW, Hong CC and Hopkins CR. (2013) Synthesis and structure-activity relationships of a novel and selective bone morphogenetic protein receptor (BMP) inhibitor derived from the pyrazolo[1.5-a]pyrimidine scaffold of dorsomorphin: the discovery of ML347 as an ALK2 versus ALK3 selective MLPCN probe. *Bioorg Med Chem Lett* **23**: 3248-52 [PMID:23639540]
8. Johnston CJC, Smyth DJ, Kodali RB, White MPJ, Harcus Y, Filbey KJ, Hewitson JP, Hinck CS, Ivens A and Kemter AM *et al.*. (2017) A structurally distinct TGF- β mimic from an intestinal helminth parasite potently induces regulatory T cells. *Nat Commun* **8**: 1741 [PMID:29170498]
9. Li HY, Wang Y, Heap CR, King CH, Mundla SR, Voss M, Clawson DK, Yan L, Campbell RM and Anderson BD *et al.*. (2006) Dihydropyrrolopyrazole transforming growth factor-beta type I receptor kinase domain inhibitors: a novel benzimidazole series with selectivity versus transforming growth factor-beta type II receptor kinase and mixed lineage kinase-7. *J Med Chem* **49**: 2138-42 [PMID:16539403]
10. Mace PD, Cutfield JF and Cutfield SM. (2006) High resolution structures of the bone morphogenetic protein type II receptor in two crystal forms: implications for ligand binding. *Biochem Biophys Res Commun* **351**: 831-8 [PMID:17094948]
11. Melisi D, Ishiyama S, Scwabas GM, Fleming JB, Xia Q, Tortora G, Abbruzzese JL and Chiao PJ. (2008) LY2109761, a novel transforming growth factor beta receptor type I and type II dual inhibitor, as a therapeutic approach to suppressing pancreatic cancer metastasis. *Mol Cancer Ther* **7**: 829-40 [PMID:18413796]
12. Weber D, Kotzsch A, Nickel J, Harth S, Seher A, Mueller U, Sebald W and Mueller TD. (2007) A silent H-bond can be mutationally activated for high-affinity interaction of BMP-2 and activin type IIB receptor. *BMC Struct Biol* **7**: 6 [PMID:17295905]
13. Wodicka LM, Ciceri P, Davis MI, Hunt JP, Floyd M, Salerno S, Hua XH, Ford JM, Armstrong RC and Zarrinkar PP *et al.*. (2010) Activation state-dependent binding of small molecule kinase inhibitors: structural insights from biochemistry. *Chem Biol* **17**: 1241-9 [PMID:21095574]
14. Wroblewski ST, Moslin R, Lin S, Zhang Y, Spergel S, Kempson J, Tokarski JS, Strnad J, Zupa-Fernandez A and Cheng L *et al.*. (2019) Highly Selective Inhibition of Tyrosine Kinase 2 (TYK2) for the Treatment of Autoimmune Diseases: Discovery of the Allosteric Inhibitor BMS-986165. *J Med Chem* **62**: 8973-8995 [PMID:31318208]
15. Yingling JM, McMillen WT, Yan L, Huang H, Sawyer JS, Graff J, Clawson DK, Britt KS, Anderson BD and Beight DW *et al.*. (2018) Preclinical assessment of galunisertib (LY2157299 monohydrate), a first-in-class transforming growth factor- β receptor type I inhibitor. *Oncotarget* **9**: 6659-6677 [PMID:29467918]