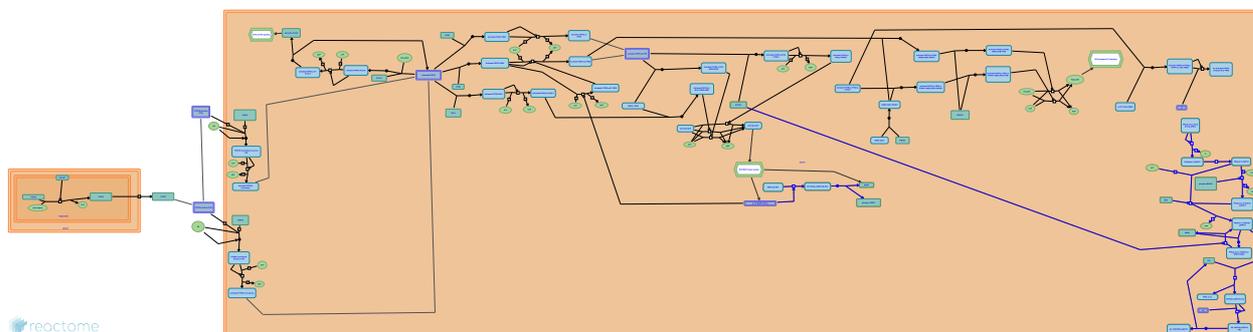


# Spry regulation of FGF signaling



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](#).

## 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: 73

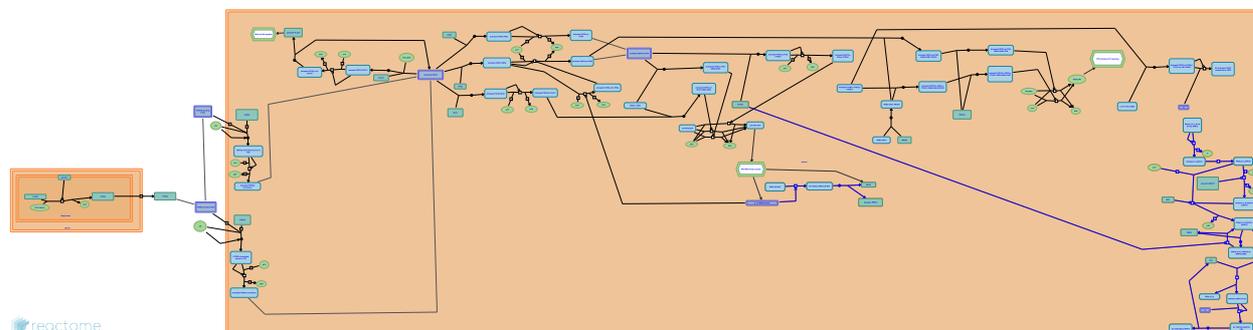
This document contains 1 pathway and 11 reactions ([see Table of Contents](#))

## Spry regulation of FGF signaling ↗

**Stable identifier:** R-CFA-1295596

**Compartments:** cytosol, plasma membrane

**Inferred from:** [Spry regulation of FGF signaling \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

## PPA2A dephosphorylates SPRY2 ↗

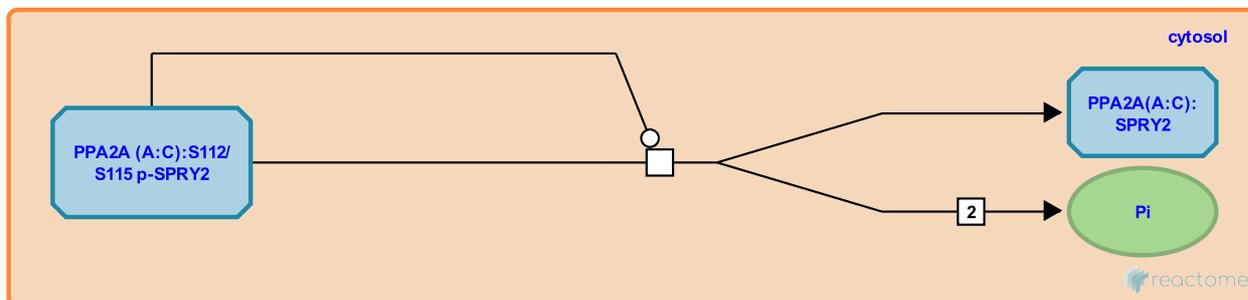
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295632

**Type:** transition

**Compartments:** cytosol

**Inferred from:** [PPA2A dephosphorylates SPRY2 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Followed by:** [SPRY2 translocates to the plasma membrane](#)

## SPRY2 translocates to the plasma membrane ↗

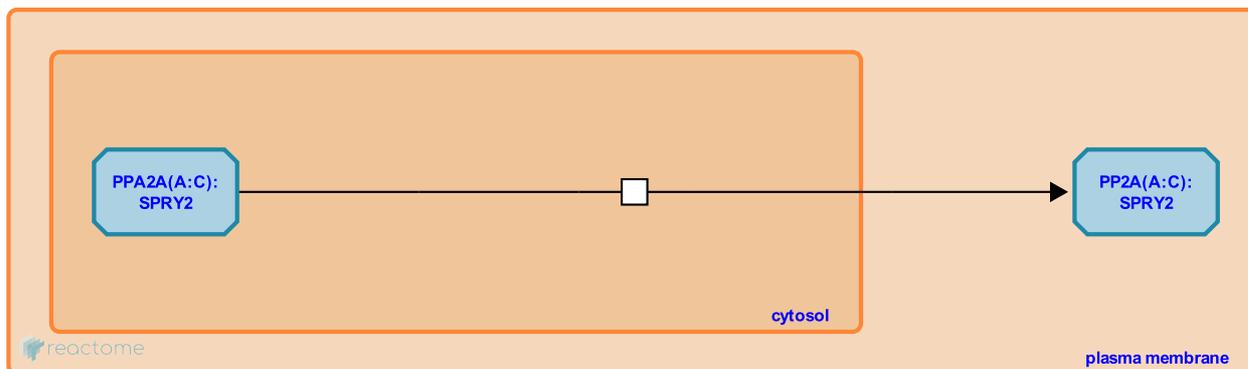
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295599

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [SPRY2 translocates to the plasma membrane \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [PPA2A dephosphorylates SPRY2](#)

**Followed by:** [SPRY2 is phosphorylated by phosphorylated MNK1](#), [SRC phosphorylates SPRY2 on Y55 and Y227](#)

## SRC phosphorylates SPRY2 on Y55 and Y227 ↗

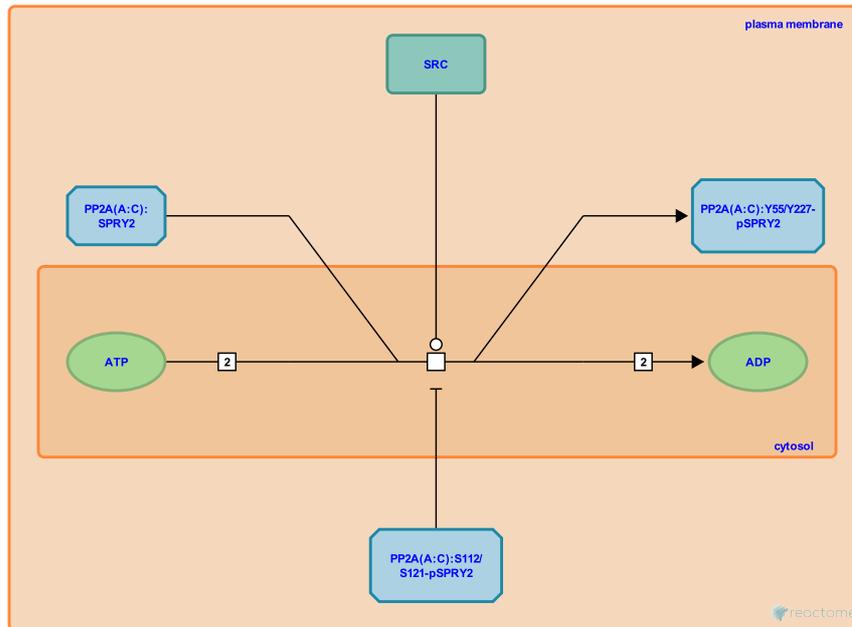
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295609

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [SRC phosphorylates SPRY2 on Y55 and Y227 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SPRY2 translocates to the plasma membrane](#)

**Followed by:** [SPRY2 binds GRB2](#), [SPRY2 binds CBL](#)

## SPRY2 binds GRB2 ↗

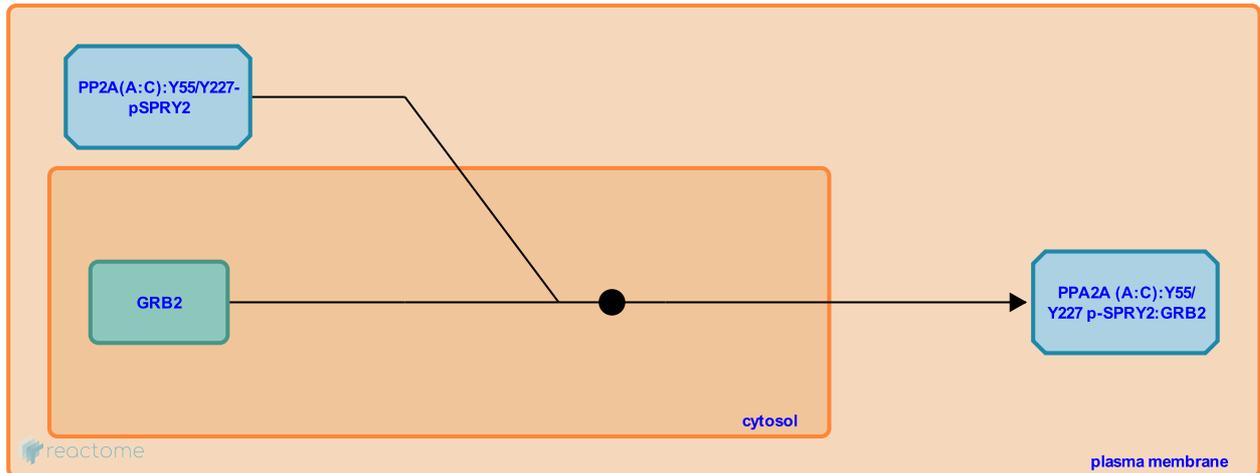
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295613

**Type:** binding

**Compartments:** cytosol, plasma membrane

**Inferred from:** [SPRY2 binds GRB2 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SRC phosphorylates SPRY2 on Y55 and Y227](#)

**Followed by:** [PPTN11 dephosphorylates SPRY2](#)

## SPRY2 binds CBL ↗

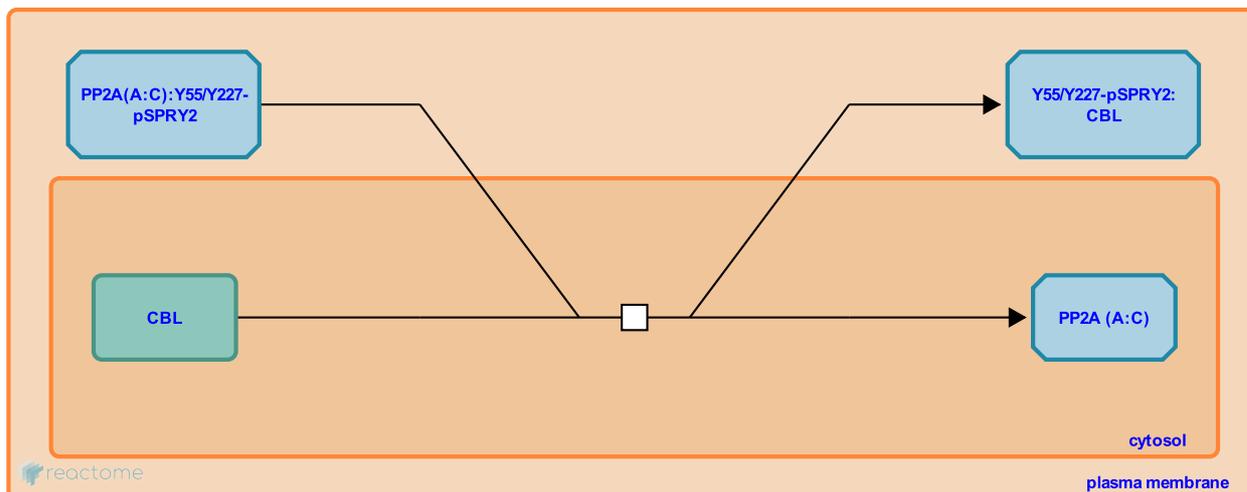
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295622

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [SPRY2 binds CBL \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SRC phosphorylates SPRY2 on Y55 and Y227](#)

**Followed by:** [Phosphorylated SPRY2 is ubiquitinated by CBL](#)

## Phosphorylated SPRY2 is ubiquitinated by CBL ↗

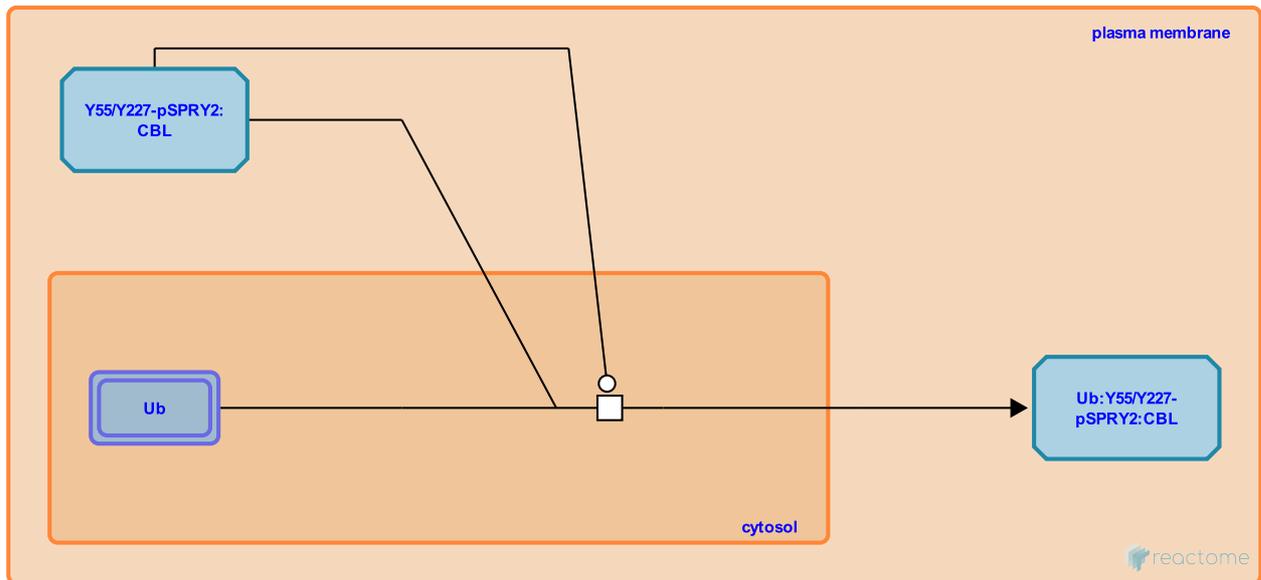
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-934604

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [Phosphorylated SPRY2 is ubiquitinated by CBL \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SPRY2 binds CBL](#)

**Followed by:** [CBL dissociates from ubiquitinated p-SPRY2](#)

## CBL dissociates from ubiquitinated p-SPRY2 ↗

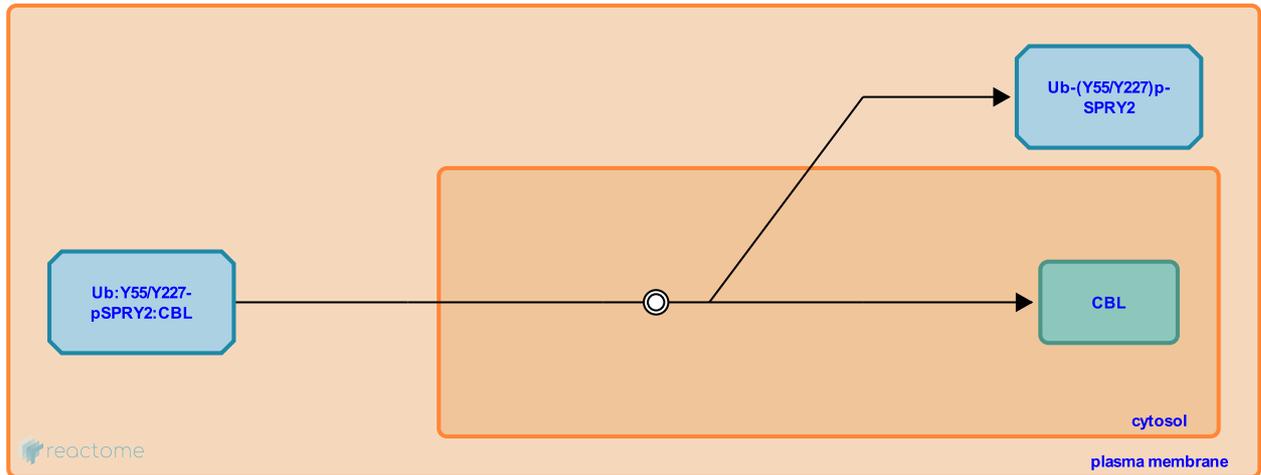
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295621

**Type:** dissociation

**Compartments:** cytosol, plasma membrane

**Inferred from:** [CBL dissociates from ubiquitinated p-SPRY2 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [Phosphorylated SPRY2 is ubiquitinated by CBL](#)

## SPRY2 is phosphorylated by phosphorylated MNK1 [↗](#)

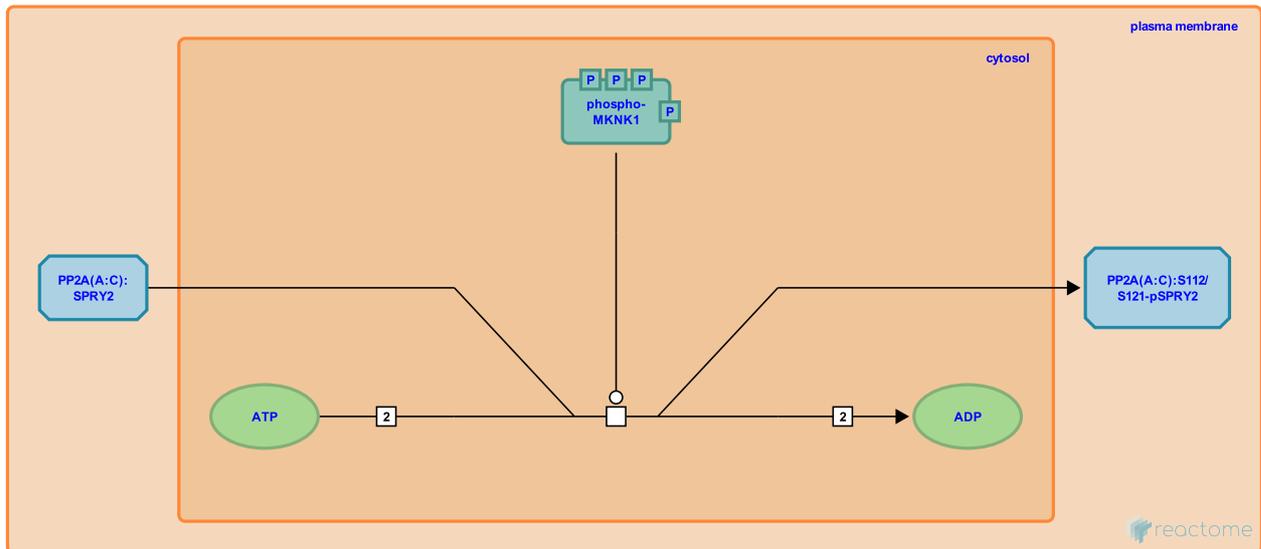
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-934559

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [SPRY2 is phosphorylated by phosphorylated MNK1 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SPRY2 translocates to the plasma membrane](#)

## SPRY2 is serine phosphorylated in response to MAPK activation ↗

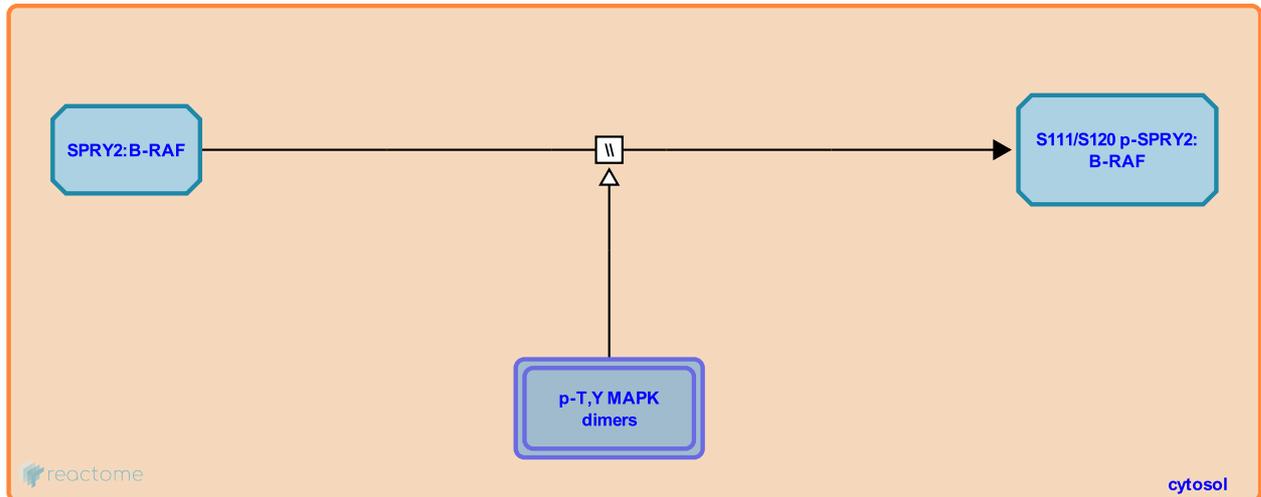
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295634

**Type:** omitted

**Compartments:** cytosol

**Inferred from:** [SPRY2 is serine phosphorylated in response to MAPK activation \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Followed by:** [B-RAF dissociates from S111/S120 p-SPRY2](#)

## B-RAF dissociates from S111/S120 p-SPRY2 ↗

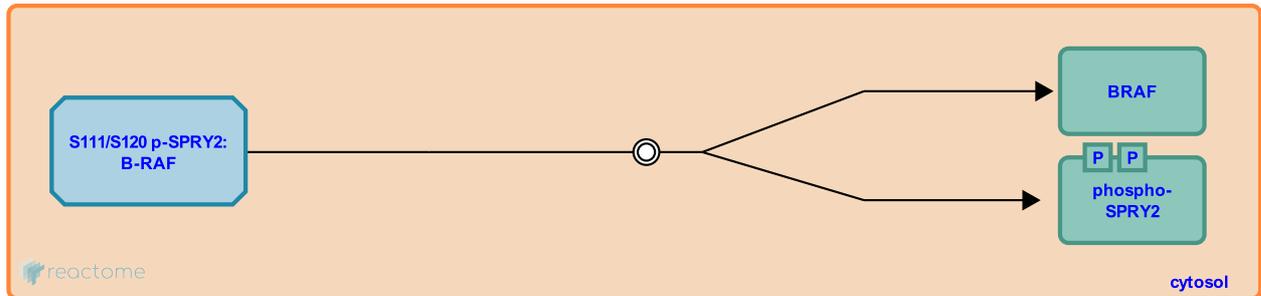
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1295604

**Type:** dissociation

**Compartments:** cytosol

**Inferred from:** [B-RAF dissociates from S111/S120 p-SPRY2 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SPRY2 is serine phosphorylated in response to MAPK activation](#)

## PPTN11 dephosphorylates SPRY2 ↗

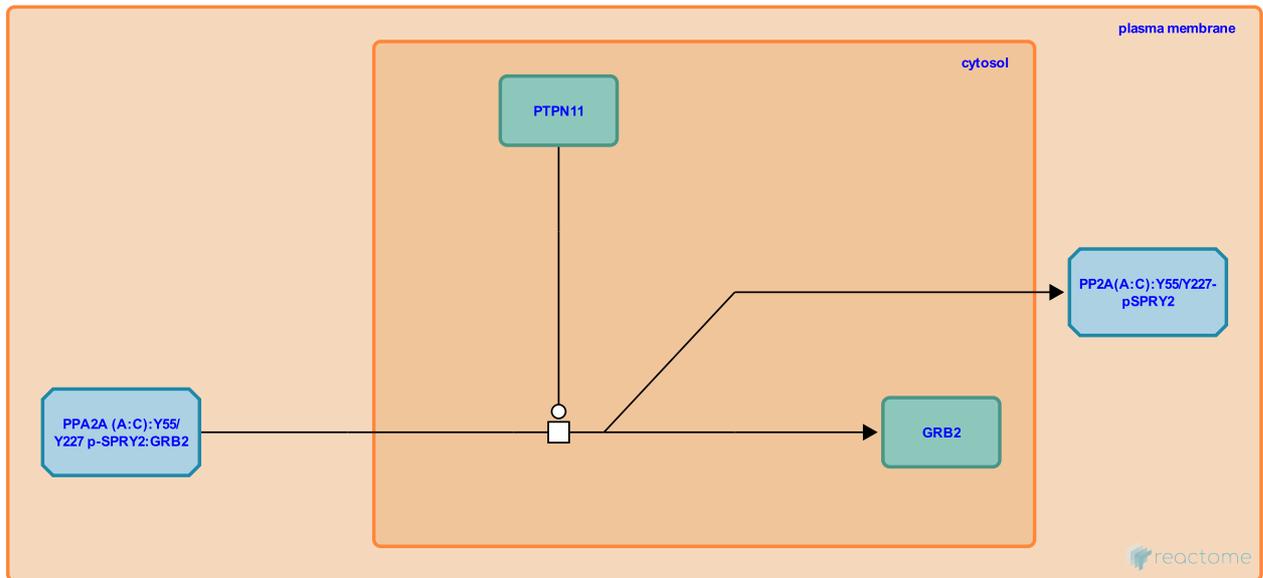
**Location:** [Spry regulation of FGF signaling](#)

**Stable identifier:** R-CFA-1549564

**Type:** transition

**Compartments:** cytosol, plasma membrane

**Inferred from:** [PPTN11 dephosphorylates SPRY2 \(Homo sapiens\)](#)



This event has been computationally inferred from an event that has been demonstrated in another species.

The inference is based on the homology mapping from PANTHER. Briefly, reactions for which all involved PhysicalEntities (in input, output and catalyst) have a mapped orthologue/paralogue (for complexes at least 75% of components must have a mapping) are inferred to the other species. High level events are also inferred for these events to allow for easier navigation.

[More details and caveats of the event inference in Reactome.](/electronic_inference_compara.html) For details on PANTHER see also: <http://www.pantherdb.org/about.jsp>

**Preceded by:** [SPRY2 binds GRB2](#)

# Table of Contents

Introduction	1
☰ Spry regulation of FGF signaling	2
↳ PPA2A dephosphorylates SPRY2	3
↳ SPRY2 translocates to the plasma membrane	4
↳ SRC phosphorylates SPRY2 on Y55 and Y227	5
↳ SPRY2 binds GRB2	6
↳ SPRY2 binds CBL	7
↳ Phosphorylated SPRY2 is ubiquitinated by CBL	8
↳ CBL dissociates from ubiquitinated p-SPRY2	9
↳ SPRY2 is phosphorylated by phosphorylated MNK1	10
☰ SPRY2 is serine phosphorylated in response to MAPK activation	11
↳ B-RAF dissociates from S111/S120 p-SPRY2	12
↳ PPTN11 dephosphorylates SPRY2	13
Table of Contents	14