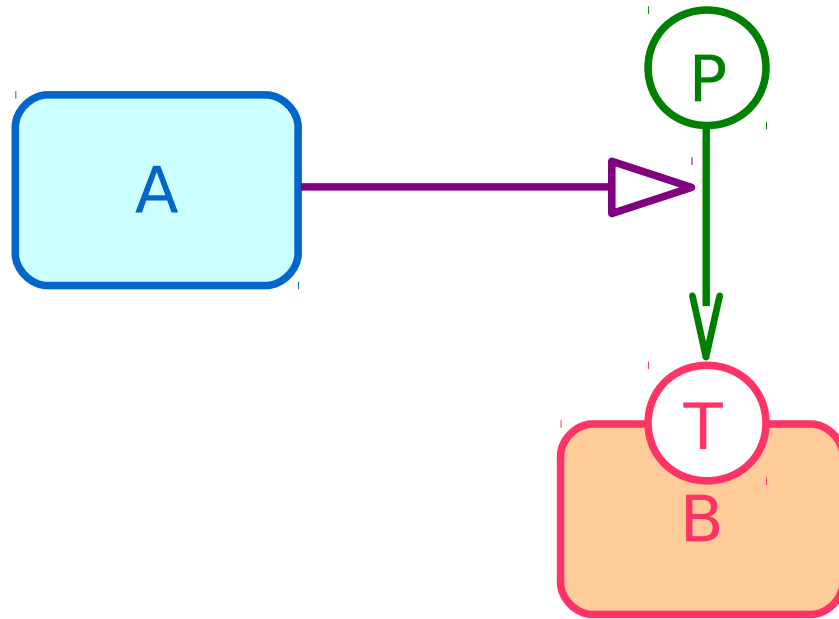


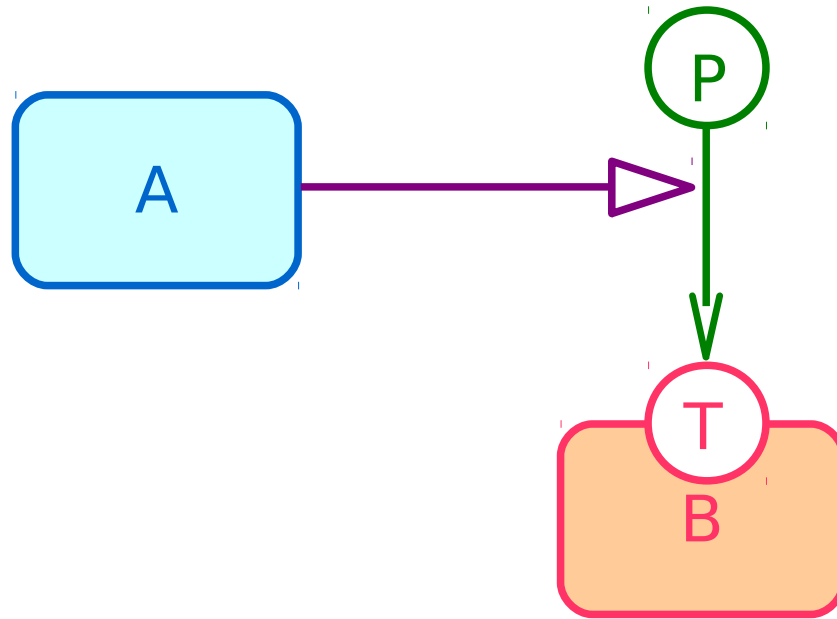
Report on the status of SBGN ER and proposed modifications/extensions

Entity Relationships can be viewed as rules



If **A** exists, the **assignment of the value P** to the **state variable T of B** is increased

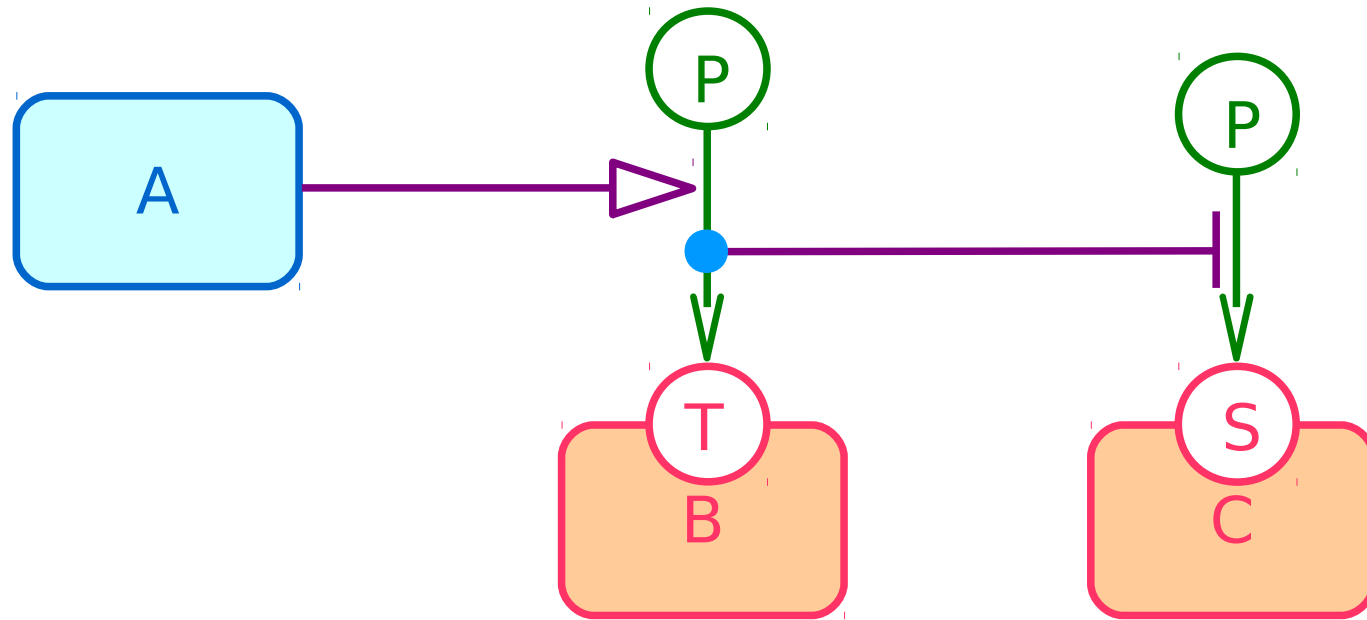
Entity Relationships can be viewed as rules



If **A** exists, the **assignment of the value P** to the **state variable T of B** is increased

(**A** stimulates the **phosphorylation** of **B on the threonine**)

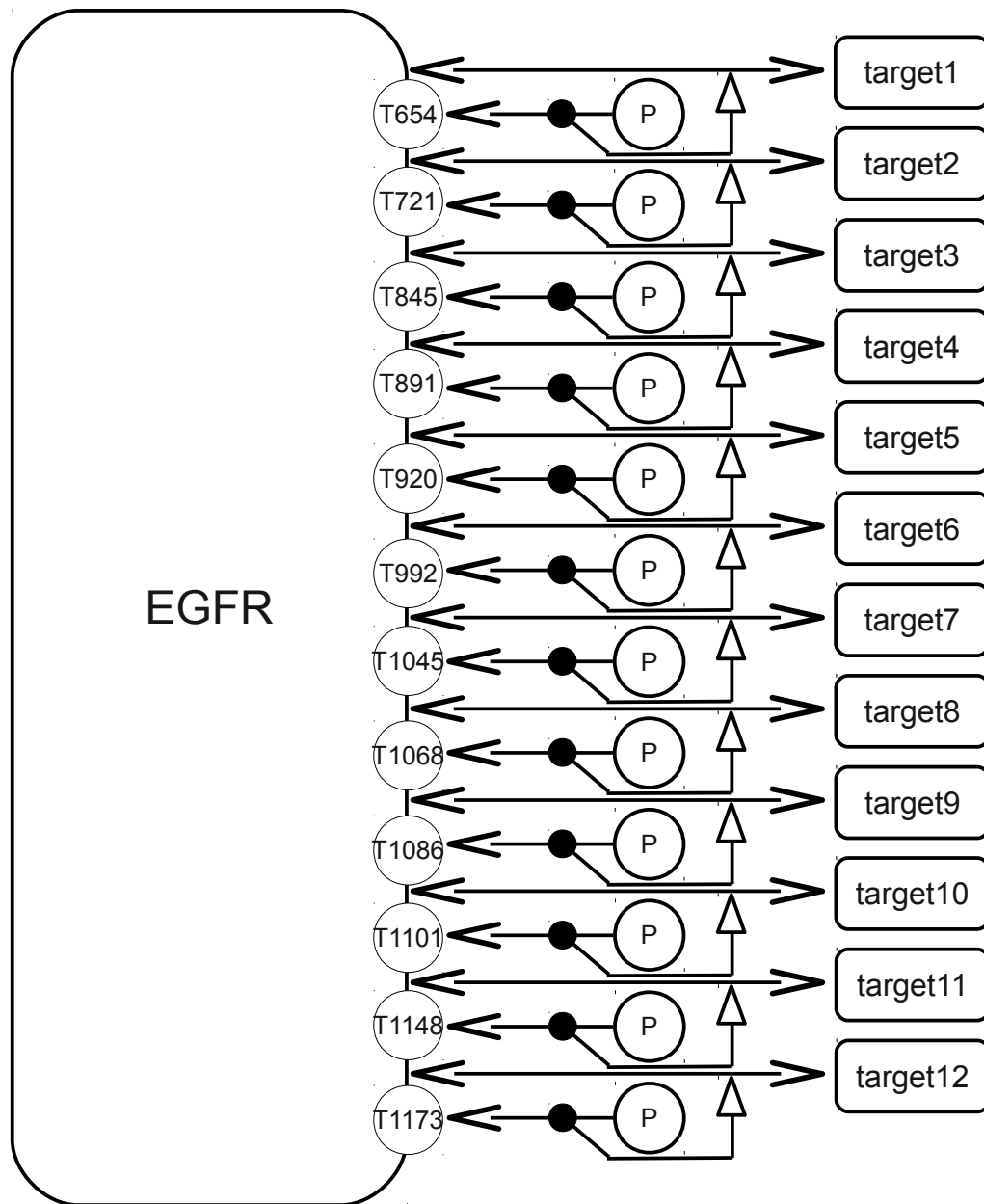
Entity Relationships can be viewed as rules



If **A exists**, the **assignment of the value P** to the **state variable T of B** is **increased**

If **P** is assigned to the **state variable T of B**, the **assignment of the value P** to the **state variable S of C** is **decreased**

Multi-state and combinatorial explosion



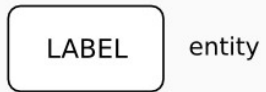
Process Descriptions:
“once a state variable value,
always a state variable value”

$2^{12} = 4096$ states
(i.e. EPN glyphs) for EGFR
and 4096 complexes between
EGFR and targets

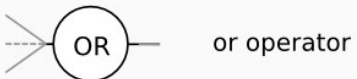
Entity Relationships L1 V1.2 reference card

Entity Nodes

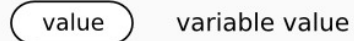
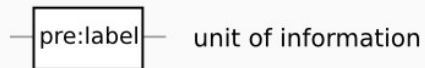
Interactors



Logical Operators

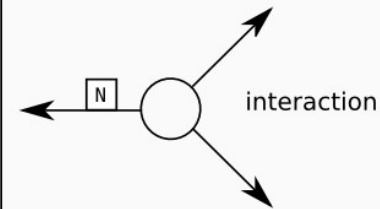


Auxiliary units

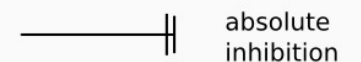
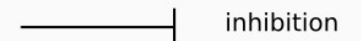
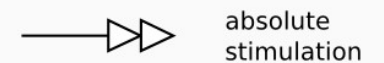


Relationship Nodes

Statements



Influence



reference nodes



Entity Relationships L1 V1 syntax

symbols \ Arc	<i>assignment</i>	<i>interaction</i>	<i>modulation</i>	<i>stimulation</i>	<i>inhibition</i>	<i>necessary stimulation</i>	<i>absolute stimulation</i>	<i>absolute inhibition</i>	<i>logic arc</i>
<i>entity</i>		IO	I	I	I	I	I	I	I
<i>outcome</i>		I(1)O(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)
<i>and</i>			I(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)O
<i>or</i>			I(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)O
<i>not</i>			I(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)O(1)
<i>delay</i>			I(1)	I(1)	I(1)	I(1)	I(1)	I(1)	I(1)O(1)
<i>perturbing agent</i>			I	I	I	I	I	I	I
<i>unit of information</i>		IO							
<i>state variable</i>	I(1)O(1)								
<i>modulation</i>				O	O	O	O	O	
<i>stimulation</i>				O	O	O	O	O	
<i>inhibition</i>				O	O	O	O	O	
<i>necessary stimulation</i>				O	O	O	O	O	
<i>absolute stimulation</i>				O	O	O	O	O	
<i>absolute inhibition</i>				O	O	O	O	O	
<i>assignment</i>				O	O	O	O	O	
<i>interaction</i>				O	O	O	O	O	
<i>phenotype</i>				O	O	O	O	O	

Example of Entity Relationships L1 V1 semantics

3.4.2 Influences

A *modulation* (Section 2.4.3.1) linking an *entity node* E and a relationship R means: “If E exists then R is either reinforced or weakened”.

A *stimulation* (Section 2.4.3.2) linking an *entity node* E and a relationship R means: “If E exists then R is reinforced” or “If E exists then the probability of R is increased”.

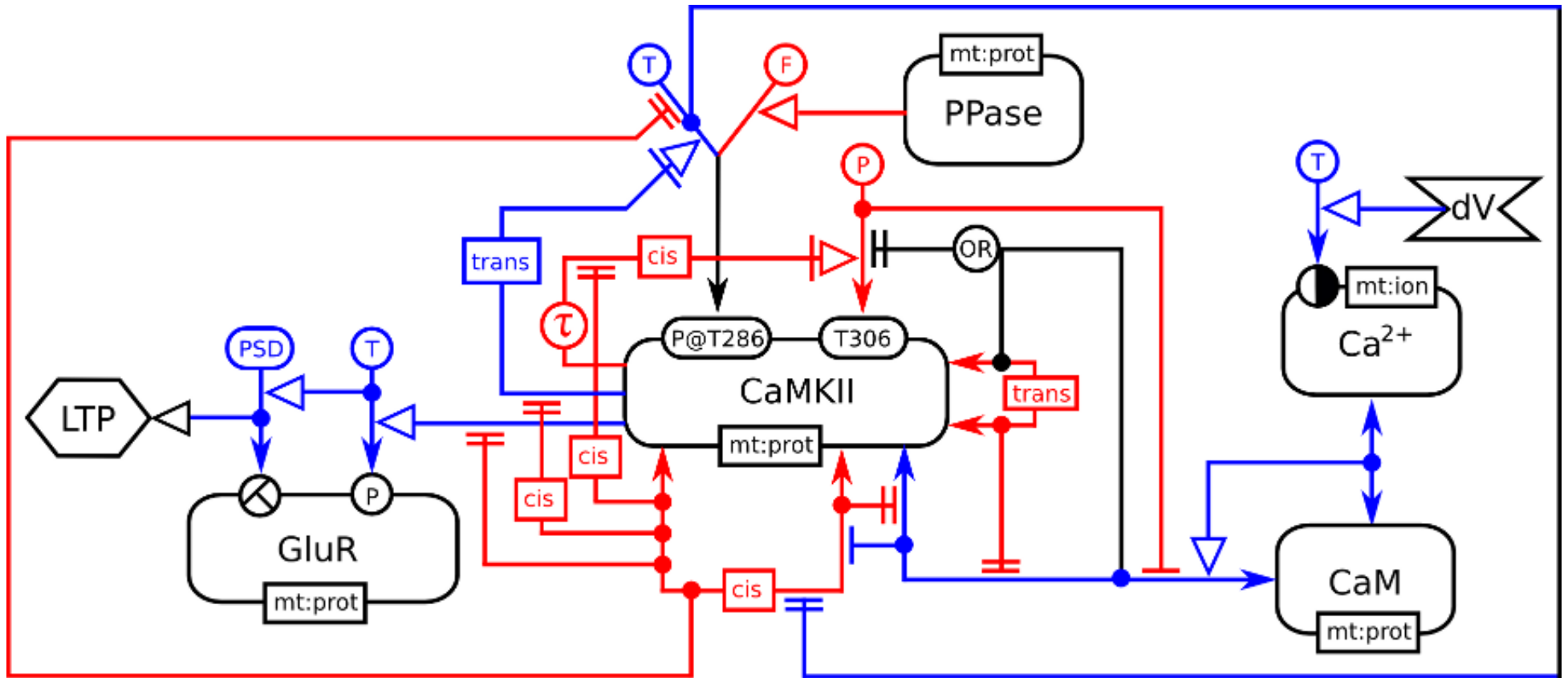
An *absolute stimulation* (Section 2.4.3.6) linking an *entity node* E and a relationship R means: “If E exists then R always takes place”.

A *necessary stimulation* (Section 2.4.3.4) linking an *entity node* E and a relationship R means: “ R only takes place if E exists.”

An *inhibition* (Section 2.4.3.3) linking an *entity node* E and a relationship R means: “If E exists then R is weakened” or “If E exists then the probability of R is lowered”.

An *absolute inhibition* (Section 2.4.3.5) linking an *entity node* E and a relationship R means: “If E exists then R never takes place”.

ER map of calcium-regulated synaptic plasticity



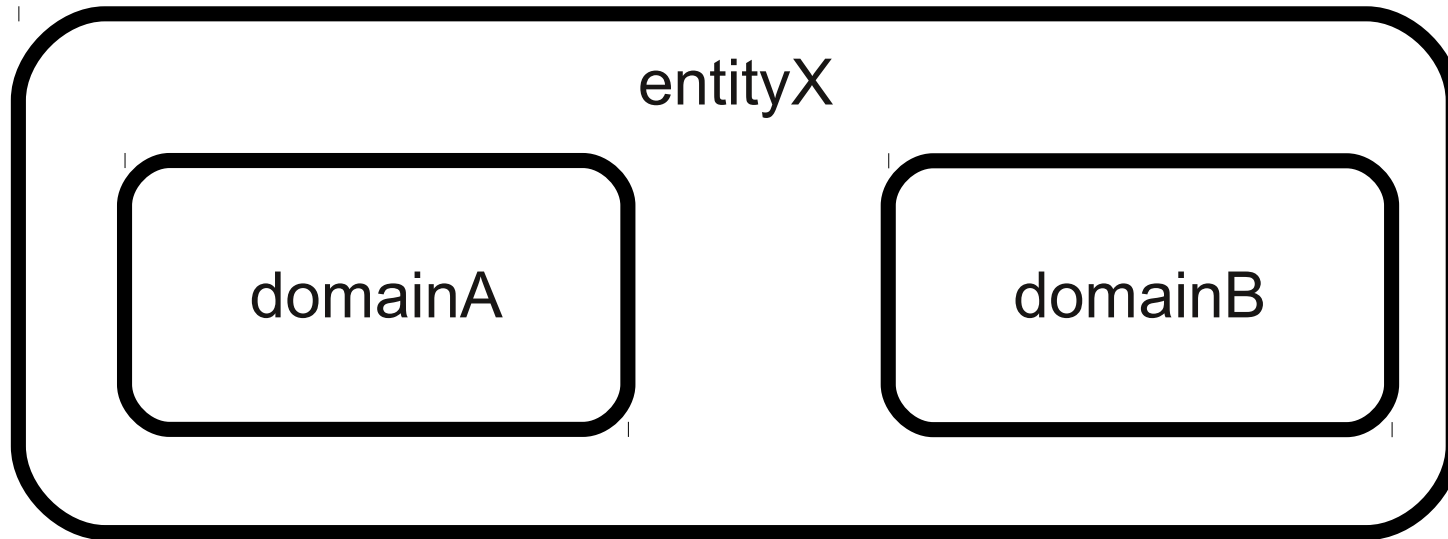
increases synaptic weight

decreases synaptic weight

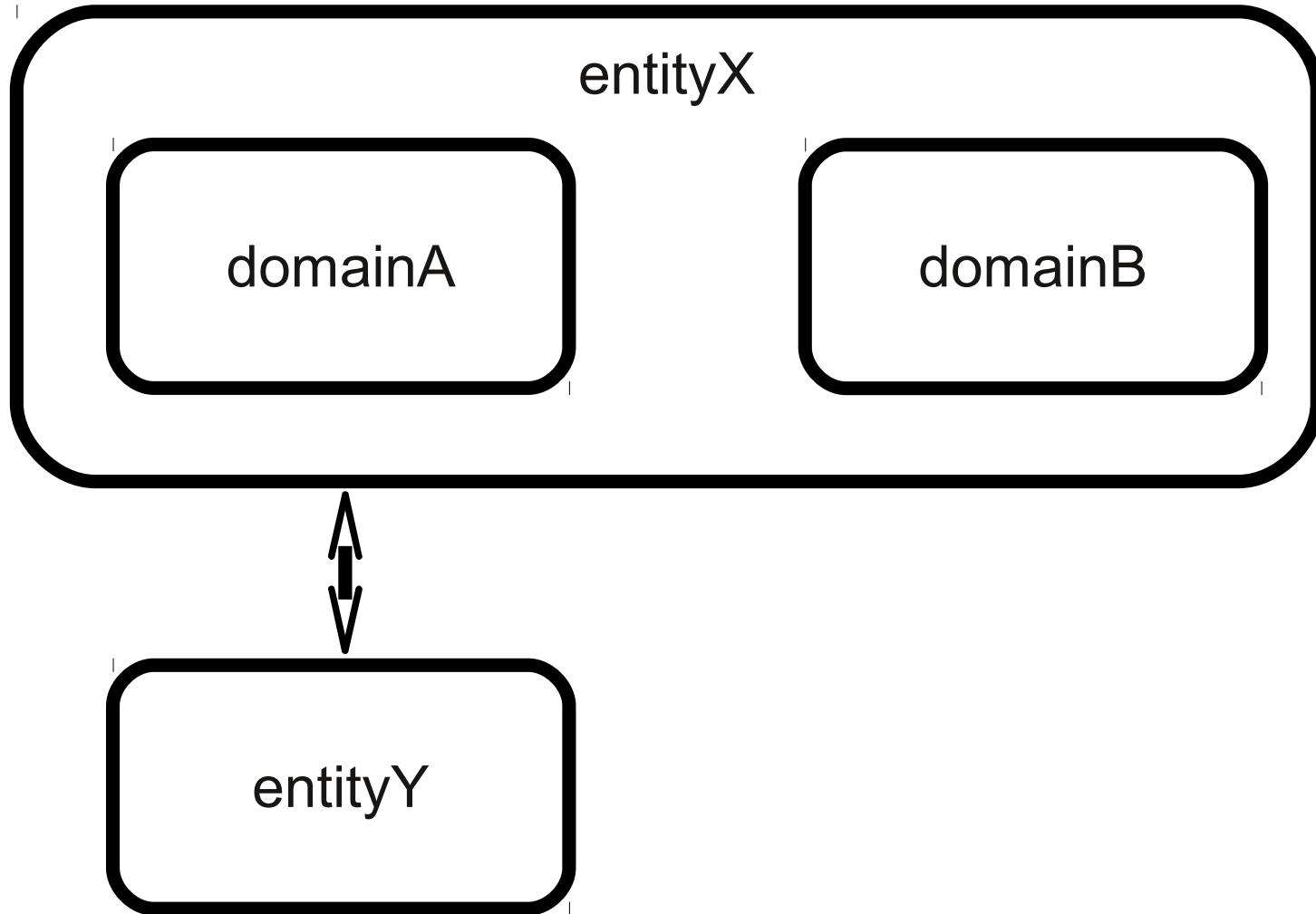
Entity Relationships L1 V2

nested entities

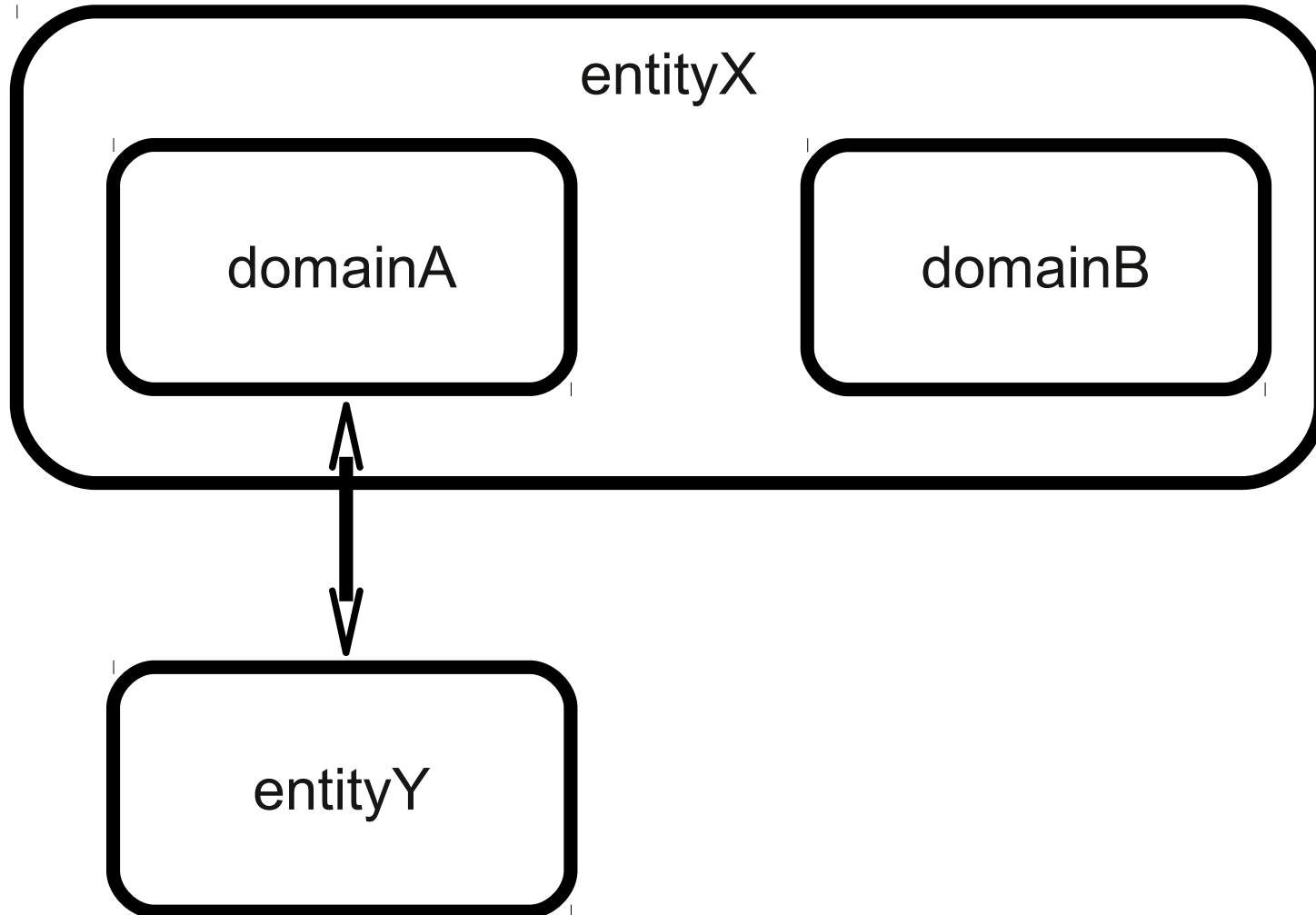
A and B are part of X



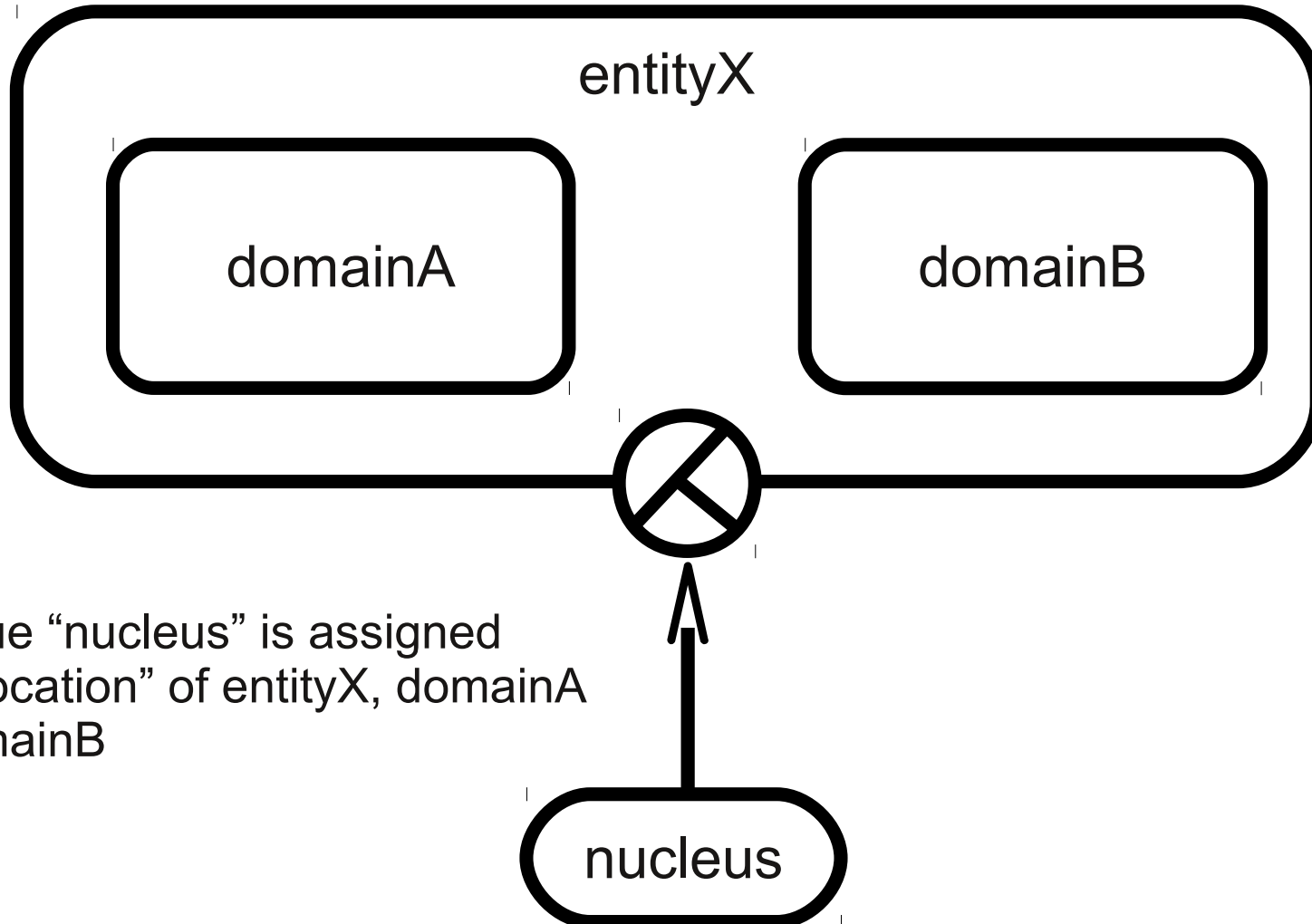
X interacts with Y



A of X interacts with Y

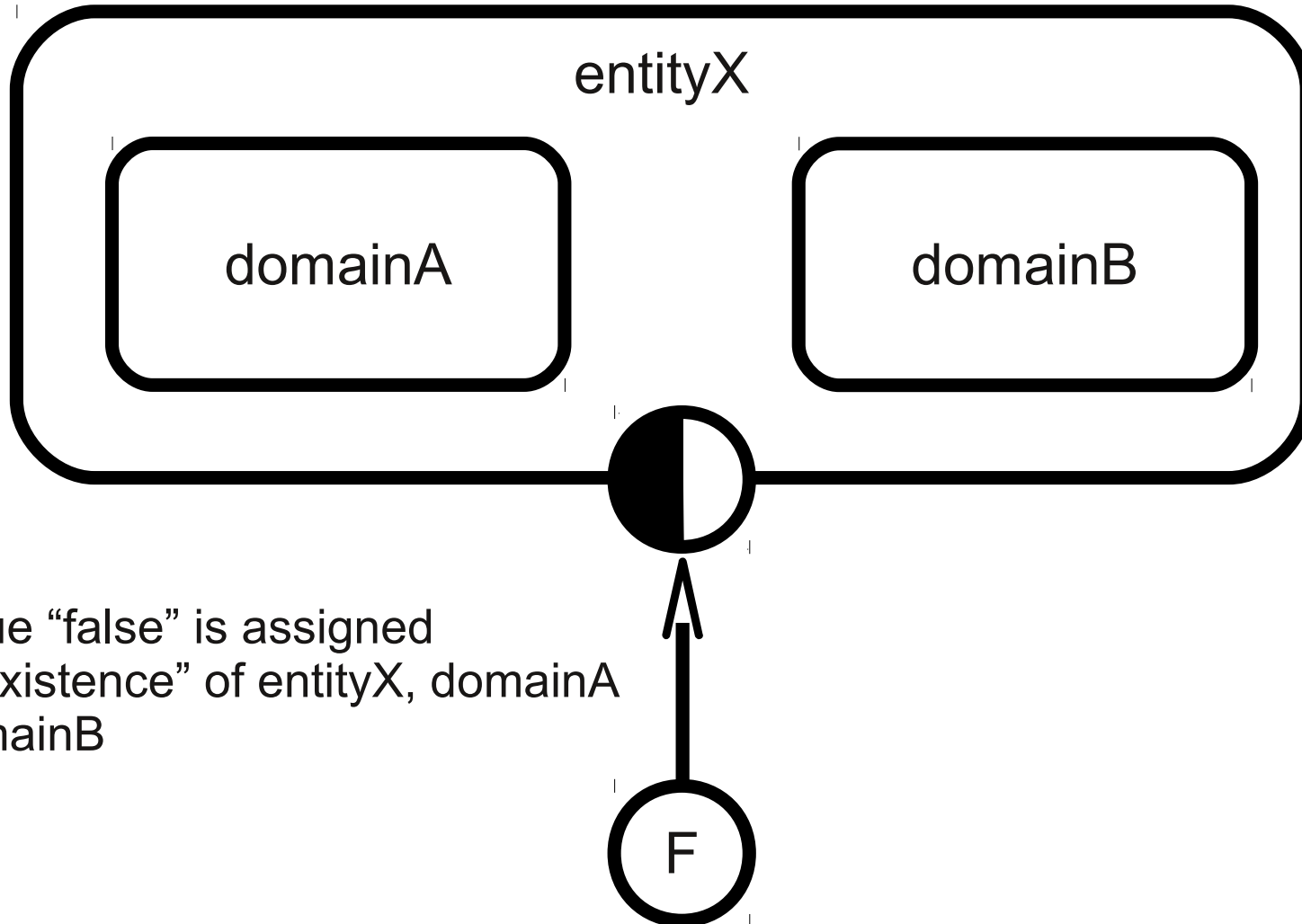


Translocation of X in the nucleus



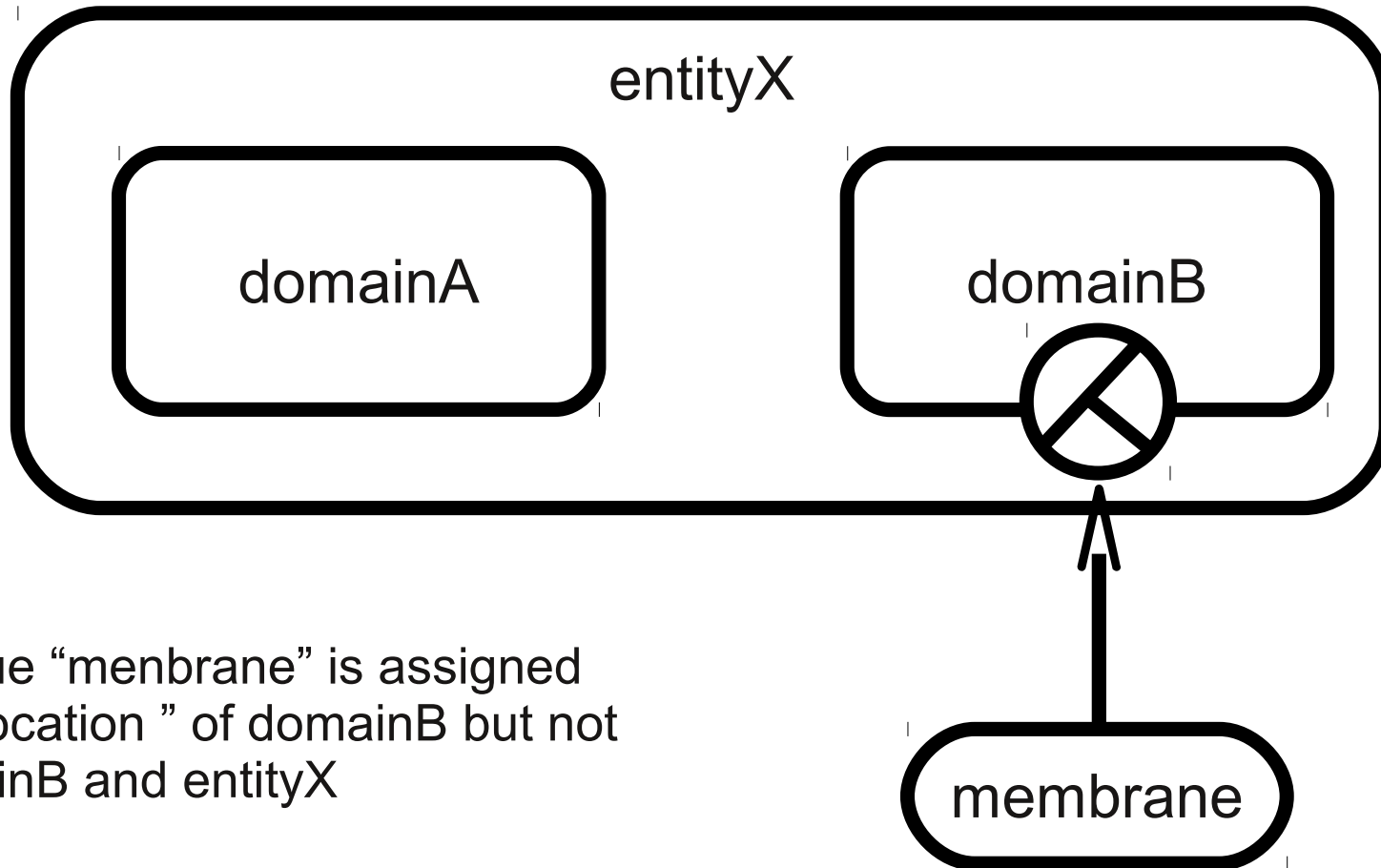
The value "nucleus" is assigned to the "location" of entityX, domainA and domainB

degradation of X



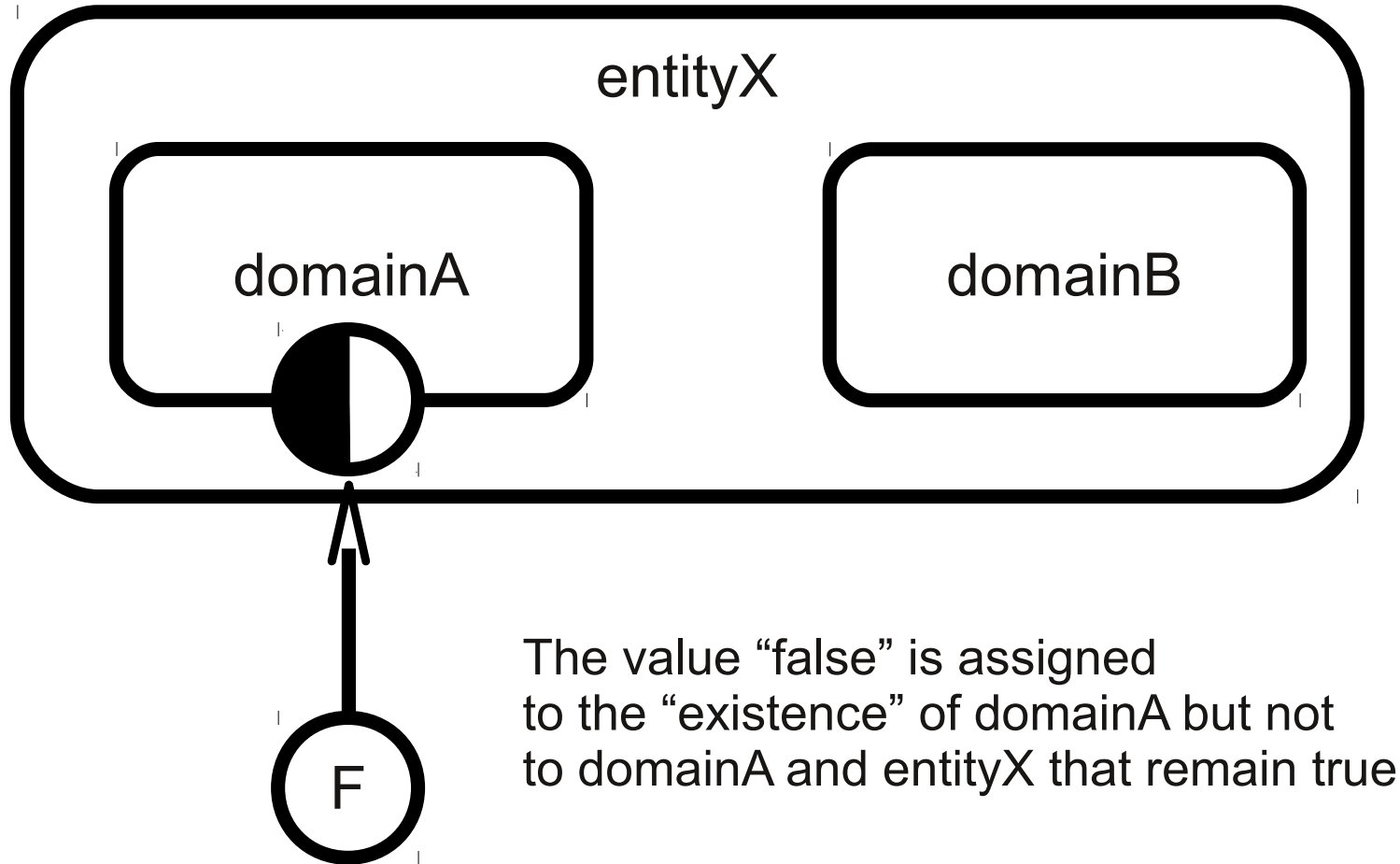
The value "false" is assigned to the "existence" of entityX, domainA and domainB

insertion of B of X in the membrane

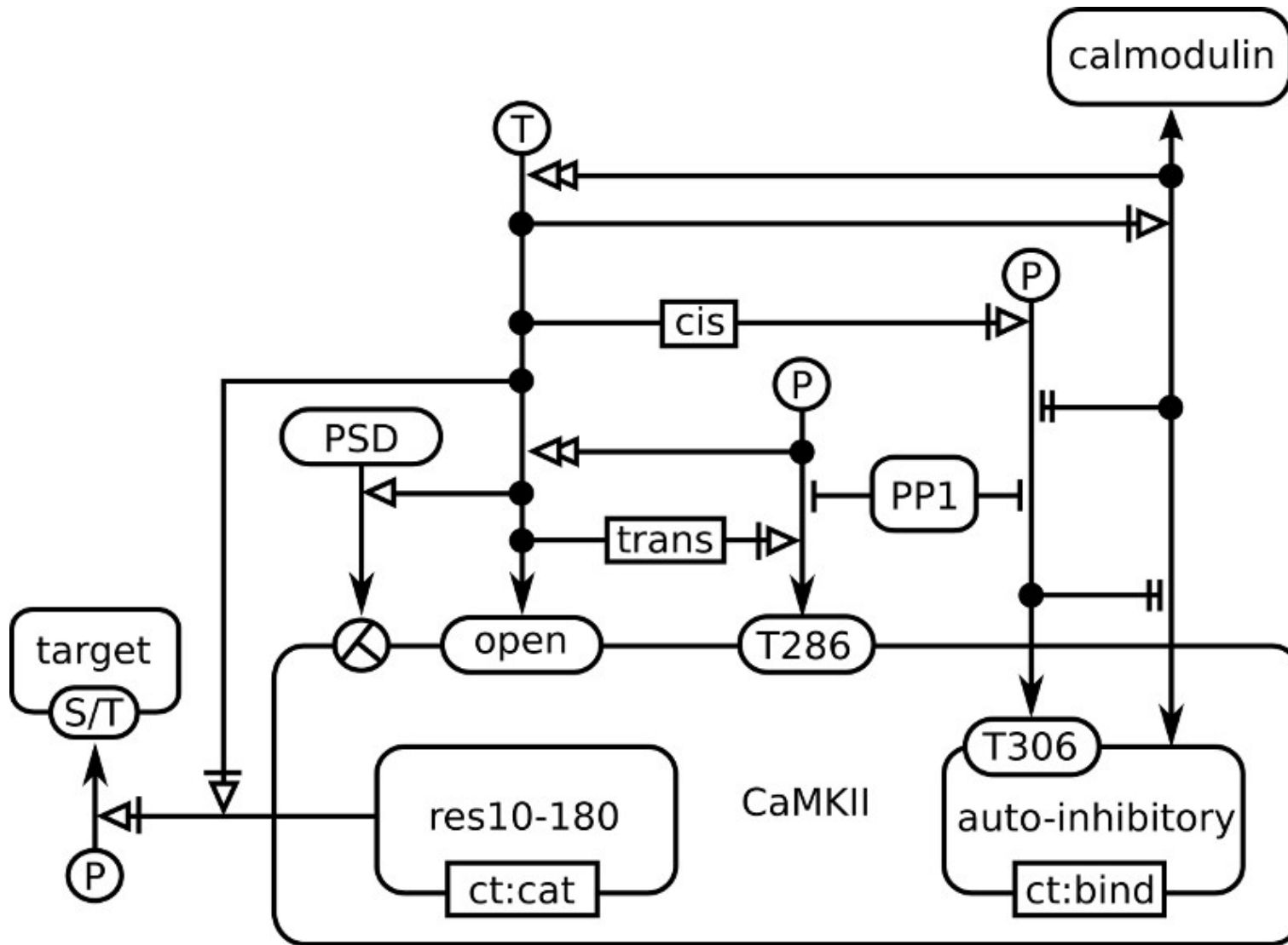


The value "membrane" is assigned to the "location" of domainB but not to domainB and entityX

degradation of domain A of X



Real example: CaMKII

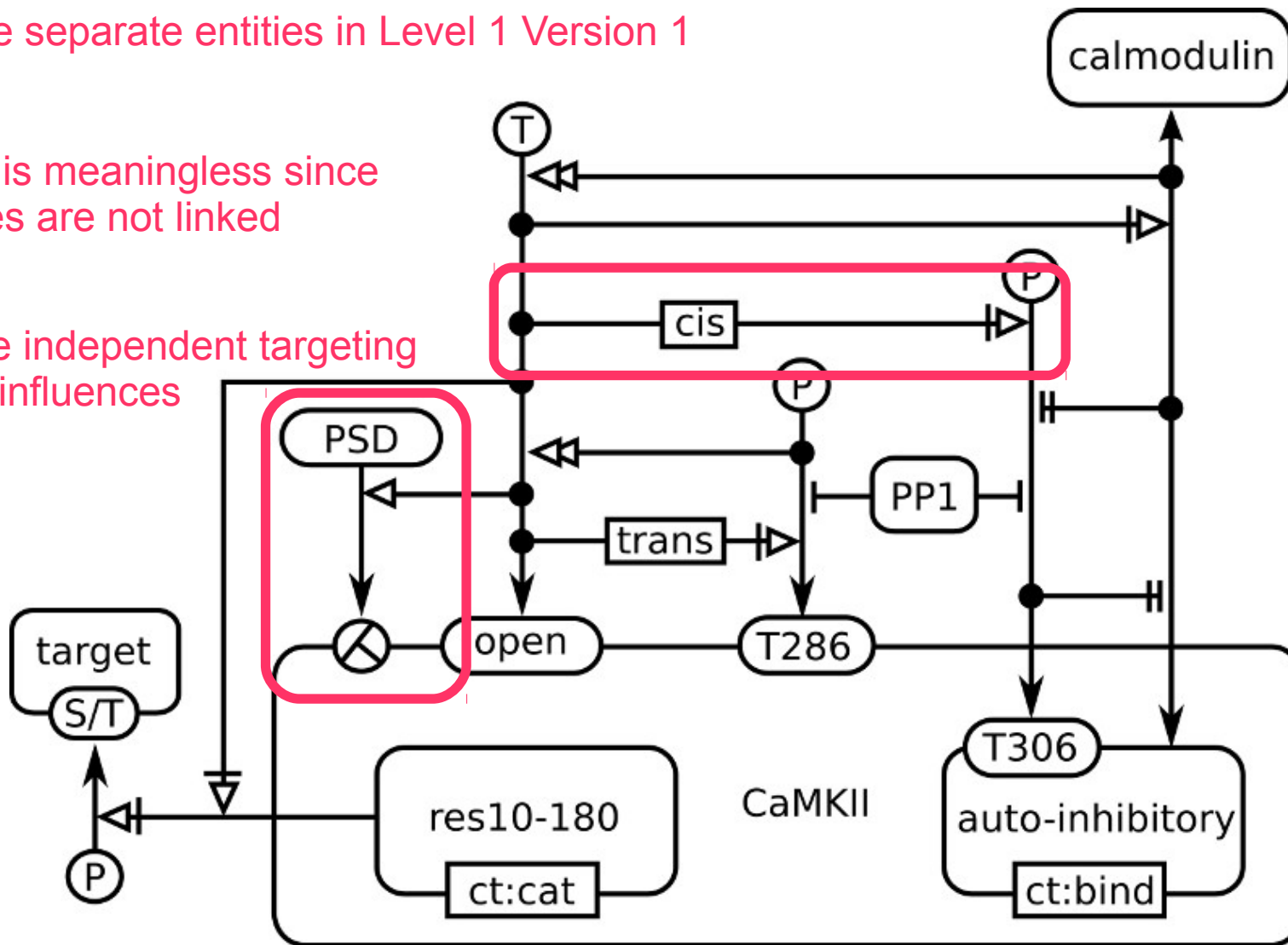


Real example: CaMKII

Three separate entities in Level 1 Version 1

“CIS” is meaningless since entities are not linked

Three independent targeting
With influences



Issues pending in HARMONY

- 1) Possibility to identify groups of entities and statements, for instance defining pathways [NLN]
- 2) Differentiating Entities (and outcomes) representing continuants and occurrents. Proposal is to follow PD and AF guidelines (round corners for continuants and acute ones for occurrents) [AL and NLN]
- 3) Outcomes on influences. Purpose is to differentiate the actualisation of an influence from the the effect of the influence [NLN]

Vote on groups

Question 1. Are-you in favour of introducing a "group" feature in SBGN languages?

Choice	Votes	Fraction
Yes	10	90.9%
No	1	9.1%
I do not know	0	0%.

Question 2: Should-we specify the way a group is displayed?

Choice	Votes	Fraction
Yes	7	63.6%
No	4	36.4%
I do not know	0	0%

Question 3: If we were to advise a way to represent groups, what should it be (multiple answers possible)?

Choice	Votes	Fraction
A spatial grouping	4	36.4%
A contour	7	63.6%
A background	8	72.7%
Highlighting glyps	4	36.4%
Unsure	2	18.2%
None of the above	0	0%

Vote on groups

Question 1. Are-you in favour of introducing a "group" feature in SBGN languages?

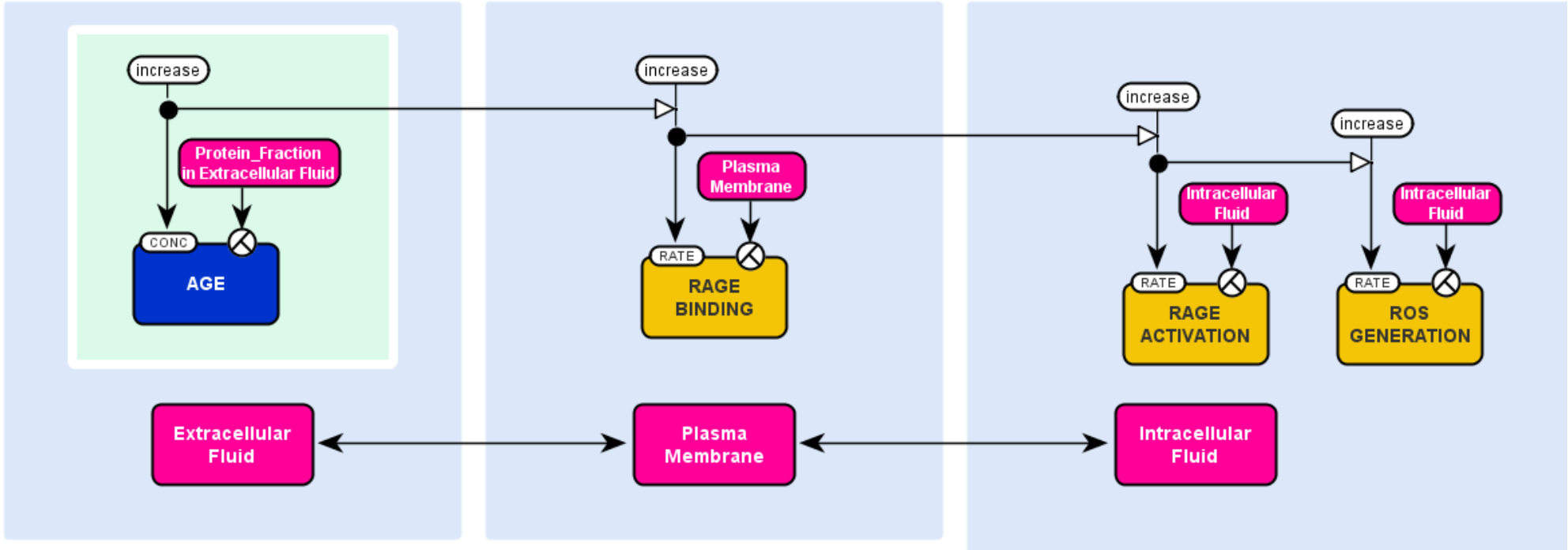
Choice	Votes	Fraction
Yes	10	90.9%
No	1	9.1%
I do not know	0	0%.

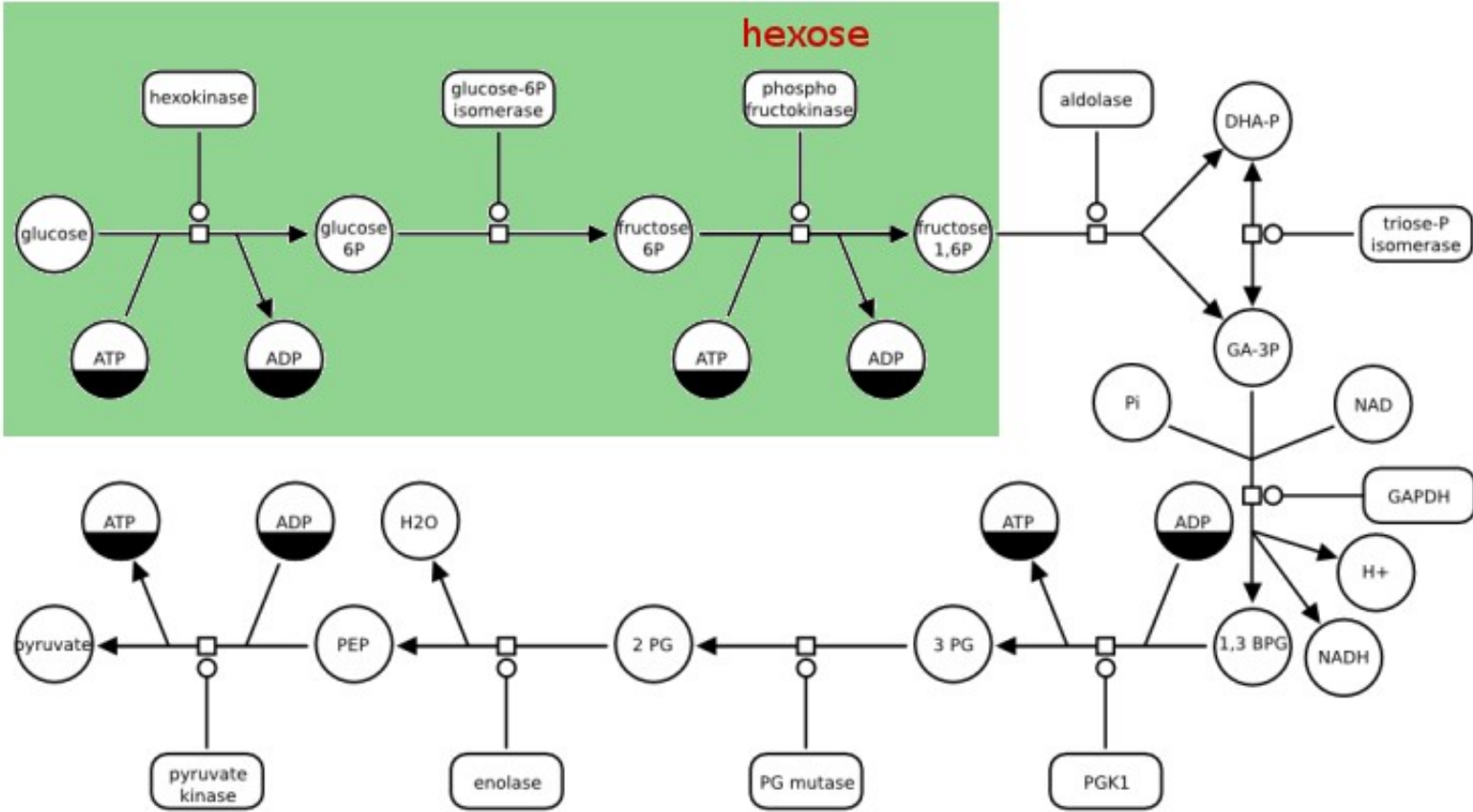
Question 2: Should-we specify the way a group is displayed?

Choice	Votes	Fraction
Yes	7	63.6%
No	4	36.4%
I do not know	0	0%

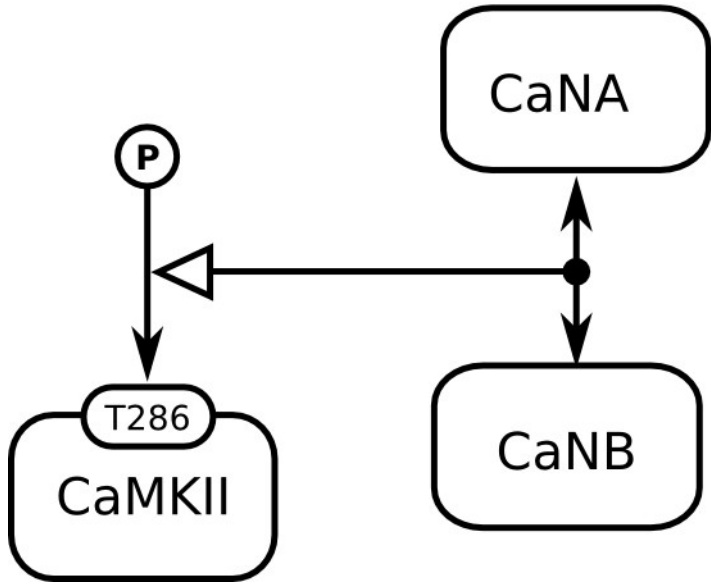
Question 3: If we were to advise a way to represent groups, what should it be (multiple answers possible)?

Choice	Votes	Fraction
A spatial grouping	4	36.4%
A contour	7	63.6%
A background	8	72.7%
Highlighting glyphs	4	36.4%
Unsure	2	18.2%
None of the above	0	0%

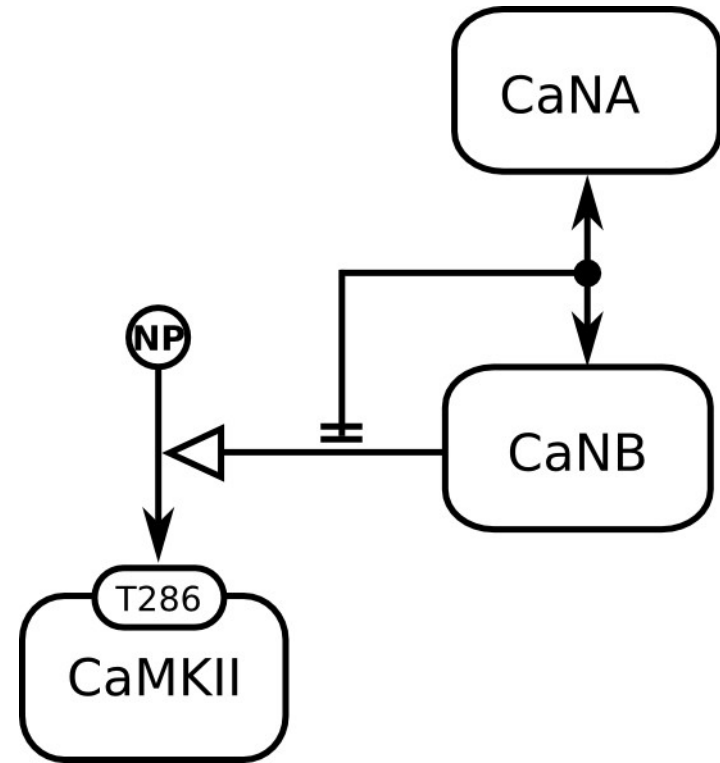
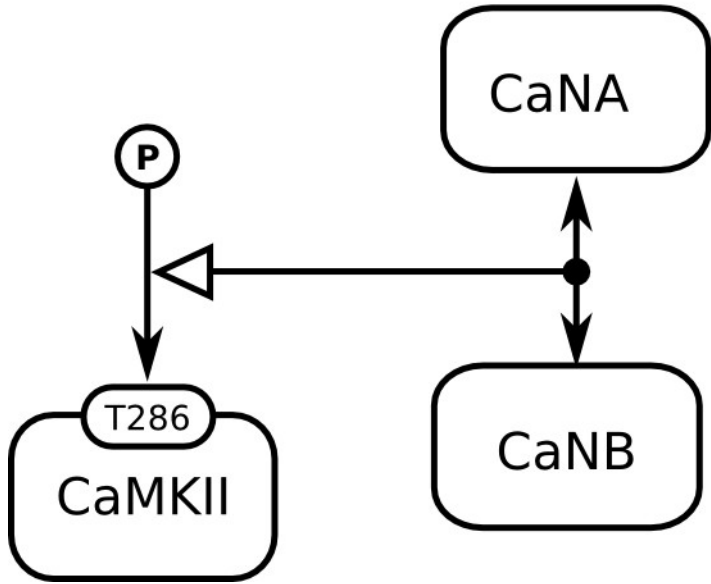




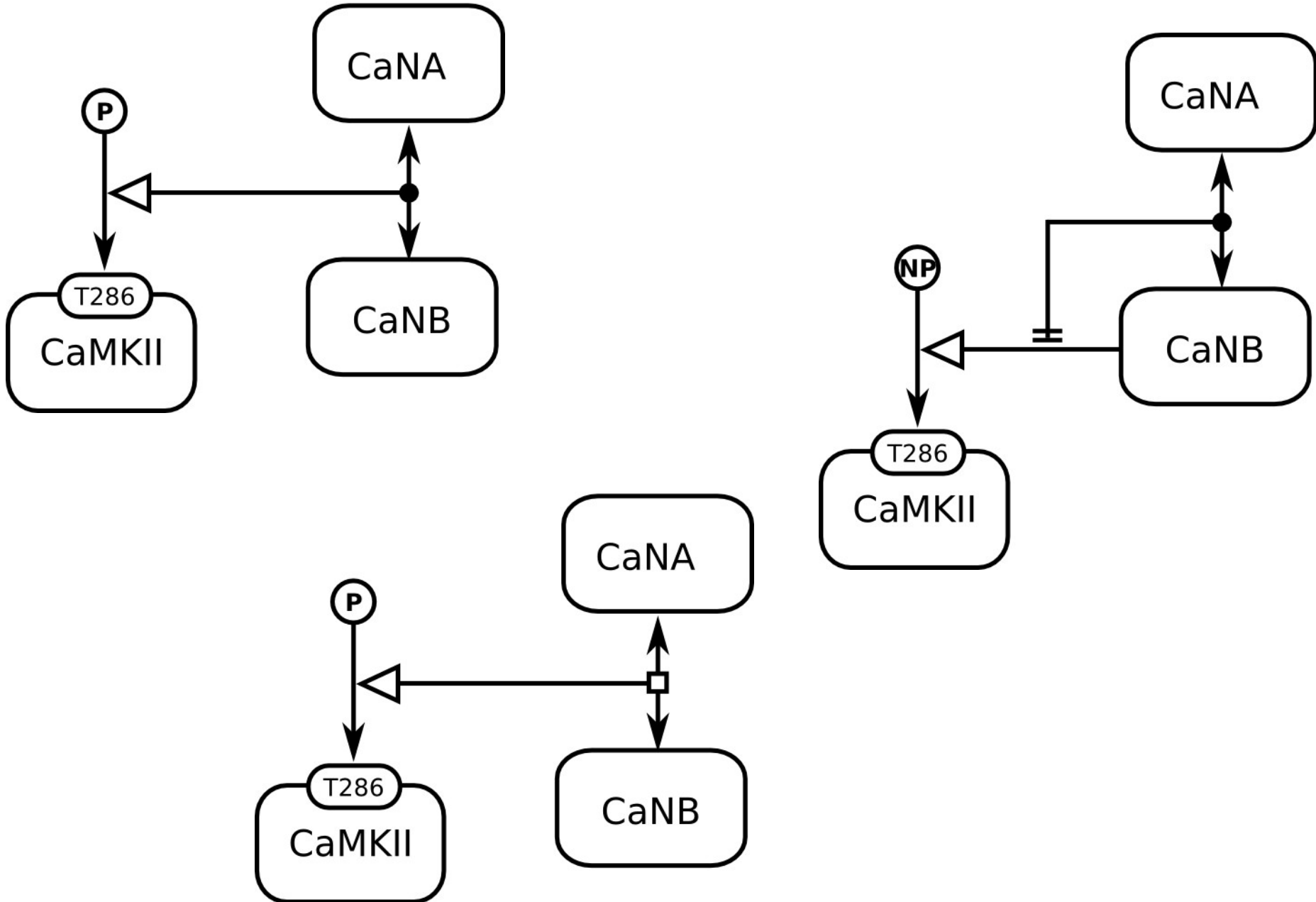
Continuant Vs occurrent outcomes



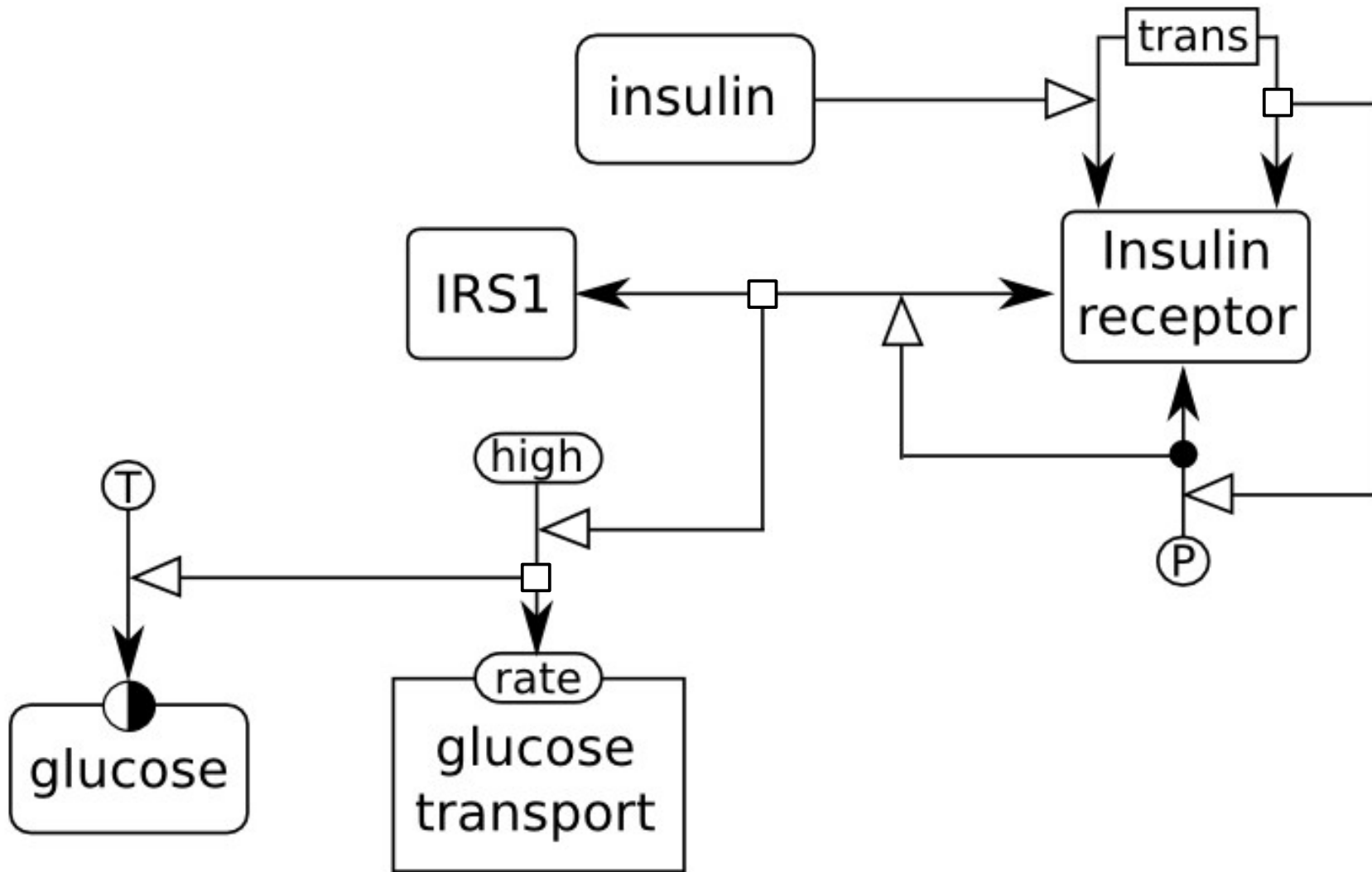
Continuant Vs occurrent outcomes



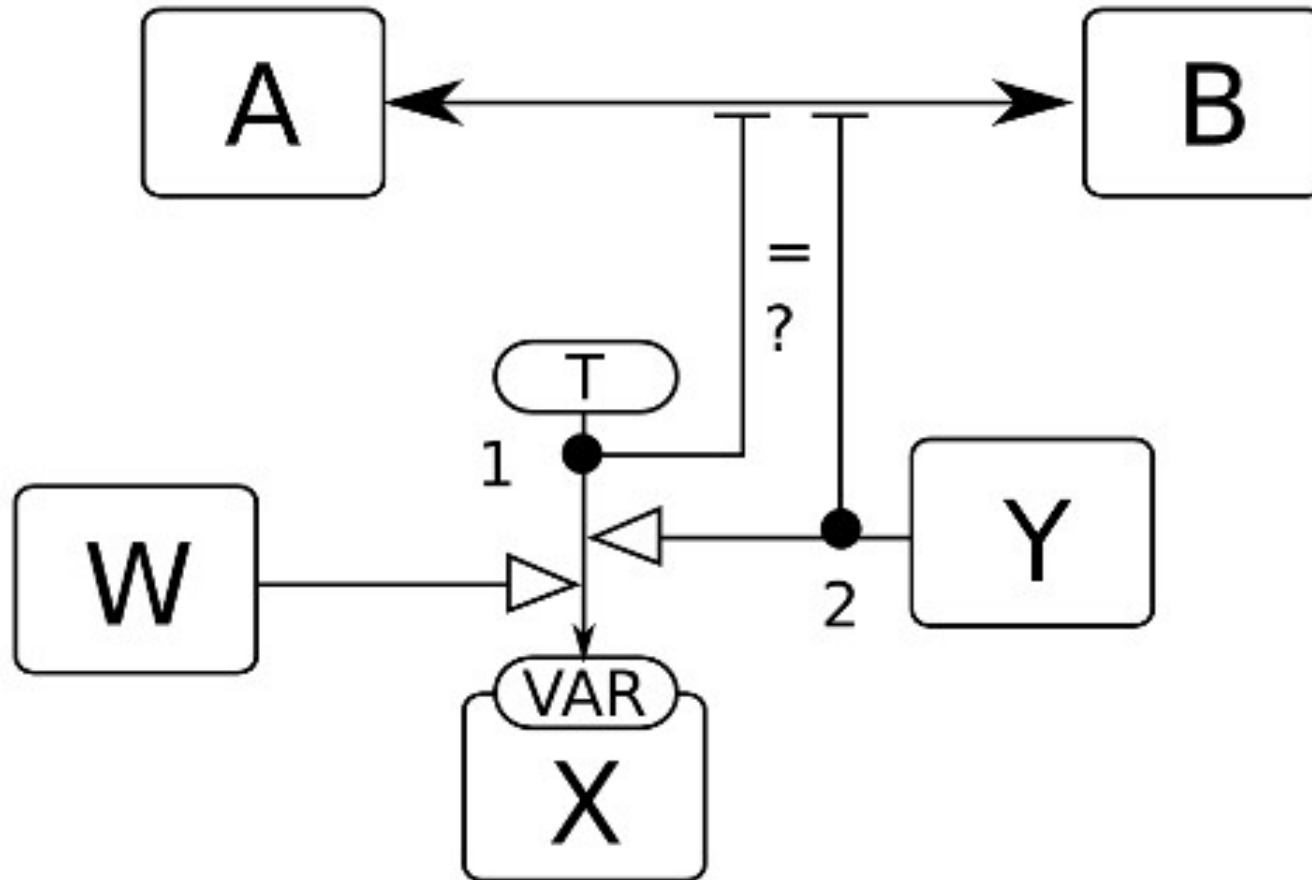
Continuant Vs occurrent outcomes



Continuant Vs occurrent entities



Outcome on influences



Better representation of rule-based models

Any other issue you want to discuss?