Model: Angiogenesis
Type: P/T Net
Origin: Academic

since
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Christian Rohr
christian.rohr@b-tu.de

This form is a summary description of the model entitled "Angiogenesis" proposed for the Model Checking Contest @ Petri Nets. Models can be given in several instances parameterized by scaling parameters. Colored nets can be accompanied by one or many equivalent, unfolded P/T nets. Models are given together with property files (possibly, one per model instance) giving a set of properties to be checked on the model.

Description

Angiogenesis, defined as the formation of new vessels from the existing ones, is a topic of great interest in all areas of human biology, particularly to scientists studying vascular development, vascular malformation and cancer biology. Angiogenesis is a complex process involving the activities of many growth factors and relative receptors, which trigger several signaling pathways resulting in different cellular responses. The Petri net was introduced in [1] and refined in [2].
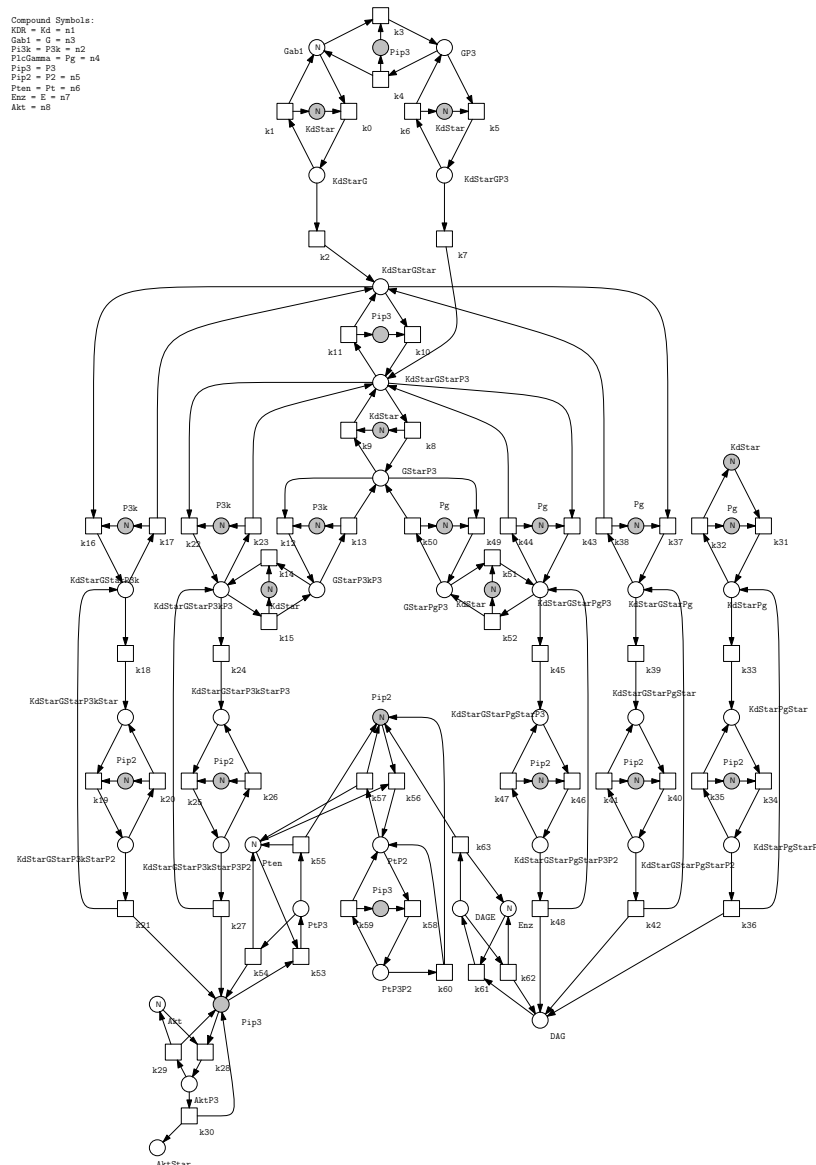
Graphical representation with parameter N. The gray coloured places are logic/fusion places.

References

1 L. Napione, D. Manini, F. Cordero, A. Horvath, A. Picco, M. D. Pierro, S. Pavan, M. Sereno, A. Veglio, F. Bussolino, and G. Balbo. On the Use of Stochastic Petri Nets in the Analysis of Signal Transduction Pathways for Angiogenesis Process. In Proc. CMSB 2009, pages 281295. LNCS/LNBI 5688, Springer, 2009.

2 F. Cordero, A. Horvath, D. Manini, L. Napione, M. D. Pierro, S. Pavan, A. Picco, A. Veglio, M. Sereno, F. Bussolino, and G. Balbo. Simplification of a complex signal transduction model using invariants and flow equivalent servers. Theor. Comput. Sci., 412(43):6036–6057, 2011.

Scaling parameter

Parameter name	Parameter description	Chosen parameter values
N	initial number of tokens on places Akt, Enz, Gab1, KdStar, P3k, Pg, Pip2 and Pten	1, 5, 10, 15, 20, 25, 50

Size of the model

Although the model is parameterized, its size does not depend on parameter values.

number of places:	39
number of transitions:	64
number of arcs:	185

Structural properties

ordinary — *all arcs have multiplicity one* .... ✔

simple free choice — *all (different) transitions with a shared input place have no other input place* .... ✗ (a)

state machine — *every transition has exactly one input place and exactly one output place* .... ✗ (b)

marked graph — *every place has exactly one input transition and exactly one output transition* .... ✗ (c)

connected — *there is an undirected path between every two nodes (places or transitions)* .... ✔ (d)

strongly connected — *there is a directed path between every two nodes (places or transitions)* .... ✗ (e)

source place(s) — *one or more places have no input transitions* .... ✗ (f)

sink place(s) — *one or more places have no output transitions* .... ✔ (g)

source transition(s) — *one or more transitions have no input places* .... ✗ (h)

sink transitions(s) — *one or more transitions have no output places* .... ✗ (i)

loop-free — *no transition has an input place that is also an output place* .... ✔ (j)

conservative — *for each transition, the number of input arcs equals the number of output arcs* .... ✗ (k)

subconservative — *for each transition, the number of input arcs equals or exceeds the number of output arcs* .... ✗ (l)

nested units — *places are structured into hierarchically nested sequential units* (m) .... ✗

(a) 40 arcs are not free choice, e.g., the arc from place "p6" (which has 2 outgoing transitions) to transition "t0" (which has 2 input places).

(b) 57 transitions are not of a state machine, e.g., transition "t0".

(c) 38 places are not of a marked graph, e.g., place "p5".

(d) stated by CÆSAR.BDD version 2.0 on all 7 instances (1, 5, 10, 15, 20, 25, and 50).

(e) from place "p2" one cannot reach place "p0".

(f) stated by CÆSAR.BDD version 2.0 on all 7 instances (1, 5, 10, 15, 20, 25, and 50).

(g) place "p2" is a sink place.

(h) stated by CÆSAR.BDD version 2.0 on all 7 instances (1, 5, 10, 15, 20, 25, and 50).

(i) stated by CÆSAR.BDD version 2.0 on all 7 instances (1, 5, 10, 15, 20, 25, and 50).

(j) stated by CÆSAR.BDD version 2.0 on all 7 instances (1, 5, 10, 15, 20, 25, and 50).

(k) 57 transitions are not conservative, e.g., transition "t0".

(l) 33 transitions are not subconservative, e.g., transition "t1".

(m) the definition of Nested-Unit Petri Nets (NUPN) is available from http://mcc.lip6.fr/nupn.php

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Behavioural properties

safe — *in every reachable marking, there is no more than one token on a place* .... ? [(n)]
deadlock — *there exists a reachable marking from which no transition can be fired* .... ✔ [(o)]
reversible — *from every reachable marking, there is a transition path going back to the initial marking* .... ✘ [(p)]
quasi-live — *for every transition t, there exists a reachable marking in which t can fire* .... ✘
live — *for every transition t, from every reachable marking, one can reach a marking in which t can fire* .... ✘ [(q)]

Size of the marking graphs

Parameter	Number of reachable markings	Number of transition firings	Max. number of tokens per place	Max. number of tokens per marking
$N = 1$	110 [(r)]	288 [(s)]	N [(t)]	8 [(u)]
$N = 5$	4.2735E+7 [(v)]	4.8687E+8 [(w)]	N [(x)]	40 [(y)]
$N = 10$	8.2265E+11 [(z)]	1.5636E+13 [(aa)]	N [(ab)]	80 [(ac)]
$N = 15$	1 115 538 966 669 107 [(ad)]	?	N	≥ 120
$N = 20$	351 820 047 967 344 849 [(ae)]	?	N	≥ 160
$N = 25$	43 090 329 340 850 957 348 [(af)]	?	N	≥ 200
$N = 50$	?	?	N	≥ 400

[(n)] stated by CÆSAR.BDD version 2.0 to be true on 1 instance(s) out of 7, and false on the remaining 6 instance(s).

[(o)] Checked by Marcie on 2013-12-13; confirmed at MCC'2014 by Tapaal on 2 instances and by Lola on 5 instances.

[(p)] has dead states.

[(q)] has dead states.

[(r)] given in [2] and computed by Marcie on 2013-12-13; confirmed by CÆSAR.BDD version 2.0; confirmed at MCC'2014 by Marcie, PNMC, PNXDD, Stratagem, and Tapaal.

[(s)] computed at MCC'2014 by Marcie.

[(t)] confirmed at MCC'20214 by Marcie, PNMC, and Tapaal.

[(u)] computed at MCC'2014 by Marcie, PNMC, and Tapaal.

[(v)] exact value 42 734 935 given in [2] and computed by Marcie on 2013-12-13; confirmed at MCC'2014 by Marcie, PNMC, and PNXDD.

[(w)] computed at MCC'2014 by Marcie.

[(x)] confirmed at MCC'2014 by Marcie and PNMC.

[(y)] computed at MCC'2014 by Marcie and PNMC.

[(z)] exact value 822 645 885 495 computed by Marcie on 2013-12-13; confirmed at MCC'2014 by Marcie, PNMC, and PNXDD.

[(aa)] computed at MCC'2014 by Marcie.

[(ab)] confirmed at MCC'2014 by Marcie and PNMC.

[(ac)] computed at MCC'2014 by Marcie and PNMC.

[(ad)] computed by Marcie on 2013-12-13.

[(ae)] computed by Marcie on 2013-12-13.

[(af)] computed by Marcie on 2013-12-13.

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