

# On the Importance of the Deadlock Trap Property for Monotonic Liveness

Monika Heiner<sup>1</sup>, Cristian Mahulea<sup>2</sup>, Manuel Silva<sup>2</sup>

<sup>1</sup> Department of Computer Science, Brandenburg University of Technology  
Postbox 10 13 44, 03013 Cottbus, Germany

`monika.heiner@tu-cottbus.de`

<sup>2</sup> Instituto de Investigación en Ingeniería de Aragón (I3A),  
Universidad de Zaragoza, Maria de Luna 1, E-50018 Zaragoza, Spain  
`cmahulea@unizar.es`, `silva@unizar.es`

**Abstract.** In Petri net systems, liveness is an important property capturing the idea of no transition (action) becoming non-fireable (unattainable). Additionally, in some situations it is particularly interesting to check if the net system is (marking) monotonically live, i.e., it remains live for any marking greater than the initial one. In this paper, we discuss structural conditions preserving liveness under arbitrary marking increase. It is proved that the deadlock trap property (DTP) is a necessary condition for liveness monotonicity of ordinary nets, and necessary and sufficient for some subclasses. We illustrate also how the result can be used to study liveness monotonicity for non-ordinary nets using a simulation preserving the firing language. Finally, we apply these conditions to several case studies of biomolecular networks.

## 1 Motivation

Petri nets are a natural choice to represent biomolecular networks. Various types of Petri nets may be useful – qualitative, deterministically timed, stochastic, continuous or hybrid ones, depending on the available information and the kind of properties to be analysed. Accordingly, the integrative framework demonstrated by several case studies in [GHR<sup>+</sup>08], [HGD08], [HDG10] applies a family of related Petri net models, sharing structure, but differing in their kind of kinetic information.

A key notion of the promoted strategy of biomodel engineering is the *level concept*, which has been introduced in the Petri net framework in [GHL07]. Here, a token stands for a specific amount of mass, defined by the total mass divided by the number of levels. Thus, increasing the token number to represent a certain amount of mass means to increase the resolution of accuracy.

This procedure silently assumes some kind of behaviour preservation while the marking is increased (typically multiplied by a factor) to represent a finer granularity of the mass flowing through the network. However, as it is well-known in Petri net theory, liveness is not monotonic with respect to (w.r.t.) the initial marking for general Petri nets. Thus, there is no reason to generally assume that

there is no significant change in the possible behaviour by marking increase. Contrary, under liveness monotonicity w.r.t. the initial marking we can expect continuization (fluidization) to be reasonable. However, only a particular kind of monotonicity seems to be needed for continuization: *homothetic liveness*, i.e., liveness preservation while multiplying the initial marking by  $k$  [RTS99], [SR02].

At structural level, (monotonic) liveness can be considered using transformation (reduction) rules [Ber86], [Sil85], [Mur89], [Sta90], the classical analysis for ordinary nets based on the Deadlock Trap Property (DTP) [Mur89], [Sta90], or the results of *Rank Theorems*, which are directly applicable to non-ordinary nets [TS96], [RTS98]. In this paper, we concentrate on the DTP, which will initially be used for ordinary net models, and later extended to non-ordinary ones.

This paper is organized as follows. We start off with recalling relevant notions and results of Petri net theory. Afterwards we introduce the considered subject by looking briefly at two examples, before turning to our main result yielding a necessary condition for monotonic liveness. We demonstrate the usefulness of our results for the analysis of biomolecular networks by a variety of case studies. We conclude with an outlook on open issues.

## 2 Preliminaries

We assume basic knowledge of the standard notions of place/transition Petri nets, see e.g. [DHP<sup>+</sup>93], [HGD08], [DA10]. To be self-contained we recall the fundamental notions relevant for our paper.

### Definition 1 (Petri net, syntax).

A Petri net is a tuple  $\mathcal{N} = \langle P, T, \mathbf{Pre}, \mathbf{Post} \rangle$ , and a Petri net system is a tuple  $\Sigma = \langle \mathcal{N}, \mathbf{m}_0 \rangle$ , where

- $P$  and  $T$  are finite, non-empty, and disjoint sets.  $P$  is the set of places.  $T$  is the set of transitions.
- $\mathbf{Pre}, \mathbf{Post} \in \mathbb{N}^{|P| \times |T|}$  are the pre- and post-matrices, where  $|\cdot|$  is the cardinality of a set, i.e., its number of elements. For a place  $p_i \in P$  and a transition  $t_j \in T$ ,  $\mathbf{Pre}(p_i, t_j)$  is the weight of the arc connecting  $p_i$  to  $t_j$  (0 if there is no arc), while  $\mathbf{Post}(p_i, t_j)$  is the weight of the arc connecting  $t_j$  to  $p_i$ .
- $\mathbf{m}_0 \in \mathbb{N}_{>0}^{|P|}$  gives the initial marking.
- $\mathbf{m}(p)$  yields the number of tokens on place  $p$  in the marking  $\mathbf{m}$ . A place  $p$  with  $\mathbf{m}(p) = 0$  is called empty (unmarked) in  $\mathbf{m}$ , otherwise it is called marked (non-empty). A set of places is called empty if all its places are empty, otherwise marked.
- The preset and postset of a node  $x \in P \cup T$  are denoted by  $\bullet x$  and  $x \bullet$ . They represent the input and output transitions of a place  $x$ , or the input and output places of a transition  $x$ . More specifically, if  $t_j \in T$ ,  $\bullet t_j = \{p_i \in P \mid \mathbf{Pre}(p_i, t_j) > 0\}$  and  $t_j \bullet = \{p_i \in P \mid \mathbf{Post}(p_i, t_j) > 0\}$ . Similarly, if  $p_i \in P$ ,  $\bullet p_i = \{t_j \in T \mid \mathbf{Post}(p_i, t_j) > 0\}$  and  $p_i \bullet = \{t_j \in T \mid \mathbf{Pre}(p_i, t_j) > 0\}$ . We extend both notions to a set of nodes  $X \subseteq P \cup T$  and define the set of all prenodes  $\bullet X := \bigcup_{x \in X} \bullet x$ , and the set of all postnodes  $X \bullet := \bigcup_{x \in X} x \bullet$ .

- A node  $x \in P \cup T$  is called source node, if  $\bullet x = \emptyset$ , and sink node if  $x \bullet = \emptyset$ . A boundary node is either a sink or a source node (but not both, because we assume a connected net).

**Definition 2 (Petri net, behaviour).** Let  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  be a net system.

- A transition  $t$  is enabled at marking  $\mathbf{m}$ , written as  $\mathbf{m}[t]$ , if  $\forall p \in \bullet t : \mathbf{m}(p) \geq \text{Pre}(p, t)$ , else disabled.
- A transition  $t$ , enabled in  $\mathbf{m}$ , may fire (occur), leading to a new marking  $\mathbf{m}'$ , written as  $\mathbf{m}[t]\mathbf{m}'$ , with  $\forall p \in P : \mathbf{m}'(p) = \mathbf{m}(p) - \text{Pre}(p, t) + \text{Post}(p, t)$ .
- The set of all markings reachable from a marking  $\mathbf{m}_0$ , written as  $[\mathbf{m}_0]$ , is the smallest set such that  $\mathbf{m}_0 \in [\mathbf{m}_0]$ ,  $\mathbf{m} \in [\mathbf{m}_0] \wedge \mathbf{m}[t]\mathbf{m}' \Rightarrow \mathbf{m}' \in [\mathbf{m}_0]$ .
- The reachability graph (RG) is a directed graph with  $[\mathbf{m}_0]$  as set of nodes, and the labelled arcs denote the reachability relation  $\mathbf{m}[t]\mathbf{m}'$ .

**Definition 3 (Behavioural properties).** Let  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  be a net system.

- A place  $p$  is  $k$ -bounded (bounded for short) if there is a positive integer number  $k$ , serving as an upper bound for the number of tokens on this place in all reachable markings of the Petri net:  $\exists k \in \mathbb{N}_0 : \forall \mathbf{m} \in [\mathbf{m}_0] : \mathbf{m}(p) \leq k$ .
- A Petri net system is  $k$ -bounded (bounded for short) if all its places are  $k$ -bounded.
- A transition  $t$  is dead at marking  $\mathbf{m}$  if it is not enabled in any marking  $\mathbf{m}'$  reachable from  $\mathbf{m}$ :  $\nexists \mathbf{m}' \in [\mathbf{m}] : \mathbf{m}'[t]$ .
- A transition  $t$  is live if it is not dead in any marking reachable from  $\mathbf{m}_0$ .
- A marking  $\mathbf{m}$  is dead if there is no transition which is enabled in  $\mathbf{m}$ .
- A Petri net system is deadlock-free (weakly live) if there are no reachable dead markings.
- A Petri net system is live (strongly live) if each transition is live.

**Definition 4 (Net structures).** Let  $\mathcal{N} = \langle P, T, \mathbf{Pre}, \mathbf{Post} \rangle$  be a Petri net.  $\mathcal{N}$  is

- Homogeneous (HOM) if  $\forall p \in P : t, t' \in p \bullet \Rightarrow \text{Pre}(p, t) = \text{Pre}(p, t')$ ;
- Ordinary (ORD) if  $\forall p \in P$  and  $\forall t \in T$ ,  $\text{Pre}(p, t) \leq 1$  and  $\text{Post}(p, t) \leq 1$ ;
- Extended Simple (ES) (sometimes also called asymmetric choice) if it is ORD and  $\forall p, q \in P : p \bullet \cap q \bullet = \emptyset \vee p \bullet \subseteq q \bullet \vee q \bullet \subseteq p \bullet$ ;
- Extended Free Choice (EFC) if it is ORD and  $\forall p, q \in P : p \bullet \cap q \bullet = \emptyset \vee p \bullet = q \bullet$ .

**Definition 5 (DTP).** Let  $\mathcal{N} = \langle P, T, \mathbf{Pre}, \mathbf{Post} \rangle$  be a Petri net.

- A siphon (structural deadlock, co-trap) is a non-empty set of places  $D \subseteq P$  with  $\bullet D \subseteq D \bullet$ .
- A trap is a non-empty set of places  $Q \subseteq P$  with  $Q \bullet \subseteq \bullet Q$ .
- A minimal siphon (trap) is a siphon (trap) not including a siphon (trap) as a proper subset.
- A bad siphon is a siphon, which does not include a trap.

- An empty siphon (trap) is a siphon (trap), not containing a token.
- The Deadlock Trap Property (DTP) asks for every siphon to include an initially marked trap, i.e., marked at  $\mathbf{m}_0$ .

The DTP can be reformulated as: minimal siphons are not bad and the maximal traps included are initially marked.

**Definition 6 (Semiflows).** Let  $\mathcal{N} = \langle P, T, \mathbf{Pre}, \mathbf{Post} \rangle$  be a net.

- The token flow matrix (or incidence matrix if the net is pure, i.e., self-loop free) is a matrix  $\mathbf{C} = \mathbf{Post} - \mathbf{Pre}$ .
- A place vector is a vector  $\mathbf{y} \in \mathbb{Z}^{|P|}$ ; a transition vector is a vector  $\mathbf{x} \in \mathbb{Z}^{|T|}$ .
- A P-semiflow is a place vector  $\mathbf{y}$  with  $\mathbf{y} \cdot \mathbf{C} = \mathbf{0}$ ,  $\mathbf{y} \geq \mathbf{0}$ ,  $\mathbf{y} \neq \mathbf{0}$ ;
- a T-semiflow is a transition vector  $\mathbf{x}$  with  $\mathbf{C} \cdot \mathbf{x} = \mathbf{0}$ ,  $\mathbf{x} \geq \mathbf{0}$ ,  $\mathbf{x} \neq \mathbf{0}$ .
- The support of a semiflow  $\mathbf{x}$ , written as  $\text{supp}(\mathbf{x})$ , is the set of nodes corresponding to the non-zero entries of  $\mathbf{x}$ .
- A net is conservative if every place belongs to the support of a P-semiflow.
- A net is consistent if every transition belongs to the support of a T-semiflow.
- In a minimal semiflow  $\mathbf{x}$ ,  $\text{supp}(\mathbf{x})$  does not contain the support of any other semiflow  $\mathbf{z}$ , i.e.,  $\nexists$  semiflow  $\mathbf{z} : \text{supp}(\mathbf{z}) \subset \text{supp}(\mathbf{x})$ , and the greatest common divisor of  $\mathbf{x}$  is 1.
- A mono-T-semiflow net (MTS net) is a consistent and conservative net that has exactly one minimal T-semiflow.

For convenience, we give vectors (markings, semiflows) in a short-hand notation by enumerating only the non-zero entries. Finally, we recall some well-known related propositions (see for example [Mur89], [Sta90]), which might be useful for the reasoning we pursue in this paper.

**Proposition 1 (Basics).**

1. An empty siphon remains empty forever. A marked trap remains marked for ever.
2. If  $R$  and  $R'$  are siphons (traps), then  $R \cup R'$  is also a siphon (trap).
3. A minimal siphon (trap) is a P-strongly-connected component, i.e., its places are strongly connected.
4. A deadlocked Petri net system has an empty siphon.
5. Each siphon of a live net system is initially marked.
6. If there is a bad siphon, the DTP does not hold.
7. A source place  $p$  establishes a bad siphon  $D = \{p\}$  on its own, and a sink place  $q$  a trap  $Q = \{q\}$ .
8. If each transition has a pre-place, then  $P \bullet = T$ , and if each transition has a post-place, then  $\bullet P = T$ . Thus, in a net without boundary transitions, the whole set of places is a siphon as well as a trap (however, not necessarily minimal ones).
9. For a P-semiflow  $\mathbf{x}$  it holds  $\bullet \text{supp}(\mathbf{x}) = \text{supp}(\mathbf{x}) \bullet$ . Thus, the support of a P-semiflow is siphon and trap as well (however, generally not vice versa).

**Proposition 2 (DTP and behavioural properties).**

1. An ordinary Petri net without siphons is live.
2. If  $\mathcal{N}$  is ordinary and the DTP holds for  $\mathbf{m}_0$ , then  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  is deadlock-free.
3. If  $\mathcal{N}$  is ES and the DTP holds for  $\mathbf{m}_0$ , then  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  is live.
4. Let  $\mathcal{N}$  be an EFC net.  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  is live iff the DTP holds.

We conclude this section with a proposition from [CCS91], which might be less known.

**Proposition 3 (MTS net and behavioural properties).** *Liveness and deadlock-freeness coincide in mono-T-semiflow net systems.*

**3 Monotonic Liveness**

If a property holds for a Petri net  $\mathcal{N}$  with the marking  $\mathbf{m}_0$ , and it also holds in  $\mathcal{N}$  for any  $\mathbf{m} \geq \mathbf{m}_0$ , then it is said to be *monotonic* in the system  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$ . In this paper we are especially interested in monotonic liveness.

**Definition 7 (Monotonic liveness).**

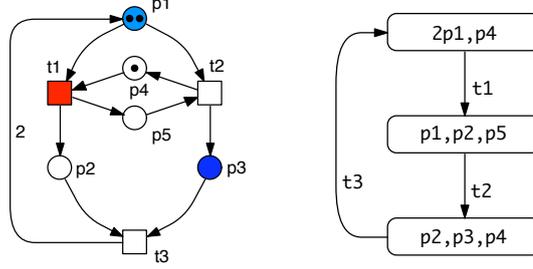
Let  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  be a Petri net system. It is called *monotonically live*, if being live for  $\mathbf{m}_0$ , it remains live for any  $\mathbf{m} \geq \mathbf{m}_0$ .

We are looking for conditions, at best structural conditions, preserving liveness under arbitrary marking increase. To illustrate the problem, let's consider a classical example [Sta90], [SR02].

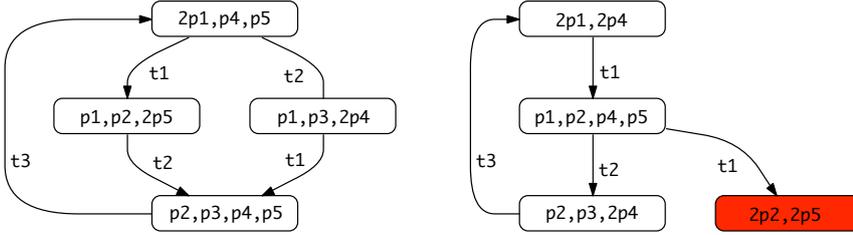
*Example 1.* The net  $\mathcal{N}$  in Figure 1 is ES, conservative, consistent, and covered by one T-semiflow. It is live for the given initial marking  $\mathbf{m}_1 = (2p_1, p_4)$ . Adding a token to place  $p_5$  yields the initial marking  $\mathbf{m}_2 = (2p_1, p_4, p_5)$  and the net system remains live for  $\mathbf{m}_2 \geq \mathbf{m}_1$ . However, adding a token to  $p_4$  yields the initial marking  $\mathbf{m}_3 = (2p_1, 2p_4)$  and the net behaviour now contains finite firing sequences, i.e., it can run into a deadlock (dead state). Thus, the net system is not live for  $\mathbf{m}_3 \geq \mathbf{m}_1$ . It is not monotonically live.

How to distinguish both cases? The net has two (minimal) bad siphons  $D_1 = \{p_1, p_2\}$  and  $D_2 = \{p_1, p_3\}$ . There is no chance to prevent these siphons from getting empty for arbitrary markings.  $D_1$  can potentially be emptied by firing  $t_2 \in D_1 \bullet \setminus \bullet D_1$ , and  $D_2$  by firing  $t_1 \in D_2 \bullet \setminus \bullet D_2$ . The latter case destroyed the liveness for  $\mathbf{m}_3$  as it will equally occur for all initial markings allowing transition sequences containing one of the troublemakers, in this example  $t_1$  and  $t_2$ , sufficiently often.  $\square$

One lesson learnt from the previous example is, a net does not have to make use of the additional tokens. Thus, all behaviour (set of transition sequences), which is possible for  $\mathbf{m}$  is still possible for  $\mathbf{m}'$ , with  $\mathbf{m} \leq \mathbf{m}'$ . However, new tokens may allow for additional system behaviour, which is actually well-known in Petri net theory, see Proposition 4.



**Fig. 1.** A mono-T-semiflow and ES Petri net  $\mathcal{N}$  and its reachability graph for the marking  $\mathbf{m}_1 = (2p_1, p_4)$ , generating the language  $\mathcal{L}_{\mathcal{N}}(\mathbf{m}_1) = (t_1 t_2 t_3)^* \{\varepsilon, t_1, t_1 t_2\}$ . The siphon  $\{p_1, p_3\}$  does not contain a trap, i.e., it is a bad siphon. If the initial marking is increased, it can potentially become empty by firing of  $t_1$ .

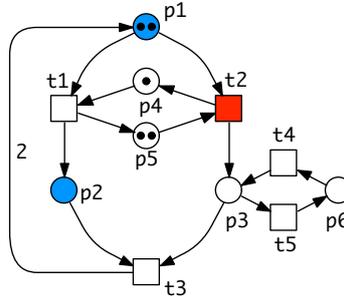


**Fig. 2.** Two other reachability graphs for the net  $\mathcal{N}$  in Figure 1 for the initial markings  $\mathbf{m}_2 = (2p_1, p_4, p_5)$  and  $\mathbf{m}_3 = (2p_1, 2p_4)$ ; both are greater than  $\mathbf{m}_1$ . Obviously,  $\mathcal{L}_{\mathcal{N}}(\mathbf{m}_1) \subset \mathcal{L}_{\mathcal{N}}(\mathbf{m}_2)$ ,  $\mathcal{L}_{\mathcal{N}}(\mathbf{m}_1) \subset \mathcal{L}_{\mathcal{N}}(\mathbf{m}_3)$ , but  $\langle \mathcal{N}, \mathbf{m}_3 \rangle$  is not live while  $\langle \mathcal{N}, \mathbf{m}_1 \rangle$  is live.

**Proposition 4.** For any net  $\mathcal{N}$  and two markings  $\mathbf{m}$  and  $\mathbf{m}'$ , with  $\mathbf{m} \leq \mathbf{m}'$ , it holds  $\mathcal{L}_{\mathcal{N}}(\mathbf{m}) \subseteq \mathcal{L}_{\mathcal{N}}(\mathbf{m}')$  [BRA83]; nevertheless,  $\langle \mathcal{N}, \mathbf{m} \rangle$  may be live while  $\langle \mathcal{N}, \mathbf{m}' \rangle$  not.

Example 1 is a mono-T-semiflow net, i.e., a net, where liveness and deadlock-freeness coincide (see Proposition 3). We look briefly at Example 2 to understand that this does not generally hold if there are several T-semiflows breathing life into the net.

*Example 2.* The net  $\mathcal{N}$  in Figure 3 is a slight extension of Example 1. It is ES, conservative, consistent and covered by two T-semiflows:  $\mathbf{x}_1 = (t_1, t_2, t_3)$ ,  $\mathbf{x}_2 = (t_4, t_5)$ . It is live for the initial marking  $\mathbf{m}_1 = (2p_1, p_4)$ .



**Fig. 3.** An ES Petri net which is not mono-T-semiflow. It is live for the initial marking  $\mathbf{m}_1 = (2p_1, p_4)$ . The siphon  $\{p_1, p_2\}$  is bad. So it can potentially become empty by firing  $t_2$  sufficiently often. This happens for the initial marking  $\mathbf{m}_2 = (2p_1, p_4, 2p_5)$ , making the net non-live, however keeping it deadlock-free (observe that  $\{p_3, p_6\}$  behaves as a trap if the firing of  $t_3$  is blocked forever).

The net has the following minimal siphons  $D_1 = \{p_1, p_2\}$ ,  $D_2 = \{p_1, p_3, p_6\}$ , and  $D_3 = \{p_4, p_5\}$ ; the first two are bad siphons. With the initial marking  $\mathbf{m}_2 = (2p_1, p_4, 2p_5)$ ,  $D_1$  can become empty by firing twice  $t_2 \in D_1 \bullet \setminus \bullet D_1$ , which destroys the liveness, without causing a dead state. The transitions  $t_4, t_5$  are live, the others not. Thus, the net system is not live, but deadlock-free.  $\square$

The loss of liveness is not necessarily monotonic itself; i.e., a net may be live for  $\mathbf{m}_1$ , non-live for a marking  $\mathbf{m}_2$  with  $\mathbf{m}_2 \geq \mathbf{m}_1$ , and live again for a marking  $\mathbf{m}_3$  with  $\mathbf{m}_3 \geq \mathbf{m}_2$  (which works for all examples in this paper). Liveness may also be lost by marking multiples (*homothetic markings*). Examples 1 and 2 are homothetically live, Example 3 in Section 5 not.

#### 4 Monotonic Liveness of Ordinary Nets

Let us turn to liveness criteria suitable for our objective looking at *ordinary* nets first. Liveness criteria not relying on the marking obviously ensure monotonic liveness. Unfortunately, there are only a few.

First of all, there are some structural reduction rules, see, e.g., [Sil85], [Ber86], [Mur89], [SR99]. To give a sample, the following reduction rule is easy to accept: a source transition is live, and all its post-places are unbounded. The transition and its post-places can be deleted (for analysis purposes); the reduction can be iterated as many times it is applicable. Sometimes, this kind of reasoning allows to decide liveness (for examples, see Section 6).

Besides structural reduction we have the DTP, which in most cases does depend on the marking, but it is obviously monotonic w.r.t. the marking: if each siphon contains a marked trap at  $\mathbf{m}$ , then – of course – it contains a marked

trap at  $\mathbf{m}' \geq \mathbf{m}$ . Thus, the DTP-related conclusions on behavioural properties in Proposition 2 are monotonic as well:

**Proposition 5 (Monotonic DTP).**

1. *An ordinary net without siphons is monotonically live.*
2. *An ordinary net system which holds the DTP is monotonically deadlock-free.*
3. *A live ES net system which holds the DTP is monotonically live.*
4. *An EFC net system is monotonically live iff the DTP holds.*

Proposition 5.1 can be considered as a special case of the DTP. Then, there must be source transitions (see Proposition 1.8), and the net is not strongly connected and not bounded.

**Lemma 1.** *Let be  $\mathcal{N}$  an ordinary Petri net. If  $\mathcal{N}$  is monotonically live, then there are no bad siphons.*

*Proof.* We will prove its reverse – if there exist a bad siphon, then the net system is not monotonically live – by contradiction. Let  $P_S$  be a bad siphon. Then there exist troublemaking transitions  $\Theta_i \in P_S \bullet \setminus \bullet P_S$ . There must be such transitions, because otherwise  $P_S \bullet = \bullet P_S$ , and then the siphon  $P_S$  would be a trap as well.

Since the net system is monotonically live, the marking of the places  $P \setminus P_S$  can be increased in such a way that it will never restrict the firing of the transitions  $P_S \bullet$ , i.e., the transitions depending on the siphon. Therefore, we can consider the subnet restricted to  $P_S$  in isolation.

We will show that the subsystem restricted to  $P_S$  can be emptied eventually by increasing the marking, hence cannot be monotonically live. Obviously we can assume that  $P_S$  is a minimal siphon. We consider two cases.

(1) The siphon has no forks ( $t_j$  is a *fork* if  $|t_j \bullet| > 1$ ). Based on the P-strongly-connectedness (see Proposition 1.3), there exists at least one path from each place  $p \in P_S$  to one of the troublemakers  $\Theta_i$ . Moving a token from  $p$  to  $\bullet \Theta_i$  does not increase the marking of any other place of  $P_S$  not belonging to the considered path. Obviously, this path can contain joins ( $t_j$  is a *join* if  $|\bullet t_j| > 1$ ), but we can add any tokens that are missing in the input places of the join. Firing the join, the marking of the places in the siphon is not increased. Using this process we can move the tokens from any  $p \in P_S$  to some  $\bullet \Theta_i$ , and by firing  $\Theta_i$  when it is enabled,  $P_S$  can be emptied. Thus, the net system can not be monotonically live.

(2) On the contrary, let us assume that there exists at least one fork  $t_j$  and let  $p_1, p_2 \in t_j \bullet$  be its output places. For the same reason as discussed in case (1), there exists a directed path from both places to one or several troublemakers. If all paths from  $p_1$  to any troublemaker  $\Theta_i$  contain  $t_j$ , then they form a trap. This is impossible because siphons are assumed to be bad. By symmetry, in the case in which the paths from  $p_2$  to troublemakers contain  $t_j$ , there exists a trap as well.

Finally, let us assume that there exists a path from  $p_1$  to a troublemaker  $\Theta_i$  and one path from  $p_2$  to a troublemakers  $\Theta_k$ , none of them containing  $t_j$ .

On both paths the same kind of reasoning can be applied (in an iterative way if several forks appear). Therefore, the siphon can be emptied even if firing  $t_j$  increases the tokens in  $P_S$ .  $\square$

Lemma 1 helps to preclude monotonic liveness for Examples 1 and 2 as well as for all other non-monotonically live examples we are aware of.

**Theorem 1.** *Let be  $\mathcal{N}$  an ordinary Petri net. If  $\langle \mathcal{N}, \mathbf{m}_0 \rangle$  is monotonically live, then the DTP holds.*

*Proof.* The structural check of the DTP can have three possible outcomes.

1. If there are no siphons, then the DTP holds trivially and the net is monotonically live (see Proposition 5.1).
2. If there are bad siphons, then the DTP does not hold for any initial marking and the net is not monotonically live (see Lemma 1).
3. If each siphon includes a trap, then the *maximal* trap  $P_T$  in every minimal siphon  $P_S$  has to be initially marked to fulfill the DTP. Because we assume liveness of the net system, there has to be at least one token in each minimal siphon (see Proposition 1.5). Let us assume that a token is not in  $P_T$ , but in a place  $p \in P_S \setminus P_T$ . If there exists at least one path without forks from  $p$  to a troublemaking transition  $\Theta_i \in P_S \bullet \setminus \bullet P_S$  not containing any transition belonging to the trap,  $\bullet P_T$ , then  $p$  can be emptied using the same reasoning as used in the proof of Lemma 1, case (1). Therefore the net can not be live. If the path from  $p$  to a troublemaking transition  $\Theta_i \in P_S \bullet \setminus \bullet P_S$  contains a fork, then the output places of the fork will be marked when  $p$  is emptied, and the paths from the output places of the forks to the output should be considered separately.

Finally, if *all* paths from  $p$  to the troublemaking transitions contain at least one transition  $\bullet P_T$ , then the trap  $P_T$  is not maximal since  $P_T$  together with all places belonging to the above mentioned paths (including all non-minimal ones) from  $p$  to transitions  $\bullet P_T$  are also a trap.  $\square$

According to Theorem 1, the DTP establishes a necessary condition for monotonic liveness, which complements Proposition 5.3.

**Corollary 1.** *A live ES net system is monotonically live iff the DTP holds.*

Moreover, for those systems for which deadlock-freeness is equivalent to liveness, the DTP is a sufficient criteria for liveness monotonicity. This leads, for example, to the following theorem:

**Theorem 2.** *Let be  $\mathcal{N}$  an ordinary mono-T-semiflow Petri net which for  $\mathbf{m}_0$  fulfills the DTP. Then the system  $\langle \mathcal{N}, \mathbf{m} \rangle$  is live for any  $\mathbf{m} \geq \mathbf{m}_0$ .*

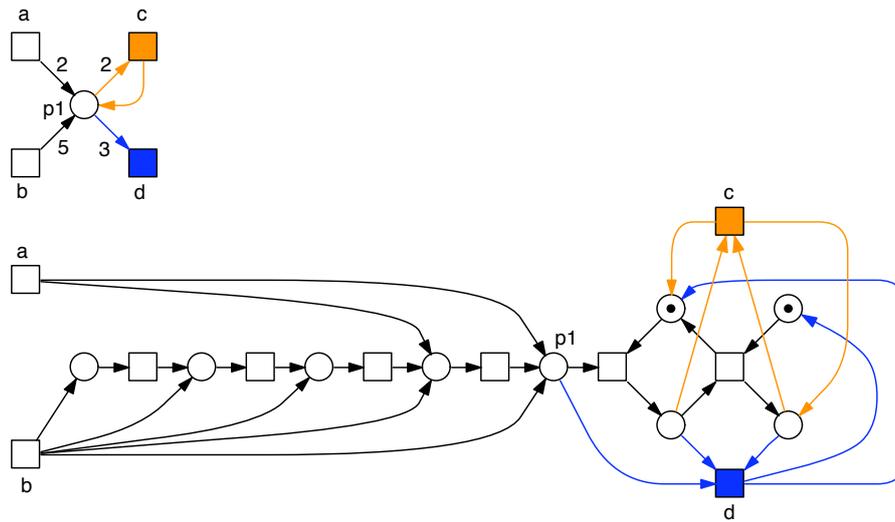
*Proof.* It follows from Proposition 5.2 (DTP and deadlock-freeness monotonicity) and Proposition 3 (equivalence of liveness and deadlock freeness in

mono-T-semiflow net systems).  $\square$

Therefore, the DTP is a sufficient criterion for monotonic liveness of ordinary mono-T-semiflow net systems as well. In summary, while the DTP is in general neither necessary nor sufficient for liveness, it turns out to be the case to keep alive ordinary ES nets or ordinary mono-T-semiflow nets under any marking increase.

## 5 Monotonic Liveness of Non-ordinary Nets

It is well-known that non-ordinary nets can be simulated under interleaving semantics by ordinary ones [Sil85] (see Figure 4 for an example). Let us look on the net structures we get by this simulation to learn how far the results for ordinary nets of Section 4 can be uplifted to non-ordinary nets.



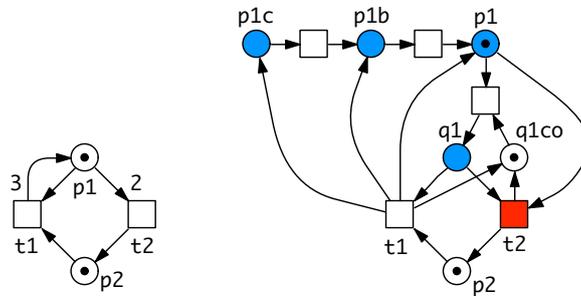
**Fig. 4.** A general principle to *simulate* a non-ordinary net system by an ordinary net system (here, the firing language of the second net projected on  $\{a, b, c, d\}$  is always equal to that of the first) [Sil85].

*Example 3.* We take a non-ordinary net from [SR02] and consider its simulation by an ordinary net, which we construct according to the general principle demonstrated in Figure 4.

The two net systems in Figure 5 are conservative, consistent, and live for the given initial marking. The ordinary net on the right hand side is not ES, and it

has two minimal bad siphons  $\{q_1, p_1, p_{1b}, p_{1c}\}$ ,  $\{p_2, p_1, p_{1b}, p_{1c}\}$ . Thus, according to Lemma 1, it is not monotonically live. Because our simulation preserves the projection of the firing language, in particular, preserves monotonicity of liveness. Thus, we conclude that the model on the left hand side is not monotonically live. Indeed, both nets are not live for any initial marking with an even number of tokens in  $p_1$ , but live for infinitely many other markings greater than or equal to  $(1, 1)$ .  $\square$

As a consequence of firing simulation by the ordinary net systems of the non-ordinary ones (preserving always the markings of the places involved in the head of the tail and complement, here  $q_1$  and  $q_{1co}$ ), liveness monotonicity can be studied on the ordinary simulation.



**Fig. 5.** A non-ordinary Petri net system and its simulation by an ordinary one. Both systems are non-live for any initial marking with an even number of tokens in  $p_1$ , and live for any other odd marking. Note that the markings of  $q_1$  and  $q_{1co}$  should not be increased in order to keep the language simulation in the right hand model. The net system on the right has a bad siphon  $\{q_1, p_1, p_{1b}, p_{1c}\}$  that can potentially become empty by firing  $t_2$  sufficiently often.

## 6 Applications

We consider a variety of test cases of our benchmark repository to demonstrate the helpfulness of the DTP for biomolecular networks. The following list sketches some basic characteristics. The essential analysis results are summarized in Table 1. All models hold the DTP, they are consistent and (supposed to be) live. For non-ordinary nets, the DTP refers to its simulation by an ordinary one.

1. **Apoptosis** (size: 37 places, 45 transitions, 89 arcs) is a signal transduction network, which governs complex mechanisms to control and execute genetically programmed cell death in mammalian cells. Disturbances in the apoptotic processes may lead to various diseases. This essential part of normal physiology for most metazoan species is not really well understood; thus

there exist many model versions. The validation by Petri net invariants of the model considered here is discussed in [HKW04], [HK04].

2. **RKIP** (size: 11 places, 11 transitions, 14 arcs) models the core of the influence of the Raf-1 Kinase Inhibitor Protein (RKIP) on the Extracellular signal Regulated Kinase (ERK) signalling pathway. It is one of the standard examples used in the systems biology community. It has been introduced in [CSK<sup>+</sup>03]; the corresponding qualitative, stochastic, continuous Petri nets are scrutinized in [GH06], [HDG10].
3. **Biosensor** (size: 6 places, 10 transitions, 21 arcs) is a gene expression network extended by metabolic activity. The model is a general template of a biosensor, which can be instantiated to be adapted to specific pollutants. It is considered as qualitative, stochastic, and continuous Petri net in [GHR<sup>+</sup>08] to demonstrate a model-driven design of a self-powering electrochemical biosensor.
4. **Hypoxia** (size: 14 places, 19 transitions, 56 arcs) is one of the well-studied molecular pathways activated under hypoxia condition. It models the Hypoxia Induced Factor (HIF) pathway responsible for regulating oxygen-sensitive gene expression. The version considered here is discussed in [YWS<sup>+</sup>07]; the corresponding qualitative and continuous Petri nets are used in [HS10] to determine the core network.
5. **Lac operon** (size: 11 places, 17 transitions, 41 arcs) is a classical example of prokaryotic gene regulation. We re-use the simplified model discussed in [Wil06]. Its corresponding stochastic Petri net is considered in [HLGM09].
6. **G/PPP** (size: 26 places, 32 transitions, 76 arcs) is a simplified model of the combined glycolysis (G) and pentose phosphate pathway (PPP) in erythrocytes (red blood cells). It belongs to the classical examples of biochemistry textbooks, see e.g. [BTS02], and thus of systems biology as well. The model was first discussed using Petri net technologies in [Red94]. Its validation by Petri net invariants is shown in [HK04], and a more exhaustive qualitative analysis in [KH08].
7. **MAPK** (size: 22 places, 30 transitions, 90 arcs) models the signalling pathway of the mitogen-activated protein kinase cascade, published in [LBS00]. It is a three-stage double phosphorylation cascade; each phosphorylation/dephosphorylation step applies the mass action kinetics pattern. The corresponding qualitative, stochastic, and continuous Petri net are scrutinized in [GHL07], [HGD08].
8. **CC – Circadian clock** (size: 14 places, 16 transitions, 58 arcs) refers to the central time signals of a roughly 24-hour cycle in living entities. Circadian rhythms are used by a wide range of organisms to anticipate daily changes in the environment. The model published in [BL00] demonstrates that circadian network can oscillate reliably in the presence of stochastic biomolecular noise and when cellular conditions are altered. It is also available as PRISM model on the PRISM website (<http://www.prismmodelchecker.org>). Its corresponding stochastic Petri net belongs to the benchmark suite used in [SH09]. We consider here a version with inhibitor arcs modelled by co-places.

9. **Halo** (size: 37 places, 38 transitions, 138 arcs) is a cellular signaling and regulation network, describing the phototaxis in the halobacterium salinarum [NMOG03]. It models the sophisticated survival strategy, which the halobacterium developed for harsh conditions (high temperature, high salt). A light sensing system and flagellar motor switching allows the cells to swim to those places of their habitat where the best light conditions are available. The model is the result of prolonged investigations by experimentally working scientists [Mar10].
10. **Pheromone** (size: 42 places, 48 transitions, 119 arcs) is a signal transduction network of the well understood mating pheromone response pathway in *Saccharomyces cerevisiae*. The qualitative Petri net in [SHK06] extends a former ODE model [KK04]. The Petri net was validated by Petri net invariants and a partitioning of the transition set.
11. **Potato** (size: 17 places, 25 transitions, 78 arcs) describes the main carbon metabolism, the sucrose-to-starch breakdown in *Solanum tuberosum* (potato) tubers. The qualitative Petri net model was developed in cooperation with experimentally working scientists, experienced in ODE modelling. Its validation by Petri net invariants is discussed in [HK04], and a more detailed pathway exploration in [KJH05].

**Table 1.** Some biomolecular case studies; all of them hold the DTP, are consistent and live.

#	case study	multiplicities	net class	bounded	liveness shown by
1	apoptosis	ORD	ES	no	Proposition 2.1
2	RKIP	ORD	ES	yes	Proposition 2.3
3	biosensor	ORD	ES	no	Proposition 2.3
4	hypoxia	ORD	not ES	no	structural reduction
5	lac operon	HOM	not ES	no	structural reduction
6	G/PPP	HOM	not ES	no	structural reduction
7	MAPK	ORD	not ES	yes	dynamic analysis (RG)
8	CC	HOM	not ES	yes	dynamic analysis (RG)
9	halo	not HOM	not ES	yes	dynamic analysis (RG)
10	pheromone	HOM	not ES	no	by reasoning
11	potato	not HOM	not ES	no	by reasoning

Contrary, the model of signal transduction events involved in the angiogenesis processes, which is discussed in [NMC<sup>+</sup>09] as a stochastic and continuous Petri net model (size: 39 places, 64 transitions, 185 arcs) is to a large extent covered by a (non-minimal) bad siphon. Thus, even if the net is live for a certain marking  $\mathbf{m}$ , there is always a larger marking  $\mathbf{m}'$ , which will allow to remove all tokens from the bad siphon. Consequently, an arbitrary marking increase will not preserve liveness.

## 7 Tools

The Petri nets for the case studies have been constructed using Snoopy [RMH10], a tool to design and animate or simulate hierarchical graphs, among them qualitative, stochastic and continuous Petri nets as used in the case studies in Section 6. Snoopy provides export to various analysis tools as well as import and export of the Systems Biology Markup Language (SBML).

The qualitative analyses have been made with the Petri net analysis tool Charlie [Fra09], complemented by the structural reduction rules supported by the Integrated Net Analyser INA [SR99].

## 8 Conclusions

We have discussed the problem of monotonic liveness, with one of the motivations originating from bio-model engineering. We have presented a new result showing the necessity of the DTP for monotonic liveness.

Moreover, we immediately know – thanks to the well-known propositions of the DTP – that ordinary ES nets are monotonically iff the DTP holds. Furthermore, we know – because the DTP monotonically ensures deadlock freeness – that for any net class, in which liveness and deadlock freeness coincide, monotonic liveness is characterized by the DTP. We have shown one instance for this case: the mono-T-semiflow nets (MTS).

We have demonstrated the usefulness of our results by applying them to a variety of biomolecular networks.

One of the remaining open issues is: what are sufficient conditions for monotonic liveness for more general net structures? While none of our test cases is an MTS net, this line might be worth being explored more carefully, e.g. by looking at FRT nets (Freely Related T-Semiflows) [CS92] and extensions.

**Acknowledgements.** This work has been partially supported by CICYT - FEDER grant DPI2006-15390 and by the European Community's Seventh Framework Programme under project DISC (Grant Agreement n. INFSO-ICT-224498). The work of M. Heiner was supported in part by the grant DPI2006-15390 for a stay with the Group of Discrete Event Systems Engineering (GISED); the main ideas of this paper have been conceived during this period.

## References

- Ber86. G. Berthelot. Checking properties of nets using transformations. In G. Rozenberg, editor, *Advances in Petri Nets 1985*, volume 222 of *Lecture Notes in Computer Science*, pages 19–40. Springer, 1986.
- BL00. N. Barkai and S. Leibler. Biological rhythms: Circadian clocks limited by noise. *Nature*, 403(6767):267–268, 2000.
- BRA83. G. W. BRAMS. *Reseaux de Petri. Theorie et pratique (2 tomes)*. Masson, 1983.

- BTS02. J.M. Berg, J.L. Tymoczko, and L. Stryer. *Biochemistry, 5th ed.* WH Freeman and Company, New York, 2002.
- CCS91. J. Campos, G. Chiola, and M. Silva. Ergodicity and Throughput Bounds of Petri Nets with Unique Consistent Firing Count Vector. *IEEE Transactions on Software Engineering*, 17:117–125, 1991.
- CS92. J. Campos and M. Silva. Structural techniques and performance bounds of stochastic Petri net models. In *Advances in Petri Nets 1992, volume 609 of Lecture Notes in Computer Science*, pages 352–391. Springer-Verlag, 1992.
- CSK<sup>+</sup>03. K.-H. Cho, S.-Y. Shin, H.-W. Kim, O. Wolkenhauer, B. McFerran, and W. Kolch. Mathematical modeling of the influence of RKIP on the ERK signaling pathway. In *Proc. CMSB*, pages 127–141. Springer, LNCS 2602, 2003.
- DA10. R. David and H. Alla. *Discrete, Continuous, and Hybrid Petri Nets*. Springer, 2010.
- DHP<sup>+</sup>93. F. DiCesare, G. Harhalakis, J.M. Proth, M. Silva, and F.B. Vernadat. *Practice of Petri nets in manufacturing*. Chapman & Hall, 1993.
- Fra09. A. Franzke. *Charlie 2.0 - a multi-threaded Petri net analyzer*. Diploma Thesis, Brandenburg University of Technology at Cottbus, CS Dep., 2009.
- GH06. D. Gilbert and M. Heiner. From Petri nets to differential equations - an integrative approach for biochemical network analysis. In *Proc. ICATPN 2006*, pages 181–200. LNCS 4024, Springer, 2006.
- GHL07. D. Gilbert, M. Heiner, and S. Lehrack. A unifying framework for modelling and analysing biochemical pathways using Petri nets. In *Proc. CMSB*, pages 200–216. LNCS/LNBI 4695, Springer, 2007.
- GHR<sup>+</sup>08. D. Gilbert, M. Heiner, S. Rosser, R. Fulton, Xu Gu, and M. Trybilo. A Case Study in Model-driven Synthetic Biology. In *Proc. 2nd IFIP Conference on Biologically Inspired Collaborative Computing (BICC), IFIP WCC 2008, Milano*, pages 163–175, 2008.
- HDG10. M. Heiner, R. Donaldson, and D. Gilbert. *Petri Nets for Systems Biology, in Iyengar, M.S. (ed.), Symbolic Systems Biology: Theory and Methods*. Jones and Bartlett Publishers, Inc., in Press, 2010.
- HGD08. M. Heiner, D. Gilbert, and R. Donaldson. Petri nets in systems and synthetic biology. In *Schools on Formal Methods (SFM)*, pages 215–264. LNCS 5016, Springer, 2008.
- HK04. M. Heiner and I. Koch. Petri Net Based Model Validation in Systems Biology. In *Proc. ICATPN, LNCS 3099*, pages 216–237. Springer, 2004.
- HKW04. M. Heiner, I. Koch, and J. Will. Model Validation of Biological Pathways Using Petri Nets - Demonstrated for Apoptosis. *BioSystems*, 75:15–28, 2004.
- HLGM09. M. Heiner, S. Lehrack, D. Gilbert, and W. Marwan. Extended Stochastic Petri Nets for Model-Based Design of Wetlab Experiments. *Transactions on Computational Systems Biology XI*, pages 138–163, 2009.
- HS10. M. Heiner and K. Sriram. Structural analysis to determine the core of hypoxia response network. *PLoS ONE*, 5(1):e8600, 01 2010.
- KH08. I. Koch and M. Heiner. *Petri Nets, in B.H. Junker and F. Schreiber (eds.), Biological Network Analysis*, chapter 7, pages 139–179. Wiley Book Series on Bioinformatics, 2008.
- KJH05. I. Koch, B.H. Junker, and M. Heiner. Application of Petri Net Theory for Modeling and Validation of the Sucrose Breakdown Pathway in the Potato Tuber. *Bioinformatics*, 21(7):1219–1226, 2005.

- KK04. B. Kofahl and E. Klipp. Modelling the dynamics of the yeast pheromone pathway. *Yeast*, 21(10):831–850, 2004.
- LBS00. A. Levchenko, J. Bruck, and P.W. Sternberg. Scaffold proteins may biphasically affect the levels of mitogen-activated protein kinase signaling and reduce its threshold properties. *Proc Natl Acad Sci USA*, 97(11):5818–5823, 2000.
- Mar10. W. Marwan. Phototaxis in halo bacterium as a case study for extended stochastic Petri nets. *Private Communication*, 2010.
- Mur89. T. Murata. Petri Nets: Properties, Analysis and Applications. *Proc. of the IEEE* 77, 4:541–580, 1989.
- NMC<sup>+</sup>09. L Napione, D Manini, F Cordero, A Horvath, A Picco, M De 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*, pages 281–295. Springer, LNCS/LNBI 5688, 2009.
- NMOG03. T. Nutsch, W. Marwan, D. Oesterhelt, and E.D. Gilles. Signal processing and flagellar motor switching during phototaxis of *Halobacterium salinarum*. *Genome research*, 13(11):2406–2412, 2003.
- Red94. V. N. Reddy. Modeling Biological Pathways: A Discrete Event Systems Approach. Master thesis, University of Maryland, 1994.
- RMH10. C. Rohr, W. Marwan, and M. Heiner. Snoopy - a unifying Petri net framework to investigate biomolecular networks. *Bioinformatics*, 26(7):974–975, 2010.
- RTS98. L. Recalde, E. Teruel, and M. Silva. On linear algebraic techniques for liveness analysis of P/T systems. *Journal of Circuits, Systems, and Computers*, 8(1):223–265, 1998.
- RTS99. L. Recalde, E. Teruel, and M. Silva. Autonomous continuous P/T systems. In *Proc. ICATPN*, pages 107–126. Springer, LNCS 1689, 1999.
- SH09. M. Schwarick and M. Heiner. CSL model checking of biochemical networks with Interval Decision Diagrams. In *Proc. CMSB*, pages 296–312. Springer, LNCS/LNBI 5688, 2009.
- SHK06. A. Sackmann, M. Heiner, and I. Koch. Application of Petri Net Based Analysis Techniques to Signal Transduction Pathways. *BMC Bioinformatics*, 7:482, 2006.
- Sil85. M. Silva. *Las Redes de Petri: en la Automática y la Informática*. Editorial AC, 1985.
- SR99. P.H. Starke and S. Roch. *INA - The Intergrated Net Analyzer*. Humboldt University Berlin, [www.informatik.hu-berlin.de/~starke/ina.html](http://www.informatik.hu-berlin.de/~starke/ina.html), 1999.
- SR02. M. Silva and L. Recalde. Petri nets and integrality relaxations: A view of continuous Petri net models. *IEEE Transactions on Systems, Man, and Cybernetics, Part C: Applications and Reviews*, 32(4):314–327, 2002.
- Sta90. P.H. Starke. *Analysis of Petri Net Models (in German)*. B. G. Teubner, Stuttgart, Stuttgart, 1990.
- TS96. E. Teruel and M. Silva. Structure theory of equal conflict systems. *Theoretical Computer Science*, 153:271–300, 1996.
- Wil06. D.J. Wilkinson. *Stochastic Modelling for System Biology*. CRC Press, New York, 1st Edition, 2006.
- YWS<sup>+</sup>07. Y. Yu, G. Wang, R. Simha, W. Peng, F. Turano, and C. Zeng. Pathway switching explains the sharp response characteristic of hypoxia response network. *PLoS Comput Biol*, 3(8):e171, 2007.