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## Convergence of Feller semigroups with applications to some stochastic genetic models

We study convergence of semigroups related to a singular-singularly perturbed abstract Cauchy problem (compare [1] and [2]), generalizing a number of recent models of mathematical biology, including the models of gene expression [2, 3, 5] and gene regulation [4]. Particular attention is paid to irregular convergence of these semigroups, i.e. convergence outside of hydrodynamic or regular space, where convergence follows by the Trotter-Kato theorem.

Given  $v, w \in \mathbb{R}^M$ ,  $M \in \mathbb{N}$  we define a compact set  $J = \{x \in \mathbb{R}^M : v \leq x \leq w\}$ . For fixed  $N \in \mathbb{N}$  we consider a stochastic process  $\{X(t), t \geq 0\}$ , which heuristically can be described as follows.  $X(t)$  jumps between  $N + 1$  copies of  $J$  according to a Markov chain-type mechanism. Between jumps, the process moves along the integral curves of ODEs, different on each copy of  $J$ . We investigate asymptotic behavior of  $X(t)$  when jump intensities are large.

At the time  $t$ , let  $\mathbf{x}(t)$  denote the position of  $X(t)$  in  $J$  and let  $\gamma(t)$  indicate which copy of  $J$  the process moves on.  $X(t) = (\mathbf{x}(t), \gamma(t))$  is an example of a piecewise deterministic Markov process of M.H.A. Davis. Consider a sequence  $(X_n(t))_{n \geq 0}$  of such processes. Their conditional expected values are given by  $\mathbf{f}_n(x, t) := \mathbb{E}_{x_n, \gamma_n} \mathbf{f}_n(\mathbf{x}_n(t), \gamma_n(t))$ , where  $(x_n, \gamma_n) := (x_n(0), \gamma_n(0))$ . If  $\mathbf{f}_n(x, t)$  are smooth enough (e.g. are of class  $C^1$ ), they satisfy the Cauchy problems

$$\frac{\partial \mathbf{f}_n(x, t)}{\partial t} = \mathcal{A}_0 \mathbf{f}_n(x, t) + \kappa_n \mathcal{Q}_n \mathbf{f}_n(x, t), \quad \mathbf{f}_n(x, 0) = \theta_n(x), \quad n \in \mathbb{N}, \quad (1)$$

where for fixed  $n, t$ ,  $\mathbf{f}_n$  belongs to a Cartesian product  $\mathbb{B}$  of  $N + 1$  copies of  $C = C(J)$ , the space of real-valued continuous functions on the set  $J$ , equipped with the supremum norm. The operator  $\mathcal{A}_0$  with domain  $\mathcal{D}$  is an infinitesimal generator of a  $c_0$  semigroup of contractions, describing deterministic movement of the processes along integral curves of ODEs.  $\mathcal{Q}_n$  is a sequence of bounded multiplication operators in  $\mathbb{B}$ , whose entries are continuous functions on  $J$ . For  $x \in J$ , each  $\mathcal{Q}_n(x)$  is the intensity matrix of a Markov chain, governing jumps of  $X_n(t)$ .  $\kappa_n$  is a sequence of non-negative constants such that  $\kappa_n \rightarrow \infty$  for  $n \rightarrow \infty$ , describing intensity of jumps. We prove a theorem about convergence of semigroups related to (1) for  $n \rightarrow \infty$ , assuming that  $\mathcal{Q}_n$  tend in operator norm to a limit operator  $\mathcal{Q}$ . Assuming that  $\mathcal{Q}(x)$  has the stationary distribution  $\mathbf{p}_0(x)$  and that  $\mathbf{p}_0$  are Lipschitz continuous functions of  $x$ , we prove that the solutions of (1) tend to these

of

$$\frac{\partial f(x, t)}{\partial t} = \mathbf{p}_0^\top \mathcal{A}_0 f(x, t), \quad f(x, 0) = \mathbf{p}_0^\top \theta(x), \quad f \in C^1. \quad (2)$$

This result can be applied to derivation of deterministic approximations of the stochastic Kepler-Elston model of gene regulation ([4]), describing binding of regulatory proteins to regulatory sequence in the gene. Similar deterministic approximation of a stochastic mechanism is used in the Lipniacki model of gene expression ([2, 3, 5]), where random activation or inactivation of a gene stimulates production of mRNA and proteins.

#### References

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