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Structural stability and structural sensitivity are mathematical concepts that describe the (qualitative or quantitative) dependence of dynamics on the parameters and functional forms underlying a dynamical system. *Identifiability* and estimability are statistical concepts that describe the possibility, or ease, of estimating the parameters of a specified model. The mathematical and statistical concepts are closely related; if the dynamics of a model are extremely sensitive to a given parameter (or functional form), then a statistical analysis of observed dynamics should be able to estimate the parameter very precisely. Conversely, when the dynamics of a model are completely insensitive to a parameter, that parameter will be statistically unidentifiable; observing the dynamics gives no information about the parameter.

We briefly discussed existing methods for analyzing structural stability, parametric sensitivity, and statistical identifiability. These comprise bifurcation analysis in mathematics; sensitivity analysis in statistics (e.g. quantification of sensitivity and elasticity of model outcomes to parameter values; global sensitivity methods such as FAST (McRae et al., 1982) or Latin hypercube sampling (Blower et al., 1991)); and identifiability analysis in statistics (e.g. data cloning (Lele et al., 2007; Ponciano et al., 2009; Lele et al., 2010), Wald estimates of variance-covariance matrices via the Fisher information matrix, and algebraic methods (Meshkat et al., 2009; Eisenberg et al., 2013)).

Our main interest was in exploring the duality of dynamical sensitivity and statistical estimability. The consequences of sensitivity depend on the goal and "direction" of analysis (forward vs. inverse): for example, sensitivity is problematic when one is trying to predict dynamics based on a specified model with known parameters, but convenient if one is trying to estimate parameters. Sensitivity and estimability also depend on which model inputs and outputs are chosen. On the input side, one can choose to focus on the sensitivity to a prespecified set of parameters, or reparameterize the model, or consider structural sensitivity by allowing variation in the functional forms within a model (Adamson and Morozov, 2013). On the output side, considering different response variables can qualitatively change the conclusions of a sensitivity analysis (Farcas and Rossberg, 2016; Li et al., 2017).

These fields are now moving from considering sensitivity and estimability of parameters to *structural sensitivity* (Adamson and Morozov, 2013) and non-parametric estimation of functional forms ("semimechanistic modeling" (Wood, 2001) or "model-free forecasting" (Perretti et al., 2013a,b)). While statistical sensitivity analysis usually focuses on quantitative changes in model outcomes, one can also consider qualitative changes (i.e., changes in the topology of the attractors (Adamson and Morozov, 2013)). While such qualitative outcomes may sometimes be of less interest in applications, at least the case of persistence vs. extinction forms the basis of population viability analysis. Some open questions in this area:

- How can we understand the relative sensitivity of different components of a system? For example, in a nonparametric or structural-sensitivity analysis, how can we pin down which properties of an input function (e.g. particular derivatives at particular points along the function) have the greatest effect on output?
- Can we characterize and understand in a *practical*, applications-oriented way how frequent structural sensitivity is, and when it is most likely to occur (Munch et al., 2018)? Are typical ecological models naturally sensitive? Are they self-tuned/self-organized to sensitivity? When is the sensitivity biologically relevant (e.g. "dangerous" sensitivity sensu "dangerous bifurcation")?
- Can we distinguish sensitivity in a model from sensitivity in a biological system? That is, what kinds of experiments and examples could (or do) reveal sensitivity in the biological system, without mediation through a (almost certainly misspecified) model? There are a many examples showing bifurcations, structural (in)stability or sensitivity in empirical systems: (Veilleux, 1979; Fussmann et al., 2000; Cushing et al., 2001; Melbourne and Hastings, 2009).

References

- Adamson, M. W. and Morozov, A. Y. (2013). When can we trust our model predictions? Unearthing structural sensitivity in biological systems. *Proceedings of the Royal Society A: Mathematical, Physical and Engineering Sciences*, 469(2149):20120500.
- Blower, S. M., Hartel, D., Dowlatabadi, H., Anderson, R. M., and May, R. M. (1991). Drugs, Sex and HIV: A Mathematical Model for New York City. *Philosophical Transactions: Biological Sciences*, 331(1260):171–187.
- Cushing, J. M., Henson, S. M., Desharnais, R. A., Dennis, B., Costantino, R. F., and King, A. (2001). A chaotic attractor in ecology: theory and experimental data. *Chaos, Solitons & Fractals*, 12(2):219–234.
- Eisenberg, M. C., Robertson, S. L., and Tien, J. H. (2013). Identifiability and estimation of multiple transmission pathways in cholera and waterborne disease. *Journal of theoretical biology*, 324:84–102.
- Farcas, A. and Rossberg, A. G. (2016). Maximum sustainable yield from interacting fish stocks in an uncertain world: two policy choices and underlying trade-offs. *ICES Journal of Marine Science*, 73(10):2499–2508.
- Fussmann, G., Ellner, S. P., Shertzer, K. W., and N. G. Hairston, J. (2000). Crossing the Hopf bifurcation in a live predator-prey system. *Science*, 290:1358–1360.

- Lele, S. R., Dennis, B., and Lutscher, F. (2007). Data cloning: easy maximum likelihood estimation for complex ecological models using Bayesian Markov chain Monte Carlo methods. *Ecology Letters*, 10:551–563.
- Lele, S. R., Nadeem, K., and Schmuland, B. (2010). Estimability and Likelihood Inference for Generalized Linear Mixed Models Using Data Cloning. *Journal* of the American Statistical Association, 105(492):1617–1625.
- Li, S.-L., Bjørnstad, O. N., Ferrari, M. J., Mummah, R., Runge, M. C., Fonnesbeck, C. J., Tildesley, M. J., Probert, W. J. M., and Shea, K. (2017). Essential information: Uncertainty and optimal control of Ebola outbreaks. Proceedings of the National Academy of Sciences, 114(22):5659–5664.
- McRae, G. J., Tilden, J. W., and Seinfeld, J. H. (1982). Global sensitivity analysis—a computational implementation of the Fourier Amplitude Sensitivity Test (FAST). Computers & Chemical Engineering, 6(1):15–25.
- Melbourne, B. A. and Hastings, A. (2009). Highly Variable Spread Rates in Replicated Biological Invasions: Fundamental Limits to Predictability. *Science*, 325(5947):1536–1539.
- Meshkat, N., Eisenberg, M., and DiStefano III, J. J. (2009). An algorithm for finding globally identifiable parameter combinations of nonlinear ODE models using Gröbner Bases. *Mathematical biosciences*, 222(2):61–72.
- Munch, S. B., Giron-Nava, A., and Sugihara, G. (2018). Nonlinear dynamics and noise in fisheries recruitment: A global meta-analysis. *Fish and Fisheries*, 19(6):964–973.
- Perretti, C. T., Munch, S. B., and Sugihara, G. (2013a). Model-free forecasting outperforms the correct mechanistic model for simulated and experimental data. *Proceedings of the National Academy of Sciences*, 110(13):5253–5257.
- Perretti, C. T., Sugihara, G., and Munch, S. B. (2013b). Nonparametric fore-casting outperforms parametric methods for a simulated multispecies system. *Ecology*, 94(4):794–800.
- Ponciano, J. M., Taper, M. L., Dennis, B., and Lele, S. R. (2009). Hierarchical models in ecology: confidence intervals, hypothesis testing, and model selection using data cloning. *Ecology*, 90(2):356–362.
- Veilleux, B. G. (1979). An Analysis of the Predatory Interaction Between Paramecium and Didinium. *Journal of Animal Ecology*, 48(3):787–803.
- Wood, S. N. (2001). Partially specified ecological models. *Ecological Monographs*, 7(1):1–25.