

# Parameter estimation of ODE's via nonparametric estimators

Nicolas Brunel, *Université d'Evry*

## Abstract

Ordinary differential equations (ODE's) are widespread models in physics, chemistry and biology. In particular, this mathematical formalism is used for describing the evolution of complex systems and it might consist of high-dimensional sets of coupled nonlinear differential equations. In this setting, we propose a general method for estimating the parameters indexing ODE's from times series. Our method is able to alleviate the computational difficulties encountered by the classical parametric methods. These difficulties are due to the implicit definition of the model. We propose the use of a nonparametric estimator of regression functions as a first-step in the construction of an M-estimator, and we show the consistency of the derived estimator under general conditions. In the case of spline estimators, we prove asymptotic normality, and that the rate of convergence is the usual  $\sqrt{n}$ -rate for parametric estimators. Some perspectives of refinements of this new family of parametric estimators are given.

AMS 2000 subject classifications: Primary 62F99.

Keywords: Asymptotics, M-estimator, Nonparametric regression, Ordinary Differential Equation, Parametric estimation, Splines.



Full Text: [PDF](#)

Brunel, Nicolas, Parameter estimation of ODE's via nonparametric estimators, Electronic Journal of Statistics, 2, (2008), 1242-1267 (electronic). DOI: 10.1214/07-EJS132.

## References

- [1] D. K. Andrews. Asymptotic normality of series estimators for nonparametric and semiparametric regression models. *Econometrica*, 59(2):307–345, 1991. [MR1097531](#)
- [2] R. Beran. Minimum hellinger distance estimates for parametric models. *Annals of Statistics*, 5:445–463, 1977. [MR0448700](#)
- [3] A. Berlinet and C. Thomas-Agnan. Reproducing Kernel Hilbert Spaces in Probability and Statistics. Kluwer, 2004. [MR2239907](#)
- [4] P.J. Bickel and Y. Ritov. Nonparametric estimators which can be “plugged-in”. *Annals of Statistics*, 31(4):4, 2003. [MR2001641](#)
- [5] N. Brunel. Parameter estimation of ode's via nonparametric estimator. Technical report, EURANDOM, 2007.
- [6] T. Cai. On adaptive wavelet estimation of a derivative and other related linear inverse problems. *Journal of Statistical Planning and Inference*, 108:329–349, 2002. [MR1947406](#)
- [7] C. de Boor. A practical guide to splines, volume 27. Springer-Verlag, 1979. [MR1900298](#)
- [8] I. Dimatteo, C.R. Genovese, and R.E. Kass. Bayesian curve-fitting with free-knot

[9] S. Donnet and A. Samson. Estimation of parameters in incomplete data models defined by dynamical systems. *Journal of Statistical Planning and Inference*, 137(9):2815–2831, 2006. [MR2323793](#)

[10] S. Efromovich. Nonparametric curve estimation – Methods, Theory and Applications. Springer-Verlag, New-York, 1999. [MR1705298](#)

[11] S. Ellner, Y. Seifu, and R.H. Smith. Fitting population dynamic models to time-series data by gradient matching. *Ecology*, 83:2256–2270, 2002.

[12] C.P. Fall, E.S. Marland, J.M. Wagner, and J.J. Tyson, editors. Computational Cell Biology. Interdisciplinary applied mathematics. Springer, 2002. [MR1911592](#)

[13] J. Fan and I. Gijbels. Local Polynomial Modelling And Its Applications. Chapman and Hall, London, 1996. [MR1383587](#)

[14] J. Fan and Y.K. Truong. Nonparametric regression with errors in variable. *Annals of Statistics*, 21(4):1900–1925, 1993. [MR1245773](#)

[15] J. Friedman. Multivariate adaptive regression splines. *Annals of Statistics*, 19:1–141, 1991. [MR1091842](#)

[16] S. Froda and G. Colavita. Estimating predator-prey systems via ordinary differential equations with closed orbits. *Australian & New-Zealand Journal of Statistics*, 47(2):235–254, 2005. [MR2155124](#)

[17] T. Gasser and H.G. Müller. Estimating regression functions and their derivatives by the kernel method. *Scandinavian J. Statistics*, 11:171–184, 1984. [MR0767241](#)

[18] L. Goldstein and K. Messer. Optimal plug-in estimators for nonparametric functional estimation. *Annals of Statistics*, 20:1306–1328, 1992. [MR1186251](#)

[19] M.W. Hirsch, S. Smale, and R. Devaney. Differential equation, Dynamical Systems and an Introduction to Chaos, volume 60 of Pure and Applied Mathematical series. Elsevier Academic Press, 2nde edition edition, 2003. [MR2144536](#)

[20] G. Hooker and L. Biegler. Ipopt and neural dynamics: Tips, tricks and diagnostics. Technical report, Department of Biological Statistics and Computational Biology, Cornell University, 2007.

[21] J.Z. Huang. Asymptotics for polynomial spline regression under weak conditions. *Statistics and probability letters*, 65:207–216, 2003. [MR2018032](#)

[22] C. Jost and S. Ellner. Testing for predator dependence in predator-prey dynamics: a non-parametric approach. *Proceedings of the Royal Society: Biological Sciences*, 267:1611–1620, 2000.

[23] J-Y Koo and K-W Lee. B-splines estimation of regression functions with errors in variable. *Statistics and probability letters*, 40:57–66, 1998. [MR1650520](#)

[24] H.L. Koul and P. Ni. Minimum distance regression model checking. *Journal of Statistical Planning and Inference*, 119:109–141, 2004. [MR2018453](#)

[25] Y.A. Kuznetsov. Elements of Applied Bifurcation Theory. Springer-Verlag, New York, 2004. [MR2071006](#)

[26] N. Lalam and C. Klaassen. Pseudo-maximum likelihood estimation for differential equations. Technical Report 2006-18, Eurandom, 2006.

[27] T. Lee. On algorithms for ordinary least squares regression spline fitting: a comparative study. *Journal of Statistical Computation and Simulation*, 72:647–663, 2002. [MR1930486](#)

- [28] Z. Li, M.R. Osborne, and T. Prvan. Parameter estimation of ordinary differential equations. *IMA Journal of Numerical Analysis*, 25:264–285, 2005. [MR2126204](#)
- [29] J. Madar, J. Abonyi, H. Roubos, and F. Szeifert. Incorporating prior knowledge in cubic spline approximation - application to the identification of reaction kinetic models. *Industrial and Engineering Chemistry Research*, 42(17):4043–4049, 2003.
- [30] M. Pascual and S. Ellner. Linking ecological patterns to environmental forcing via nonlinear time series models. *Ecology*, 81:2767–2780, 2000.
- [31] A. A. Poyton, M.S. Varziri, K.B. McAuley, P.J. McLellan, and J.O. Ramsay. Parameter estimation in continuous-time dynamic models using principal differential analysis. *Computers and Chemical Engineering*, 30:698–708, 2006.
- [32] J.O. Ramsay. Principal differential analysis: Data reduction by differential operators. *Journal of the Royal Statistical Society. Series B*, 58:495–508, 1996. [MR1394362](#)
- [33] J.O. Ramsay, G. Hooker, J. Cao, and D. Campbell. Parameter estimation for differential equations: A generalized smoothing approach. *Journal of the Royal Statistical Society (B)*, 69:741–796, 2007. (with discussions). [MR2368570](#)
- [34] J.O. Ramsay and B. W. Silverman. *Functional Data Analysis*. Springer series in statistics. Springer, 1997. [MR2168993](#)
- [35] C.J. Stone. Optimal rates of convergence for nonparametric estimators. *Annals of Statistics*, 8:1348–1360, 1980. [MR0594650](#)
- [36] C.J. Stone. Optimal global rates of convergence for nonparametric regression. *Annals of Statistics*, 10(4):1040–1053, 1982. [MR0673642](#)
- [37] C.J. Stone and J.Z. Huang. Free knot splines in concave extended linear modeling. *Journal of Statistical Planning and Inference*, 108:219–253, 2002. [MR1947401](#)
- [38] S. Vajda, P. Valko, and A. Yermakova. A direct-indirect procedure for estimation of kinetic parameters. *Computers and Chemical Engineering*, 10:49–58, 1986.
- [39] A.W. van der Vaart. *Asymptotic Statistics*. Cambridge Series in Statistical and Probabilities Mathematics. Cambridge University Press, 1998. [MR1652247](#)
- [40] J. M. Varah. A spline least squares method for numerical parameter estimation in differential equations. *SIAM J.sci. Stat. Comput.*, 3(1):28–46, 1982. [MR0651865](#)
- [41] M.S. Varziri, A.A. Poyton, K.B. McAuley, P.J. McLellan, and J.O. Ramsay. Selecting optimal weighting factors in ipda for parameter estimation in continuous-time dynamic models. *Computers and Chemical Engineering*, 32:3011–3022, 2008.
- [42] E.O. Voit and J. Almeida. Decoupling dynamical systems for pathway identification from metabolic profiles. *Bioinformatics*, 20(11):1670–1681, 2004.
- [43] G. Wahba. *Spline models for observational data*, volume 59. SIAM, 1990. [MR1045442](#)
- [44] S. Zhou, X. Shen, and D.A. Wolfe. Local asymptotics for regression splines and confidence regions. *Annals of Statistics*, 26(5):1760–1782, 1998. [MR1673277](#)
- [45] S. Zhou and D.A. Wolfe. On derivative estimation in spline regression. *Statistica Sinica*, 10:93–108, 2000. [MR1742102](#)