Neural Trajectories Reconstruction Matlab Toolbox. v1.0 Beta

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This is a Beta version under testing, not guaranteed. Nevertheless, support of this "toolbox" for researchers is freely provided by the authors under the terms of the GNU GPL license http://www.gnu.org/licenses/gpl-faq.html#WhatDoesGPLStandFor (within reason).

This "toolbox" is based on: Balaguer-Ballester E, Lapish C, Seamans JK and Durstewitz D 2011 "Attracting Dynamics of Frontal Cortex Populations during Memory Guided Decision-Making". PLoS Comput. Biol., *In press.*

Requirements: Matlab v7.11 (2010b), Statistics Toolbox v7.4 (2010b). It can be easily adapted to work on earlier matlab versions (please contact authors if any difficulty).

The code runs either in Windows 7 or in Linux, but please remembers to change the path in the first code lines of "ntr.m" file.

For running a short demonstration of this toolbox please type ">ntr;"

*NOTE: This brief document only provides an overview. Nevertheless, the code is profusely commented. For a detailed help, please move to the deploy directory and type:

">help filename", where "filename" is one of the files located in the main deployment directory.

Please, feel free to contact the authors for any question (kindly see upper email addresses).

I. Overview

This toolbox performs statistical analyses on nonlinear time-series of multivariate neural responses –of different kinds- during a cognitive task. Therefore it is assumed that the separate cognitive epoch's tasks are observable (e.g. a stimulus presentation, a successful choice, reward acquisition, movement to a definite place like an arm) and are labelled accordingly. This requirement will be relaxed in future versions.

It is also assumed that all multivariate neural responses are time-ordered and simultaneously recorded. Convergence or divergence of neural responses trajectories will be evaluated in those states defined by the distinct time-windows corresponding to cognitively relevant epochs of the task. Statistical analyses will indicate the probability of those states to "behave" like attracting regions of neural responses (or rather they will suggest more likely transient dynamics). This toolbox is intended to be used in different types of neurophysiological response. Nevertheless it was developed and testes only on electrophysiological data (not on BOLD or EEG responses at least in the present moment).

An optimal reconstruction of neural trajectories properties may require high-dim, nonlinear neural interactions spaces. For that reason, kernel methods will be used in the statistical analyses of such state spaces.

A demonstrative dataset is provided, please type ">ntr;". Besides in Balaguer et al. (2011), a brief overview of such techniques can be found in Durstewitz and Balaguer (2010), Statistical Approaches for Reconstructing Neuro-Cognitive Dynamics from High-Dimensional Neural Recordings. Neuroforum 4/10: 266-276 and in references therein (e.g. Chuckland et al., 1999 etc.). Automatic "data-smoothing" methods (e.g. for estimate firing rates from spike trains) are not provided within this version. Please see e.g. Yu et al. (2009) for this means.

II. How-to

II.1) Drop a ".mat" or ASCII-type file (e.g. named 'file_name.mat') on "./data" folder, where "./" indicates the directory containing this entire toolbox or "deployment directory" (Note: use ".\" for Linux). This 'file_name.mat' should contain a matlab matrix named "data" of dimension "number of time bins x (dimensionality of neural responses+3)", where "dimensionality of neural responses" refers to the number of simultaneously recorded neurons, voxels, electrodes etc. and/or delayed version of them. The dataset structure is the following:

- Columns "1:end-3" of "Data" matrix: Must contain neural responses over time.
- Column "end-2" of "Data" matrix: Must contain natural numbers>0, labelling the different stimulus or behavioural "epochs" in which the experimentalist segments the task. "-1" should encode "no-labelled" time-bins.

- Column "end-1" of "Data" matrix: Must contain natural numbers>0, they are alternative labelling used only for trial trajectory display (number of labels has to be smaller than 8. See section code for more info). Those labels typically represent "phases" of the task, containing different "epochs". If all "epochs" are to be displayed, this column has to be a copy of "end-2" one.
- Last column of "Data" matrix: Must contain natural numbers>0, labelling the different trials of the task. Trials typically represents repetitions of the experiment, containing each the same "phases" of the task.
- II.2) Open "kspaces_configuration.m" and setup the configuration parameters. Please find a detailed description of the parameters by typing ">help kspaces_config".
- II.3) Once in the deployed directory, type ">ntr('file_name');". Alternatively, one can load in the workspace the data matrix variable formatted as indicated in I.1 (e.g. "Data_matrix_name") and type ">ntr(Data_matrix_name);".

III. Outputs

Output is controlled by parameters specified in "kspaces_config.m" file. They are subdivided (for clarity purposes) into "Basic" and "Advanced" parameters. In this brief note, only few parameters are described. Please type ">help kspaces_config" and read "setup_config.m" function (and on each file of this toolbox) for more information.

III.1) Configuration of the algorithm is achieved by the next parameters:

- "order": Indicates the maximum order of the polynomial products among units' responses (please see "multinom_kernel.m" for more information). This product order is the parameter defining each high-dimensional space.
- "regularization": Ensures the validity of the analyses in future dataset by
 "penalizing" the complexity of the Fisher Discriminant Criterion in highdimensional spaces. It has to be tuned by validation analyses for each particular
 kind of data (please see "kfd_multiv.m"; "class_trajec.m" and "kfd_cross_val.m"
 for more information).
- "is_shuffled_events": If positive, shuffles blocks as specified by the labels in "end-2" data column (task-epoch labels). Please see "shuff_data.m" for more information. This serves for creating non-parametric block-permutation bootstrap analyses in which short-term autocorrelation is typically preserves.
- "is_shuffled_within_events": If positive, preserves blocks as specified by the labels in "end-2" data column (task-epoch labels), but shuffles all time-bins within such task-epochs (please see "shuff_data.m" for more information). This serves

- for creating non-parametric bootstrap analyses in which temporal contingency is destroyed.
- "do_dcm": If positive, performs a delay-coordinate map for disambiguating trajectories i.e. dimensions constructed from delayed versions of units' responses will be added to the original multivariate recordings space. An unwanted trajectory crossing occurs when all neural responses dimensions corresponding to two or more district task-epochs have the same values in the same time bin. Delay-coordinate maps may typically avoid those "non-deterministic" ambiguities. Other advanced parameters control how this map will be performed (see "dcm.m" for more information).

III.2) Low-dimensional displays:

- Single-trial trajectory display. Only shown if the configuration field "trial_disp>0" or "make_video>0". Displays a single-trial trajectory for orders 1 and the selected one for visual comparison (left and right plots respectively), coloured according to the task-phases selected in the Column "end-1" of "Data" matrix (see II.1). This plot is obtained using *kernelized* principal components analyses (K-PCA, Schölkopf et al, 1998) of a delay-coordinate map. The display includes only one trial, the one selected in the configuration field "trial_disp" (see "kspaces_config.m"). Alternatively, if the configuration field "make_video>0", a video of such trial will be played instead. More information by typing ">help visualization" (see also "visualization.m")
- Multiple trials flow display. Only shown if the configuration field "trials_flow_disp>0". Displays flow field for orders 1 and the selected one for visual comparison (left and right plots respectively), coloured according to the task-epochs selected in column "end-2" of "Data" matrix (see II). Those were plots into the three main first discriminant axes (or two if there are less than three task-epochs), obtained by multivariate *kernel*-fisher discriminant analyses (K-FDA, Mika et al., 2000). The display includes several consecutive trials simultaneously, the lower and upper and ones appear in the 1x2 matrix "trials_flow_disp" (see "kspaces_configuration.m").
- III.3) Command window statistical reports, showing out-of-sample predictions (cross-validation) in high-dimensional spaces (defined by the multiple activity interactions; see previous paragraphs) of task-epochs vs. neural states associations. Two types of statistics can be found: classical (comparing percentage of miss-classified vectors for each task-epoch and across different validation sets, see below) and a trajectory-based analysis (comparing percentage of divergent trajectories for each behavioral epoch).
 - Multiple-discriminant analyses. A Two-class FDA i.e. a maximum discriminating solution corresponding will be used for trajectory analyses. In addition, if

- "is_multiple_discriminant>0", an optimal discriminating subspace of dimensions = number of task-epochs-1 will be generated i.e. all task-epochs will be simultaneously analysed (default).
- Full statistics report. Default displays do not show statistical tests, intermediate statistical reports and warnings related to numerical problems in the analyses. However, iIf "full_stats_disp>0", information-theoretic measures, parametric and non-parametric normality tests of different kind and all warnings and intermediate reports will be displayed (please see "class_trajec.m" and "kfd_multiv.m" for more details). Warning: command window may become too "crowded", not recommended as default approach.
- Cross-validation. Two kinds of out-of-sample predictions are supported. If "causal_xval<0", standard leave-one-out method will be performed: Each ith-trial is removed in turn, then optimum discriminant directions are computed using all-but-the ith trial; while this trial will become the ith-validation set. However, if "causal_xval>0", causality-preserved n-fold cross-validation will be performed instead: last j-trials are removed in each j-th validation block. The non-removed trials form the reference set (which thus is smaller for increasing j-th-validation blocks). There will be n=(m/2)-1 validation blocks, always occurring "after" the reference set.