Dendritica

Version 1.0

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1. General Introduction

Dendritica is a program package for relating dendritic geometry and signal propagation. The programs are based on those used for the simulations described in the following paper:

<u>Vetter, P., Roth, A. & Häusser, M. (2001).</u> Action potential propagation in dendrites depends on dendritic morphology. *Journal of Neurophysiology*, 85: 926-937.

Dendritica can functionally be divided into three main parts:

- interactive morphological analysis and electrophysiological simulation of single cells
- automated batch simulations across a set of morphologies using the same simulation parameters
- automated analysis of batch simulation runs

Dendritica requires NEURON 4.1.1 with some modifications described in Appendix 1. It was tested for NEURON 4.1.1 on Linux and SGI IRIX. Some modifications to the *Dendritica* code may be necessary in order to run it on older or newer versions of NEURON.

We are very grateful to Muki Rapp, Diana Smetters, Nelson Spruston, Greg Stuart, and the contributors to the Duke-Southampton Neuronal Morphology Archive (accessible via http://www.neuro.soton.ac.uk) for allowing us to use their neuronal reconstructions for this project. We also thank Alain Destexhe (Destexhe@iaf.cnrs-gif.fr) and Zach Mainen (mainen@cshl.org) for providing NEURON code. Please note that while the programs in Dendritica are freely available, they are protected by the GNU public licence, and we request that you acknowledge us if you use the programs for a publication. If you require further information, please do not hesitate to contact any of the authors: Philipp Vetter (p.vetter@ucl.ac.uk), Arnd Roth (roth@mpimf-heidelberg.mpg.de), or Michael H usser (m.hausser@ucl.ac.uk).

1.1 Directory structure

The directory structure is defined automatically when you untar the package. The structure must be respected for most of the functions to work properly.

The directory dendritica-1.0 contains all files. There are three subdirectories:

```
batch_back/ batch_forward/ batch_forward2/
```

indicating different types of simulation runs, i.e. looking a backpropagating action potentials and forward propagating action potentials. The structure of the subdirectories is identical, however.

```
batch back/back:
      aphalf.hoc
                                 gui.hoc
     batch1.hoc
                                 help.hoc
     batch2.hoc
                                 impedance.hoc
     batch3.hoc
                                init.hoc
     batchrun
                                mod/
                               neuronprefs.hoc
    bp
      enuspike_p21 output.hoc electrophysiology.hoc parse.hoc
      dendspike p21
                                referenceAP p18@200um act0
      figures.hoc
      forward.hoc
                                settings.hoc
      geometry.hoc
                                statistics.hoc
      graphics.hoc
```

The *.hoc files contain the code for the NEURON interpreter. It is split into several files according to what the procedures/functions do. The directory mod/ contains all mod files necessary to create the special executable of NEURON. The mod files for the simulations are in the subdirectory kvz_naz.dendspike_p21 and referenceAP_p18@200um_act0 are saved waveforms that can be played in during simulations.

```
batch_back/data:
    act0/ cells/ geometry/
```

act0 contains simulation results that depend on the active model used, while geometry contains simulations results that are independent. cells contains the morphologies of all the cells used.

```
batch_back/neuron_output:
```

This contains ascii files with correlation analysis from a batch run.

2. Getting Started

Most of the data presented in the paper are directly accessible through the graphical user interface (GUI). (Note that the optimization routine to find halfdecay max is an exception).

2.1 A sample session

- To create the program special go to dendritica-1.0/batch_back/mod/kvz_naz
- To create special type >nrnivmod1
- Move special to dendritica-1.0/batch_back/, then load the GUI with
 >special gui.hoc -
 - Dialog box: Welcome to Progagation Geometry [Load Cell] [Statistics]
- Choose [Load Cell]
 - Dialog box: Please pick a neuron and an active model [Neuron] [Conductances]
- Specify Neuron [Nigra] -> [Nigra2], and Conductance [standard conductances] (act0 is the setting used in the paper), and then [Accept]. The Morphology is loaded, and subsequently the panels Electrophysiology and Main (see below) are called.

[Voltage Clamp] runs a Voltage clamp simulation with the Electrode standardly located at the soma. This takes about a minute on a PentiumII, depending also on the morphology being simulated. The position can be changed manually by clicking on [Location]. The size of the waveform can be changed by entering a value under [scaling]. The standard waveform is somatic AP (p18), but others can be chosen from the pull-down menu [waveform].

[Current Clamp] clamps a constant current, standardly at the soma. The electrode location can be changed manually using [Location]. The magnitude of the current can be specified by changing [Amplitude].

[Synapse] simulates a synapse. The parameters and location of the synapse can be altered by clicking on [Location] and/or [gmax].

[Input Resistance] calculates the input resistance at the soma.

[g_na threshold] calculates the Na-channel density for full backpropagation. This simulation can take >30 minutes!

- Press [Voltage Clamp]. After the simulation is over, 4 figures are plotted:
- (1) Voltage traces at soma, node and dendrite
- (2) The AP amplitude as a function of distance from the soma
- (3) Simulation settings
- (4) Plot of rate of change of peak voltage as a function of distance from the soma

 The same plots are obtained when simulating [Current Clamp] or [Synapse].

2.2 Options available from the GUI

```
[Load Neuron]
[Clear Screen]
[Graphs]
```

```
->[New Graph]
   ->[Which Sections]
   ->[Which Lengths]
   ->[Simulation]
   ->[Geometric Values]
   ->[Functional Values]
   ->[Simulation X against distance]
   ->[Geometric individual]
   ->[Impedance individual]
[Other Panels]
   ->[Electrophysiology]
   ->[Statistics]
   ->[Channels]
   ->[Simulation settings]
   ->[Geometry]
[Miscellaneous]
   ->..
[Quit]
```

- Choose [Graphs] -> [Which Sections] -> [all] to plot figures (2) & (4) with data from all segments of the Morphology.
- Choose [Graphs] -> [Which Lengths] -> [electrotonic] to plot figures (2) & (4) in electrotonic space (X-axis!).
- Choose [Other Panels] -> [Simulation settings] to call up a panel allowing to change simulation duration, and time step.
- Choose [Other Panels] -> [Conductances] to call up a panel which allows setting of the active membrane properties.
- To re-run the simulation, simply press [Voltage Clamp] in the panel Electrophysiology. Units as in the paper.
- Choose [Other Panels] -> [Geometry] to call up a panel which allows the Axon to be removed [Remove Axon] or added to the morphology [Connect Axon].
- [Graphs] -> [Geometric Values] Plots 5 figures that have been calculated in a batch simulation (see next section)
- (1) Branchpoint and Termination histogram as a function of distance from the soma
- (2) Cumulative membrane area as a function of distance from the soma
- (3) Rate of change in membrane against distance from the soma
- (4) Rall ratio distribution of branchpoints (smoothed)
- (5) number of sections at a given distance from the soma.

2.3 On-line help

An on-line help can be accessed from the command line. All functions and procedures of the package can be listed with the command

```
oc>hlp()
parse.hoc
get()
get_somadist()
connect_axon()
...
```

The listing is sorted according to the .hoc files the functions and procedures are defined in. To get more information about a particular function, e.g. the function get, type

```
oc>hlp("get")
get cell $s1
use ActiveModel $s2
load data if numarg=3
```

2.4 Basic commands for running simulations

There are four basic commands to run simulations from the command line (see hlp() for details)

- get() loads a morphology, it s simulation results, gets it ready for simulations
- sim() runs a simulation
- fig() plots vectors
- spaceplot() dumps a spaceplot on disk

2.5 What happens when loading a cell

get() loads morphologies from ../data/cells/<name of cell>, specifies name and spine_density in neuronprefs.hoc (structure MyCell) inserts passive membrane properties and channels parse.hoc, then sets the parameters as specified in settings.hoc. To facilitate analysis, the morphology is split into soma and dendrites (note Purkinje cells have two types of dendrites), and SectionLists are specified in parse.hoc accordingly.

```
dist_switch()
if (n == 1) distlist = trunk
if (n == 2) distlist = all
if (n == 3) distlist = branchpoints
if (n == 4) distlist = terminations
if (n == 5) distlist = branchpt_noend
if (n == 6) distlist = all noend
```

Simulation results (calculated previously) are loaded from ../data/act0/ and ../data/geometry" into vectors. These vectors can be printed using pt(<vectorname>), or plotted using fig(<vectorname>). Vectors can be plotted against each other as fig(<vectorx>,<vectory>) (see hlp("fig")).

2.6 What happens during simulations

Most functions to do with simulations are in electrophysiology.hoc (see hlp() for details). Simulations come in three flavours - voltage clamp/current clamp/synapse, which is set by the flag simMode. sim() brings the cell to resting potential with rest(), then inserts the appropriate PointProcess. The unused PointProcesses are parked on a dummy section. sim then calls simcore() which is equivalent to run(). Because some values, like the AP half-width require knowledge of the AP-waveform, simcore() has to be called twice, so that these values can be calculated.

To be able to plot them, type

```
>sim_calc()
```

which creates the following vectors that can be plotted against distance from the soma.

- vpk peak voltage
- amp AP amplitude
- vmax maximum velocity

```
plat - peak latency
olat - onset latency
half - half distance
dvdr - spatial derivative of peak voltage
e.g.
>sim()
>dist_switch(2) // all sections [optional]
>L_switch(0) // physical lengths [optional]
>sim_calc()
>fig(dist,vpk)
```

3. Batch Simulations

Batch simulations allow for the automated generation and saving of simulation results across a wide range of morphologies using the same set of simulation parameters. Because these calculations are computationally intensive, it is more efficient to invoke batch() without using the GUI. Simulation runs take >24 hours on a PentiumII 450 MHz.

3.1 Examples

```
>batch(17,act0)
```

performs all the calculations in conjuction with action potential backpropagation, using the standard active model "act0".

```
>batch(18,act0)
```

performs all the calculations when the action potential is generated at a dendritic location 200 um from the soma.

```
>batch(19,act0)
```

performs all the calculations when the action potential is generated at the dendritic location from where the action potential has the greatest halfdecay distance.

3.2 How batch simulations are done

The procedure batch() is a loop which applies a function to all cells in turn. The specifics of this are defined in output.hoc Basically, calculations done in electrophysiology.hoc, geometry.hoc and impedance.hoc are saved as numbers or vectors in the directories ../data/act0 and ../data/geometry. The convention is that the directory name is the same as the vector, and the filename is the same as that of morphological data of the cell in ../data/cells.

4. Batch Analysis

4.1 A sample session

The results of the batch simulations can be analysed using the graphical user interface.

- Go to directory dendritica-1.0/batch_back/back/ and type
 >special gui.hoc Dialog box: Welcome to Progagation Geometry [Load Cell]
 [Statistics]
 Choose [Statistics]
 Dialog box: Select dataset to analyse
 [Conductances]
 [] equivalent
 - [] backpropagation
 - [] forward200
 - [] forwardhdecay
- Select [Conductances] -> [standard]
- Select [x] backpropagation
- Press [Load]

The simulation results are loaded into memory, and the panels Main and Statistics are opened

[Get_Data] [Legend]
[Average] [Single] [Double] [Triple] [] Powers
[Y]
[X1]
[X2]
[X3]

- Choose [Y] -> [1] -> [nathresholdvclamp]
- Press [Average]
 This does 3 things
- (1) plots a bar chart with cell-type averaged Na thresh values under voltage clamp
- (2) prints numerical values on command line
- (3) saves numerical values in ascii in

dendritica-1.0/batch back/neuron output/nathresholdvclamp

- Choose [X1]->[3]->[d2area max]
- Press [Single]
- Press [Legend] This correlates the maximum rate of rise in membrane area as a function of distance from the soma with nathresholdvclamp and shows a legend colour-coding the cell types. Again, 3 things are done
- (4) correlation plot
- (5) numerical values on command line
- (6) numerical values saved in

/neuron_output/nathresholdvclamp vs branchpoints_num (act0)

- Choose [] Powers
- Press [Single]

This does the same as before, but maximizes the correlation nathresholdvclamp and d2area_max^exponent, by varying the exponent.

• Choose [X2]->[3]->[diam mean]

• Press [Double]

This maximizes the correlation between nathresholdvclamp and (d2area_max^a * diam mean^b)

- Press [Clear Screen]
- Choose [X1] -> [geometric]
- Deselect (optional) Powers
- Press [Single]

This plots the 6 best correlations of geometric parameters against nathresholdvclamp, and plots a ranked list of correlations on the command line

- Press [Clear Screen]
- Type

```
>make figures()
```

This creates all the average and correlation plots shown in the paper.

Type

```
>multi correlation()
```

This will save all good single and multiple correlations into the file

dendritica-1.0/batch back/neuron output/backpropagation

4.2 Basic commands

There are six key commands for ANALYSIS/STATISTICS

- (1) get_data() loads simulation results for the whole batch of cells
- (2) averages () prints/plots cell-type average for any parameter
- (3) cplot() correlates two parameters with each other
- (4) single_corr() correlates one parameter with all geometric parameters
- (5) single corrf() correlates one parameter with all functional parameters
- (6) writevecs() writes vectors to disk

4.3 Settings

FLAGS that have to be set (use before calling get ()):

equiv 1= equivalent cylinder mode

hdecay 1= morphology is cut in two, where the halfdecay distance is maximal; distal

part removed

forward 1= morphology cut in two 200 um from soma, distal part removed

simMode 0= do voltage clamp when sim() is called

1= do current clamp when sim() is called 3= do synapse when sim() is called

electrotonicL 0= physical lengths

1= electrotonic lengths

(Note that usually, equiv=hdecay=forward=simMode=0)

4.4 What happens during correlation analysis

During correlation analysis, all simulation results are read from disk into the vector

```
data[i][j][k]
```

- $i = \{0,1,2\}$ and specifies, respectively, a functional, physically-geometric, electrotonically-geometric parameter
- $j = \{0..30\}$ for the different parameters
- $k = \{0,1,2\}$ 0 = parameter (normal), 1 = exp(parameter), 2 = ln(parameter)

It s a nuisance to specify one vector with three numbers, so there is a one-number shorthand

```
1000*i + 100*k + j {if i==0 add 3000}
```

dissect() turns shorthand into ci,cj,ck, antidissect() does the opposite. Because they are all vectors they can be plotted and manipulated as mentioned above.

To get averages of a parameter for a given cell-type

```
>averages(3014) // gets nathreshold (voltage clamp mode) averages
```

N.B. This writes the numerical values into a correctly named file

```
../neuron output/nathesholdvclamp
```

To make the correlations, the appropriate data[][][] vectors are copied into vecx and vecy, and Rcorrelation() is applied. To correlate two parameters

```
>cplot(3014,1000)
```

N.B. This writes the numerical values into a correctly named file in

```
../neuron output/nathresholdvclamp vs area max act0
```

N.B.II All such data relevant for the figures is generated automatically using make figures()

Many correlations are possible (just loop through the indices i,j,k)

and in order to make sense of the data

(single_corr()/single_corrf(), double_corr(). To make the data more easy to read, they are ranked according to their correlation coefficient in good_corr().

To look at a mix of these correlations (with and without powers | normal or equivalent cable geometries etc)

4.5 List of functional parameters

r = distance from soma

Δr = incremental distance

Parameter	#	Description	
st intensity		-	
	3001	Current needed to elicit a nodal AP in the absence of somatic/dendritic sodium channels	
Nathreshold 3		g_na that leads to a depolarization >0mV in all sections during current clamp at st_intensity	
Nathresholdvclamp	3014	<code>g_na</code> that leads to a depolarization >0mV in all sections during voltage clamp with AP waveform	
nathresholdvclamp2	3021	g na that leads to a depolarization $>0\mathrm{mV}$ in terminal sections during voltage clamp with AP waveform	
AP200	3010	AP amplitude 200 um from soma / AP amplitude at soma	
AP200_pass	3011	AP amplitude 200 um from soma / AP amplitude at soma (g_na = 0)	
AP200_half	3016	Sigmoidal fit of AP200 = f(g_na)	
		<pre>AP200 = AP200_basis + AP200_range/ { 1+exp[-(g_na - AP200_half)/AP200_steep)]}</pre>	
AP200_steep	3017	See above	
AP200_range	3018	See above	
AP200_basis	3019	See above	
Aphalf	3012	Distance from soma at which AP amplitude has decayed to 50%	
APhalf_pass	3013	Distance from soma at which AP amplitude has decayed to 50% (g_na=0)	
input_resistance	3015	Input resistance at soma	
Rfwd_min	3026	minimum somatofugal input resistance:	
		Cut morphology in half at a given point, and measure the input resistance at the end with the somatofugal portion of the morphology	
Rfwd max	3027	Maximum somatofugal input resistance	
_ Zfwd min	3022	minimum somatofugal input impedance (f=200 Hz)	
_ Zfwd max	3023	Maximum somatofugal input impedance (f=200 Hz)	
_ Rmismatch_peak	3002	Cut morphology in half at a given point	
-		Measure resting input resistance at both new ends.	
		Mismatch is defined as the ratio of somatopetal/somatofugal input resistance.	
		=> peak value of this mismatch	
Zmismatch peak	3003	Same as above, but measuring input impedance at 200 Hz	
aRmismatch_peak	3004	Same as Rmismatch peak, but measuring resistance at time, when the peak of the action potential has just reached the point of measurement.	
aZmismatch peak	3005	Analogous	
Rmismatch mean	3006	Same calculations as above, but take the mean over all points instead of peak.	
Zmismatch mean	3007	Analogous	
aRmismatch mean	3008	Analogous	
aZmismatch mean	3009	Analogous	
dZfwd max	3024	Maximum ΔZfwd/ Δr	
dZfwd min	3025	Minimum ΔZfwd/ Δr	
dRfwd max	3028	Maximum $\Delta Z fwd/\Delta r$	
dRfwd min	3029	Maximum ΔZfwd/ Δr	
aZfwd_min	3030	Same as Zfwd_min, but calculations done when action potential has just reached the point at which the cut is made.	
aZfwd max	3031	Analogous	
daZfwd max	3032	Analogous	
 daZfwd min	3033	Analogous	
aRfwd_min	3034	Analogous	
aRfwd max	3035	Analogous	
daRfwd max	3036	Analogous	
_ daRfwd min	3037	Analogous	
cZfwd min	3038	Minimum of ΔZfwd/ (Δr · Zfwd) over morphology	
cZfwd max	3039	Maximum of ΔZfwd/ (Δr · Zfwd) over morphology	
cRfwd min	3040	Minimum of Δ Rfwd/ (Δ r · Rfwd) over morphology	
cRfwd max	3041	Maximum of $\Delta Rfwd$ / ($\Delta r \cdot Rfwd$) over morphology	
caZfwd min	3042	Minimum of $\Delta Z f w d / (\Delta r \cdot Z f w d)$ over morphology when AP has just reached point	
caZfwd max	3043	Maximum of $\Delta Z f w d / (\Delta r \cdot Z f w d)$ over morphology when AP has just reached point	
caRfwd min	3044	Minimum of $\Delta Rfwd$ / ($\Delta r \cdot Rfwd$) over morphology when AP has just reached point	
caRfwd max	3045	Maximum of $\Delta Rwd/$ ($\Delta r \cdot Rfwd$) over morphology when AP has just reached point	
_	-	,	

4.6 List of geometric parameters

heranchpoints_num distance_max 1008 Maximum r area_max 1008 Mostimum to the professions area (spine corrected) taper_mean 1010 Mean taper_A(diameter)/ Ar in the somatofugal direction darea_max 1000 Maximum A(membrane area/) Ar darea maxdist 1001 r at which maximum A(membrane area/) Ar is reached ddr_pelmax 1002 First relative maximum in the change of membrane area after the first minimum change of membrane area as a twn of distance from soma; these values are calculated seeni-automatically dddr_ratio 1011 dddr_releax / preceding minimum change in membrane area as a fxn of distance d2area_max 1013 Meximum archese values are calculated seeni-automatically d2area_max 1014 Meximum archese values are calculated seeni-automatically d2area_maxist 1014 Meximum archese values are calculated seeni-automatically d2area_maxist 1015 Meximum change area at their darea maximum arcaches d2area_maxist 1016 At d2area_maxist: 1007 membrane area distal to soma/membrane area for the soma d2area_maxist percent 1015 At d2area_maxist: 1007 membrane area distal to soma/tectal membrane area d2area_maxist percent 1016 At d2area_maxist: 1007 membrane area distal to soma/tectal membrane area d2area_maxist percent 1015 At d2area_maxist: 1007 membrane area distal to soma/tectal membrane area d2area_maxist 1016 Meximum number of sections are as distal to soma/tectal membrane area realizatio_peak 1017 Meximum number of sections at a given distance from the soma sections_max sections_max sections_max 1018 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1019 Meximum number of sections at all r dam_ario_peak 1020 Meximum number of sections at all r dam_ario_peak 1021	PARAMETER	#	DESCRIPTION
distance max 1006 Maximum r reas max 1007 Total membrane area (spine corrected) taper_mean 1010 Maximum A(membrane area) / Ar in the somatofugal direction darca paxx 1010 Maximum A(membrane area) / Ar darea paxdist 1010 First relative maximum in the change of membrane area as a tran of distance from sona; these values are calculated semi-automatically dAdr_ratio 1011 Adr_relamx 1012 Adr_relamx / preceding minimum change in membrane area as a fix not distance from sona; these values are calculated semi-automatically dAdr_ratio 1011 Adr_relamx / preceding minimum change in membrane area as a fix not distance from darca maxdist 1019 Maximum rate of change in A(membrane area) As a function of distance from darca maxdir_ratio 1014 Distance from the sona at which d2area_max is reached d2area maxdir_precent 1015 At d2area_maxdist 100° membrane area distal to soma/tend membrane area raltratio_mean 1014 Adr_ream_maxdist 100° Area maxdist membrane area distal to soma/tend membrane area raltratio_mean 1015 At d2area_maxdist 100° membrane area distal to soma/tend membrane area raltratio_mean 1016 Mean of the distribution of Rall ratios obtained from the branchpoints in the morphology raltratio_mean 1019 Maximum number of sections at a given distance from the branchpoints in the morphology restriction_max sections_maxdist 1019 Feak in the distance from the sona sections_maxdist 1010 Feak in the distance between branchpoints branchdensityII noend 1021 Mean distance between branchpoints branchdensityII noend 1022 Number of branchpoints / total length of dendritic sections branchdensityII noend 1023 Number of branchpoints / total length of mon-terminal branchpoints Mean of the above distribution 1024 Same as a filantio_mean, but omitting terminal branchpoints 1025 Same as a fallratio_mean, but omitting terminal branchpoints 1026 Mean distance for branchpoints of the dendritic sections 1027 Same as rallratio_mean, but omitting terminal branchpoints 1028 Same as a fallratio_mean, but omitting terminal branchpoints 1029 Same	branchpoints num	1003	Number of branchpoints
area max 1002 Total membrane area (spine corrected) taper_mean 1010 Mean taper A(diameter)/ Ar in the somatofugal direction darea max 1000 Maximum A(membrane area)/ Ar is reached dAdr_prainx 1001 r at which maximum A(membrane area)/ Ar is reached dAdr_pratio 2012 Pirar relative maximum in the change of membrane area as a firm of distance from sona; these values are calculated semi-automatically dZarea_max 1011 Addr_cleamy / preceding minimum change in membrane area as a function of distance from the sona dZarea_maxit 1015 At dZarea_maxit 1016 At dZarea_maxidist; membrane area distal to soma/tention of distance from the sona dZarea_maxit 1015 At dZarea_maxidist; membrane area distal to soma/tention of distance from the sona dZarea_maxit 1015 At dZarea_maxidist; membrane area distal to soma/tention membrane area rallratio_mean 1016 At dZarea_maxidist; membrane area distal to soma/tenti membrane area rallratio_mean 1007 Aximm number of sections at all ratios obtained from the branchpoints in the membrane area rallratio_mean 1017 Maximm number of sections at all r rallratio_mean 1017 Mean inthic secti		1006	
Laper, mean 1010 Mean taper A(diameter) / Ar in the somatofugal direction darea max 100 Maximum A(membrane area) / Ar is reached dadr_ratio 101 First relative maximum in the change of membrane area after the first minimum change in membrane area after the first minimum change in membrane area as a fxm of distance from soma; these values are calculated semi-automatically dAdr_ratio 1013 Maximum rate of change in A(membrane area) / Ar as a function of distance from the soma at which diarca max is reached dAzera_maxAr_ratio 1015 At diarca_maxdist: 1004 membrane area distal to soma/membrane area proximal to soma distance area proximal to soma and the distribution of Rall ratios obtained from the branchpoints in the morphology rallratio_mean 1004 Mean of the distribution of Rall ratios obtained from the branchpoints in the morphology rallratio_mean 1005 Pack in the distribution of Rall ratios obtained from the branchpoints in the morphology rallratio_mean 1005 Pack in the distribution of Rall ratios obtained from the branchpoints in the morphology rallratio_mean 1007 Maximum number of sections at all r dam_mean 1007 Mean dedritic diameter branchdensityI 1018 Mean distance between branchpoints branchdensityII noend diamatio_need peak	_	1002	Total membrane area (spine corrected)
darea_maxdist 1000 waximum A(membrane area)/ Ar is reached dddr_relmax 1015 First relative maximum in the change of membrane area after the first minimum calculated semi-automatically calculated calculated semi-automatically calculated calculated semi-automatically calculated calculated semi-automatically calculated calculated s	_	1010	
dard_relmax Comparison		1000	
dAdr_relmax Size First relative maximum in the change of membrane area as a fur of distance from soma; these values are calculated semi-automatically dadr_ratio Size S	-	1001	
from somms; these values are calculated semi-automatically	_	1012	change of membrane area as a fxn of distance from soma; these values are
Carea maxdist 1014 Distance from the soma at which d2area_max is reached d2area_maxAr_ratio 1015 At d2area_maxdist: membrane area distal to soma/membrane area proximal to soma d2area_maxAr_percent 1016 At d2area_maxdist: 100*membrane area distal to soma/total membrane area rallratio_mean 1004 Mean of the distribution of Rall ratios obtained from the branchpoints in the morphology sections_max 1007 Maximum number of sections at a given distance from the soma sections maxdist 1008 r at which sections_max sections mean 1009 Mean number of sections at all r diam_mean 1017 Mean dendritic diameter branchdensityII noend 1018 Mean dendritic diameter branchdensityII noend 1028 Number of branchpoints / total length of dendritic sections diam_ratio_mean 1021 Mean of the above distribution diam_ratio_mean 1021 Mean of the above distribution diam_ratio_mean 1025 Same as diam_ratio_mean but leaving out terminal branchpoints diam_ratio_mean 1025 Same as diam_ratio_mean but leaving out terminal branchpoints diam_ratio_mean 1025 Same as diam_ratio_mean but leaving out terminal branchpoints diam_ratio_mean 1025 Same as at allratio_mean, but make the proper sections mean_stem_dendrite_diam 1026 Mean diameter of dendrites branching off from soma rallratio_noend_mean 1025 Same as at allratio_mean, but make the reminal branchpoints deq_ratio 2016 Equivalent to dAdr_relmax in the equivalent cable representation deq_max 2016 Equivalent to dAdr_relmax in the equivalent cable representation ded_maxAr_ratio 2018 Equivalent to d2_area_maxAr_ratio in the equivalent cable representation ded_maxAr_ratio 2018 Equivalent to d2_area_maxAr_ratio in the equivalent cable representation ded_maxAr_ratio 2018 Equivalent to d2_area_maxAr_ratio in the equivalent cable representation ded_maxAr_ratio 2018 Equivalent to d2_area_maxAr_ratio in the equivalent cable representation ded_maxAr_ratio 2016 Equivalent to d2_area_maxAr_ratio in the equivalent cable representation ded_maxAr_ratio 2016 Equivalent to d2_area_maxAr_ratio in the equivalent cable repre	dAdr_ratio	1011	
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branchdensityII_noend diamratio_peak local peak to the distribution of diamter ratios at branchpoints given by	branchdensity	1018	Mean distance between branchpoints
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adiam_mean 2011 diam_mean in electrotonic space abranchdensity 2007 branchdensity in electrotonic space abranchdensityII 2008 branchdensityII in electrotonic space abranchdensityII_noend 2010 branchdensityII_noend in electrotonic space adeq_max 2012 deq_max in electrotonic space	asections_mean	2005	sections_mean in electrotonic space
abranchdensity 2007 branchdensity in electrotonic space abranchdensityII 2008 branchdensityII in electrotonic space abranchdensityII_noend 2010 branchdensityII_noend in electrotonic space adeq_max 2012 deq_max in electrotonic space	ataper_mean	2006	taper_mean in electrotonic space
abranchdensityII 2008 branchdensityII in electrotonic space abranchdensityII_noend 2010 branchdensityII_noend in electrotonic space adeq_max 2012 deq_max in electrotonic space	adiam_mean	2011	diam_mean in electrotonic space
abranchdensityII_noend 2010 branchdensityII_noend in electrotonic space adeq_max 2012 deq_max in electrotonic space	abranchdensity	2007	branchdensity in electrotonic space
adeq_max 2012 deq_max in electrotonic space	abranchdensityII	2008	branchdensityII in electrotonic space
	abranchdensityII_noend	2010	branchdensityII_noend in electrotonic space
adog maydigt 2012 dog maydigt in elegtrotopic grago	adeq_max	2012	deq_max in electrotonic space
aueq_maxursc 2013 ueq_maxursc in electroconic space	adeq_maxdist	2013	deq_maxdist in electrotonic space

5. Appendix 1: Modifications to NEURON 4.1.1

To run all parts of *Dendritica* successfully, the following modifications to the NEURON source code are required.

```
diff -r nrn.new/src/ivoc/vector.c nrn.old/src/ivoc/vector.c
< // #define BYTEHEADER int BYTESWAP_FLAG=0;</pre>
< // #define BYTESWAP(_X__,_TYPE__)
   #define BYTEHEADER int BYTESWAP_FLAG=0;
   #define BYTESWAP( X , TYPE )
68,69c68,69
< #if 1
< // #include <sys/isa_defs.h>
   #if 0
• #include <sys/isa defs.h>
Only in nrn.new/src/ivoc: vector.c.byteswap
Only in nrn.new/src/ivoc: vector.c.orig
diff -r nrn.new/src/nrnoc/cabcode.c nrn.old/src/nrnoc/cabcode.c
48,49c48
< #define NSECSTACK 10000
< /* A.R. 28.12.1998 */
   #define NSECSTACK 20
Only in nrn.new/src/nrnoc: cabcode.c.orig
diff -r nrn.new/src/oc/hoc_oop.c nrn.old/src/oc/hoc_oop.c
184,185c184
< #define NTYPESTACK 10000
< /* A.R. 28.12.1998 */
   #define NTYPESTACK 30
218,219c217
< #define NTEMPLATESTACK
                          10000
< /* A.R.
                 28.12.1998 */
   #define NTEMPLATESTACK
Only in nrn.new/src/oc: hoc_oop.c.orig
```

NEURON must be recompiled for the changes to take effect.

6. Appendix 2: List of functions

parse.hoc	cosinefit()
get()	
<pre>get_somadist()</pre>	forward.hoc
connect_axon()	name_somadist()
add_axon()	\mathtt{name} _halfdecay()
remove_axon()	resize_cell()
insert_channels()	
${ t make_sectionlists()}$	output.hoc
isterminal()	batch()
${ t make_distvectors()}$	calculation()
switch()	manual()
${ t dist_switch()}$	<pre>save_geometry()</pre>
$ extbf{L}_{ extbf{switch}}()$	save_active()
${ t make_vectors()}$	save_cable()
single_vectors()	${ t save_forwardmini()}$
set_origin()	save_all()
	save_back()
help.hoc	save_fI()
hlp()	save_fII()
hlpscan()	helpme()
hlpfound()	write_numbers()
fxnscan()	write_nathreshold()
check()	geometry_read()
consistency()	active_read()
get_parents()	normforward()
find_section()	equivforward()
traces()	equivforwardII()
fxarea()	printvectors()
sectest()	printvectors_back()
which()	printvectors_forward()
alastmanhusialasu has	printvectors_forward2()
electrophysiology.hoc	make_figures()
rest()	equivZfwdwrite()
simcore()	
sim()	statistics.hoc
initsimvclamp()	<pre>get neurondata()</pre>
<pre>dvdr_calc() sim err()</pre>	add_vals()
sim_eff() sim fit()	lg()
rinput_calc()	get_data()
sim calc()	${\tt compare_activemodels()}$
threshold_calc()	c2()
forwardthreshold_calc()	c3 ()
threshold find()	c()
threshold()	dissect()
thresh()	antidissect()
APdecay()	clean()
APdecay_sensitivity()	correl_raw()
sigmoidal()	correl_func1()
sigmoidal calc()	correl_func2()
scrappy()	correl_func3()
	Rcorrelation()
impedance.hoc	single_corr()
impedance calc()	single_corrf()
impedance mismatch()	double_corr()
switch_off_intra()	triple_corr()
switch_on_intra()	clegend() averages()
get_children()	_
switch_on()	label_list() clabel()
$imp_calc()$	Clabel() Cplot()
<pre>impedance_check()</pre>	powerplot()
get_Zfwdvalues()	cplot()
get_cZ()	get geomorder()
<pre>get_APfrequencies()</pre>	multiplot()
cosine()	datalegend()
cosinefxn()	aacaregena()

```
good_corr_func()
good_doublecorr()
checkit()
multi_correlation()
write_singlecorr()
neuronprefs.hoc
add_cell()
cell name()
set_suffix()
set_spinedensity()
dendII()
dendIII()
swc format()
make_sectionrefs()
geometry.hoc
fdistance()
fL()
segL()
mindist()
maxdist()
farea()
sectionarea()
fseg()
fbranch()
get_parent()
pbranchpoint()
nextparent()
branchpoint()
get root()
ubranch()
get rall()
rall_calc()
ename()
gstep()
make_dAr()
get_gdist()
geometry_calc()
mean()
div()
equivalent calc()
spinetransform()
make_equivalent_cable()
slope darea()
slope deq()
dAdr_calc()
dAdr_write()
deq_calc()
deq_write()
estcore()
tap()
lintaper()
get link()
set_electrotonic()
graphics.hoc
flip()
pt()
P()
ar()
mx()
mn()
mod()
ceil()
fig()
figlab()
clf()
```

hist()

gauss() bar() sort() filter() rolling() rolling2() roll() writevec el() writeveca() readveca() writevec() readvec() writevecs() nvectors() nasens() spaceplot() show()