2- Single Cell Hoc in BMTK

Instructions

This document assumes you have completed the necessary steps in 01-Installing_BMTK

Setup

1. Now that BTMK is installed, visit the directory you want to **build** your model in and run BMTK's directory setup. The network directory is where the node/edge configuration files will sit. For example, the following will change directory to your desktop, create a project directory (bmtk_model), create a network directory for btmk to use, and initialize the directory for your further customization. This step will only be run once!

```
cd C:\Users\<your_username>\Desktop
mkdir my_bmtk_model
cd my_bmtk_model
python -m bmtk.utils.sim setup bionet .
```

This will create the bmtk directory structure at the present location "\$BASE.dir", and create the following files and nested directories:

Nested directories: biophys_coponents, network, output Json files: circuit_config.json, simulation_config.json

2. In Windows you may be met with compilation errors like the following:

```
C:\Users\Tyler\Desktop\my_bmtk_model\biophys_components\mechanism
s
Was unable to compile mechanism in $COMPONENTS_DIR/mechanisms
```

This is normal. You will need to compile your mod files any time they change. Run mknrndll in the (\components\mechanisms\modfiles\) directory then copy the resulting dll file to the parent directory (\components\mechanisms\)

In Linux, copy all mod files into the (components\mechanisms) directory and run nrnivmodl

HOC Build

- 1. HOC template files <u>can</u> be used by BMTK. For this example, we'll take a single HCO cell and run it with no input, in BMTK.
- 2. Download the HCO files from https://github.com/tjbanks/two-cell-hco/archive/master.zip
- 3. Extract the files in the zip directory and copy all .mod files into (.\components\mechanisms\modfiles\)
- 4. Run nrnmkdll in
 (.\components\mechanisms\modfiles\)

Copy the resulting dll file from that directory into
(.\components\mechanisms\)

5. Next, tell BMTK where to find your template files. In the root of your model directory, edit circuit_config.json. Edit the templates_dir key under components. Your file should look similar to the following (and may already exist):

```
circuit_config.json
1
 2
 3
      "manifest": {
        "$BASE DIR": ".",
 4
 5
        "$COMPONENTS DIR": "$BASE DIR/biophys components",
 6
        "$NETWORK DIR": "$BASE DIR/network"
 7
 8
      "components": {
        "morphologies dir": "$COMPONENTS DIR/morphologies",
 9
        "synaptic_models_dir": "$COMPONENTS_DIR/synaptic_models",
10
11
        "mechanisms dir": "$COMPONENTS DIR/mechanisms",
12
        "biophysical neuron models dir":
13
    "$COMPONENTS DIR/biophysical neuron templates",
14
        "point neuron models dir": "$COMPONENTS DIR/point neuron templates",
        "templates dir": "$COMPONENTS DIR/templates"
15
16
      },
17
      "networks": {
18
        "nodes": [],
19
        "edges": []
20
21
22
```

- 6. Create a templates directory under . \components if it doesn't already exist.
- 7. Create a new file named **HCOCell.hoc** in your 'templates' directory and paste the following code into that file:

```
HCOCell.hoc
1
2
   begintemplate HCOcell
3
       public NumSoma
 4
       NumSoma = 1
 5
           public soma
 6
                           // declares object soma is a public object that
7
   can be accessed by any procedures/functions
8
           create soma [NumSoma] // declares soma as a membrane
9
   compartment object
10
11
       public all, somatic, basal, apical, axonal
12
        objref all, somatic, basal, apical, axonal
13
14
        objref this
15
```

```
16
17
18
            proc init() {
19
20
            all = new SectionList()
21
            somatic = new SectionList()
22
            basal = new SectionList()
23
            apical = new SectionList()
24
             axonal = new SectionList()
25
26
27
            for i=0,NumSoma soma[i] { all.append()
28
                 somatic.append() }
29
30
                     ////// geometrical properties //////
31
                     soma[0] {
32
                            nseg=1 // create only one segment in the soma
33
                            // gives area of .628e-3 cm<sup>2</sup>
34
                            L=1000
                                            // (micrometer)
35
                            diam=9.99593
                                           // (micrometer)
36
                     }
37
38
                    ///// biophysical properties /////
39
                    soma[0] {
40
                             cm = 1 // (microF/cm2)
41
42
                            //insert the appropriate channels and give them
43
    reversal potentials and conductances
44
                             insert leak
45
                             insert cat
46
                             insert cas
47
                             insert kdr
48
                            insert ka
49
                            insert kca
50
                            insert capool
51
                            insert hyper
52
53
                            eleak = -50 // (mV)
54
                            gbar leak = .03e-3 // (siemens/cm2)
55
56
                            cao = 3
57
                            cai = 50e-6
                            gbar_cat = .02 //(.005\sim.01 siemens/cm2)
58
59
                            gbar_cas = .01 //(.001 \sim .005 siemens/cm2)
60
61
                            ek = -80
62
                            gbar kdr = .1 // (.1 \sim .5 siemens/cm2)
63
                            gbar^{ka} = .3 // (.1 \sim .5 siemens/cm2)
64
                            \frac{1}{2} gbar kca = .01 //// (.01~.05 siemens/cm2)
65
66
                            eh=-20
67
                            gbar hyper = .0002 // (.0001 \sim .0003 siemens/cm2)
68
69
                            insert na
70
                            ena = 50
                                                     // (mV)
71
                            gbar na = 0.5 // (siemens/cm2)
72
```

- 8. Things to note about this file:
 - a. There are public section references used by bmtk:
 - i. Line 11: public all, somatic, basal, apical, axonal
 - b. The sections are indexable, i.e., soma is not a single object but an array.
 - i. Line 8: create soma[NumSoma]
 - ii. Line 27: append your sections to the correct section list
 - c. This can be any hoc file that specifies a template. This will be where a good majority of your model customization will be.
 - d. Line 75: define_shape() must be called if you don't define a 3d morphology in the hoc otherwise. BMTK relies heavily on 3d locations.
- 9. Something <u>important to note</u>, which may be a limitation of BMTK: the morphology file is ALWAYS required. For any hoc file loaded you must specify an swc, however, it can be a blank file, as it will be ignored. Create the **blank.swc** file in

.\biophys components\morphologies and leave this file blank.

Later versions of BMTK may not require the blank.swc file.

```
blank.swc
```

10. Next, you'll need to create a cell builder script to tell BMTK the type of cells you want to use.

Create the file **build_network.py** in the root of your directory and add the following code:

```
build_network.py
   from bmtk.builder.networks import NetworkBuilder
2
 3
 4
   net = NetworkBuilder('hco net')
 5
   net.add nodes (cell name='HCOCell',
 6
                  model type='biophysical',
7
                  model template='hoc:HCOcell',
8
                  morphology='blank.swc',
                  HCOCell='HCOCell'
9
10
11
   net.build()
12
   net.save nodes(output dir='network')
13
```

11. Now you should be ready to build your network. In your Anaconda Prompt, in the root of your directory execute the following command to build your network:

```
python build network.py
```

A successful run may not have any output.

12. Before running your simulation you will need to tell BMTK which generated network files are to be used in your simulation. ANY time you change your network configuration (networks, edges, etc) this will need to be updated. These files were generated in the previous step and exist in the network directory. Edit circuit_config.json. Add the appropriate "networks" key values that correspond to the files generated in the network directory. Your file should look like the following:

```
circuit_config.json
1
2
 3
      "manifest": {
        "$BASE DIR": ".",
 4
        "$COMPONENTS DIR": "$BASE DIR/biophys components",
 5
 6
        "$NETWORK DIR": "$BASE DIR/network"
7
 8
      "components": {
        "morphologies dir": "$COMPONENTS DIR/morphologies",
 9
        "synaptic models dir": "$COMPONENTS DIR/synaptic models",
10
        "mechanisms dir": "$COMPONENTS DIR/mechanisms",
11
12
        "biophysical neuron models dir":
    "$COMPONENTS DIR/biophysical neuron templates",
13
        "point neuron models dir": "$COMPONENTS DIR/point neuron templates",
14
15
        "templates dir": "$COMPONENTS DIR/templates"
16
17
      "networks": {
18
        "nodes": [
19
            "nodes file": "$NETWORK DIR/hco net nodes.h5",
20
            "node types file": "$NETWORK DIR/hco net node types.csv"
21
22
        ],
        "edges": [
```

13. An important, step needed for the network to run correctly:

BMTK will need to have its default hoc loader overridden. You do this by editing your run bionet.py file to add the highlighted code:

```
run_bionet.py
   # -*- coding: utf-8 -*-
1
 2
 3
   """Simulates an example network of 14 cell receiving two kinds of
   exernal input as defined in configuration file"""
 4
 5
   import os, sys
 6
7
   from bmtk.simulator import bionet
   from bmtk.simulator.bionet.default setters.cell models import loadHOC
   bionet.pyfunction cache.add cell model(loadHOC, directive='hoc',
10
   model type='biophysical')
11
12
   def run(config file):
13
        conf = bionet.Config.from json(config file, validate=True)
14
15
        conf.build env()
16
```

14. Finally, run the simulation by executing the following in your Anaconda prompt, in the root of your project directory:

```
python run bionet.py simulation config.json
```

A successful run will output something like the following:

```
(clean) C:\Users\Tyler\Desktop\my bmtk model>python run bionet.py
simulation config.json
2018-12-30 20:23:31,719 [INFO] Created log file 2018-12-30 20:23:31,779 [INFO] Building cells.
C:\Users\Tyler\Anaconda3\envs\clean\lib\site-packages\bmtk-0.0.7-
py3.7.egg\bmtk\simulator\bionet\morphology.py:61: RuntimeWarning: invalid
value encountered in true divide
  r3dsoma /= n3dsoma
2018-12-30 20:23:31,796 [INFO] Building recurrent connections
2018-12-30 20:23:31,804 [INFO] Running simulation for 1000.000 ms with
the time step 0.001 ms
2018-12-30 20:23:31,804 [INFO] Starting timestep: 0 at t_sim: 0.000 ms
2018-12-30 20:23:31,810 [INFO] Block save every 5000 steps
2018-12-30 20:23:31,929 [INFO]
                                    step:5000 t sim:5.00 ms
2018-12-30 20:23:32,061 [INFO]
                                    step:10000 \overline{t} sim:10.00 ms
                                  step:15000 t_sim:15.00 ms
2018-12-30 20:23:32,165 [INFO]
2018-12-30 20:23:32,231 [INFO]
                                    step:20000 t sim:20.00 ms
2018-12-30 20:23:49,841 [INFO]
                                    step:985000 t sim:985.00 ms
2018-12-30 20:23:50,006 [INFO]
                                    step:990000 t sim:990.00 ms
2018-12-30 20:23:50,156 [INFO]
                                    step:995000 t sim:995.00 ms
2018-12-30 20:23:50,316 [INFO]
                                    step:1000000 t sim:1000.00 ms
2018-12-30 20:23:50,357 [INFO] Simulation completed in 18.55 seconds
```

- 15. If you receive "PermissionError: [WinError 5] Access is denied: './output'" just run the network again.
- 16. At this point your network should have ran. From here we can customize the output and view results
- 17. Edit the reports section of the **simulation_config.json** file in the root of your model directory to look like:

```
simulation_config.json
26
    "reports": {
27
           "membrane report": {
           "module": "membrane report",
28
           "cells": "all",
29
           "variable name": [
30
31
             "cai",
              '' <sub>77</sub> ''
32
33
           "file name": "cell vars.h5",
34
35
           "sections": "soma"
36
37
```

- 18. This will instruct BMTK to log the calcium and voltage levels for your cell.

 Run your network again, as we did previously, with (python run_bionet.py simulation config.json)
- 19. Create a new file named **plot_test.py** in the root of your directory and paste the following code into it:

```
plot_test.py

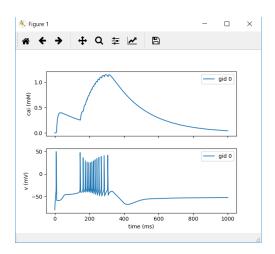
from bmtk.analyzer.cell_vars import plot_report

plot_report(config_file='simulation_config.json')

plot_report(config_file='simulation_config.json')
```

20. Executing this file with (python plot test.py) will return a plot like the following.

If you only see a single plot, that is ok, voltage is the important one here. Adding 'cai' to the simulation_config reports section is for demonstration purposes only.



For additional resources and instructions on configuring BMTK see: https://github.com/AllenInstitute/bmtk/blob/develop/docs/tutorial/Simulation_Intro.ipynb