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1 Transition to Oscillations in Two Coupled Variables

1.1 Mathematical Model

Single Variable, first-order differential equation

The equation:

$$\begin{split} \frac{dEx}{dt} &= h_{ex} - Ex + c_1 * tanh(Ex) - c_2 * tanh(In) \\ \frac{dIn}{dt} &= h_{in} - In + c_3 * tanh(Ex) - c_4 * tanh(In) \end{split}$$

where Ex and In are variables that changes with time t, h_{ex} and c_i are model parameters, tanh is the tangens hyperbolicus.

For $c_2 = c_3 = 0$, the two variables are uncoupled and independent.

The model dynamics can be characterised by scans of parameter h_ex. Here, we use a slow continuous change of this parameter to simulate spontaneous qualitative transitions of dynamics in the human EEG.

1.2 Import Functions

```
[62]: from scipy.integrate import odeint from scipy.signal import find_peaks, butter, sosfilt

from numpy import zeros, ones, tanh, mod, gradient, linspace, sign, log, meshgrid
from numpy import sqrt, fill_diagonal, ndarray, amax, amin, where from numpy import asarray, array, around, arange, corrcoef, flip, var from numpy.random import default_rng

from matplotlib.pyplot import subplots, xticks, yticks, axes
```

1.3 Model and Functions

```
[34]: def sigmoid(u):
          return tanh(u)
      def single_oscillator(y, t, h_ex, h_in, pars):
          tau_ex, tau_in, c_1, c_2, c_3, c_4 = pars
          dvdt = (
              (h_{ex} - y[0] + c_1*sigmoid(y[0]) - c_2*sigmoid(y[1]))*tau_ex,
              (h_{in} - y[1] + c_3*sigmoid(y[0]) - c_4*sigmoid(y[1]))*tau_in,
          return dydt
      def single_oscillator_plus_driving(y, t, h_ex, h_in, pars, sr, time_stop,_u
       ⇒driving, driving_strength):
          tau_ex, tau_in, c1, c2, c3, c4 = pars
          index = int(t*sr)
          if index >= time_stop*sr:
              dydt = zeros(2)
              return dydt
          h_ex_driven = h_ex + driving_strength*driving[index]
          dydt = (
              (h_ex_driven - y[0] + c1*sigmoid(y[0]) - c2*sigmoid(y[1]))*tau_ex,
                         - y[1] + c3*sigmoid(y[0]) - c4*sigmoid(y[1]))*tau_in,
         )
          return dydt
      def single_oscillator_plus_driving_plus_noise(y, t, h_ex, h_in, pars, sr,u
       stime_stop, driving, driving_strength, random_data, random_strength):
          tau_ex, tau_in, c1_ex, c2_ex, c3_in, c4_in = pars
          index = int(t*sr)
          if index >= time_stop*sr:
```

```
dydt = zeros(2)

return dydt

h_ex_driven = h_ex + driving_strength*driving[index]

dydt = (
    (h_ex_driven - y[0] + c1_ex*sigmoid(y[0]) - c2_ex*sigmoid(y[1]) +
    random_strength*random_data[index, 0])*tau_ex,
    (h_in - y[1] + c3_in*sigmoid(y[0]) - c4_in*sigmoid(y[1]) +
    random_strength*random_data[index, 1])*tau_in
)

return dydt
```

```
[122]: def plot_series(time, data, time_begin, time_end, sr):
           N = data.shape[1]//2
           name_vars = ('Ex', 'In')
           no_vars = 2*N
           fig, ax = subplots(ncols=2*N, figsize=(6, 4))
           for ind in arange(no_vars):
               ax[ind].plot(time[time_begin*sr:time_end*sr], data[time_begin*sr:

→time_end*sr, ind], linewidth=2, c='b')
               ax[ind].set_xticks(linspace(0, time_end-time_begin, 5));
               ax[ind].set_xticklabels(linspace(0, time_end-time_begin, 5));
               ax[ind].set_xlabel('Time', fontsize=12);
               ax[ind].set_ylabel(name_vars[ind], fontsize=12)
               y_min, y_max = ax[ind].get_ylim()
               ax[ind].set_yticks(linspace(y_min, y_max, 3));
               ax[ind].set_yticklabels(around(linspace(y_min, y_max, 3),1),__

¬fontsize=14);
           fig.tight_layout()
           return fig, ax
       def plot_series_statespace(time, data, time_begin, time_end, sr):
```

```
name_vars = ('Ex', 'In')
         no_vars = 2*N
         fig, ax = subplots(ncols=2*N, figsize=(6, 4))
         ax[0].plot(time[time_begin*sr:time_end*sr], data[time_begin*sr:time_end*sr,_
      \circlearrowleft0], linewidth=2, c='b')
         ax[0].set_xticks(linspace(0, time_end-time_begin, 5));
         ax[0].set_xticklabels(linspace(0, time_end-time_begin, 5));
         ax[0].set_xlabel('Time', fontsize=12);
         y_min, y_max = ax[0].get_ylim()
         ax[0].set_yticks(linspace(y_min, y_max, 3));
         ax[0].set_yticklabels(around(linspace(y_min, y_max, 3),1), fontsize=14);
         ax[0].set_ylabel(name_vars[0], fontsize=12);
         ax[1].plot(data[time_begin*sr:time_end*sr, 1], data[time_begin*sr:
      →time_end*sr, 0], linewidth=2, c='b')
         x_min, x_max = ax[1].get_xlim()
         ax[1].set_xticks(linspace(y_min, y_max, 3));
         ax[1].set_xticklabels(linspace(0, time_end-time_begin, 3));
         ax[1].set_xlabel(name_vars[1], fontsize=12);
         ax[1].set_ylabel(name_vars[0], fontsize=12)
         y min, y max = ax[1].get ylim()
         ax[1].set_yticks(linspace(y_min, y_max, 3));
         ax[1].set_yticklabels(around(linspace(y_min, y_max, 3),1), fontsize=14);
         ax[1].set_ylabel(name_vars[0], fontsize=12);
         fig.tight_layout()
         return fig, ax
[4]: def plot_bifdiagram(results_min_f, results_max_f,
                         results_min_b, results_max_b,
                         par_set):
         N = len(results_min_f)
         fig, ax = subplots()
         for xe, ye in zip(par_set, results_max_f[0]):
             if not isinstance(ye, ndarray):
                 ax.scatter(xe, ye, c='r', s=5)
```

N = data.shape[1]//2

```
else:
        ax.scatter([xe] * len(ye), ye, c='m', s=50, marker='x')
for xe, ye in zip(par_set, results_min_f[0]):
    if not isinstance(ye, ndarray):
        ax.scatter(xe, ye, c='r', s=5)
    else:
        ax.scatter([xe] * len(ye), ye, c='m', s=50, marker='x')
for xe, ye in zip(flip(par_set), results_max_b[0]):
    if not isinstance(ye, ndarray):
        ax.scatter(xe, ye, c='r', s=5)
    else:
        ax.scatter([xe] * len(ye), ye, c='b', s=20, marker='P')
for xe, ye in zip(flip(par_set), results_min_b[0]):
    if not isinstance(ye, ndarray):
        ax.scatter(xe, ye, c='r', s=5)
    else:
        ax.scatter([xe] * len(ye), ye, c='b', s=20, marker='P')
ax.set_xticks(linspace(par_min, par_max, 5));
ax.set_xticklabels(around(linspace(par_min, par_max, 5), 2), fontsize=16);
ax.set_xlabel('Parameter', fontsize=16)
ax.set_ylabel('Ex', fontsize=14)
y_min, y_max = ax.get_ylim()
ax.set_yticks(linspace(y_min, y_max, 3));
ax.set_yticklabels(around(linspace(y_min, y_max, 3),2), fontsize=14);
fig.tight_layout()
return fig, ax
```

1.4 Time Series

```
[135]: # Excitatory input parameter
h_ex_0 = -4.2
h_in_0 = -4
# Supercritical Hopf parameters
```

```
pars = (1, 1, 4, 6, 6, 0)
# Bistability parameters
# pars = (1, 1, 4, 1, 6, 0)
# Initial conditions
SEED = 123
rng = default_rng()
y_ini = rng.uniform(size=2)
\# y_ini = y[-1, :]
# Time array
time\_stop = 100
         = 1000
sr
          = linspace(start=0, stop=time_stop, num=time_stop*sr)
time
# Simulation
y = odeint(func=single_oscillator, y0=y_ini, t=time,
          args=(h_ex_0, h_in_0, pars),
          hmax=0.1)
# Show final values of all variables
print('End of run:', list(around(y[-1,:],3)))
print('')
```

End of run: [-2.076, -9.814]

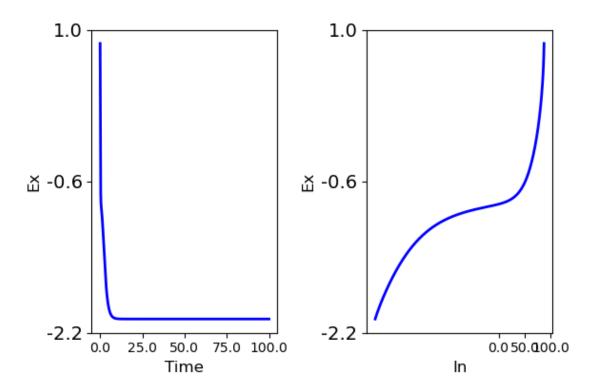
```
[130]: time_begin, time_end = 0, time_stop

fig, ax = plot_series_statespace(time, y, time_begin, time_end, sr)

title_chars = 'Figs/SNIC_Timeseries_h_ex' + str(h_ex_0) + '.png'

# fig.savefig(title_chars, format='png')
print(title_chars)
```

Figs/SNIC_Timeseries_h_ex-4.2.png



1.5 Bifurcation Diagram

```
[131]: # Initial conditions
y_ini = y[-1, :]

# Bifurcation parameter range
steps = 50

par_min, par_max = -5, 2

par_set = linspace(par_min, par_max, steps)

# Stop time
time_stop = 500
time = linspace(start=0, stop=time_stop, num=time_stop*sr)

results_max_f = dict()
results_max_inds_f = dict()
results_min_f = dict()
results_min_inds_f = dict()
rows = time.size
```

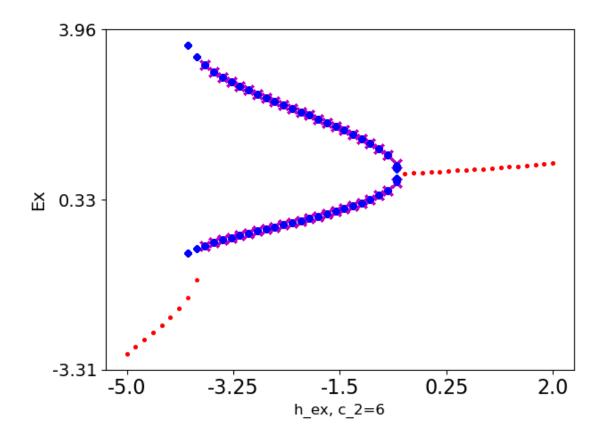
```
# Simulation "forward"
for par in par_set:
   h_ex = par
   y_f = odeint(func=single_oscillator, y0=y_ini, t=time,
             args=(h_ex, h_in_0, pars),
            hmax=0.1)
   for num, series in enumerate(y_f[rows//2:,:-1:2].T):
        if var(series) < 0.00005:</pre>
            if num not in results_max_f:
                results_max_f[num]
                                     = [series[-1]]
                results_max_inds_f[num] = [0]
                                     = [series[-1]]
                results_min_f[num]
                results_min_inds_f[num] = [0]
            else:
                results_max_f[num].append(series[-1])
                results_max_inds_f[num].append(0)
                results_min_f[num].append(series[-1])
                results_min_inds_f[num].append(0)
        else:
            y_f_max_inds = find_peaks(series, distance=100)
                      = series[y_f_max_inds[0]]
           y_f_maxs
           y_f_min_inds = find_peaks(-series, distance=100)
           y_f_mins = series[y_f_min_inds[0]]
            if num not in results_max_f:
                results max f[num]
                                     = [y_f_maxs]
                results_max_inds_f[num] = [y_f_max_inds]
                results min f[num]
                                   = [y f mins]
                results_min_inds_f[num] = [y_f_min_inds]
            else:
                results_max_f[num].append(y_f_maxs)
                results_max_inds_f[num].append(y_f_max_inds)
```

```
results_min_f[num].append(y_f_mins)
                results_min_inds_f[num].append(y_f_min_inds)
   if par != par_set[-1]:
       y_{ini} = y_{f}[-1, :]
results_max_b = dict()
results_max_inds_b = dict()
results_min_b = dict()
results_min_inds_b = dict()
# Simulation "backward"
for par in flip(par_set):
   h_ex = par
   y_b = odeint(func=single_oscillator, y0=y_ini, t=time,
             args=(h_ex, h_in_0, pars),
            hmax=0.1)
   for num, series in enumerate(y_b[rows//2:,:-1:2].T):
        if var(series) < 0.00005:</pre>
            if num not in results_max_b:
                results_max_b[num] = [series[-1]]
                results_max_inds_b[num] = [0]
                results_min_b[num]
                                     = [series[-1]]
                results_min_inds_b[num] = [0]
            else:
                results_max_b[num].append(series[-1])
                results_max_inds_b[num].append(0)
                results_min_b[num].append(series[-1])
                results_min_inds_b[num].append(0)
        else:
            y_b_max_inds = find_peaks(series, distance=100)
            y_b_maxs
                      = series[y_b_max_inds[0]]
            y_b_min_inds = find_peaks(-series, distance=100)
                     = series[y_b_min_inds[0]]
            y_b_mins
```

```
if num not in results_max_b:
                results_max_b[num]
                                     = [y_b_{maxs}]
                results_max_inds_b[num] = [y_b_max_inds]
                results_min_b[num]
                                        = [y_b_mins]
                results_min_inds_b[num] = [y_b_min_inds]
            else:
                results_max_b[num].append(y_b_maxs)
                results_max_inds_b[num].append(y_b_max_inds)
                results_min_b[num].append(y_b_mins)
                results_min_inds_b[num].append(y_b_min_inds)
    y_{ini} = y_{b}[-1, :]
print('')
print('Scan complete!')
print('')
```

Scan complete!

[132]: 'losc_Bifs_h_ex, c_2=6.png'



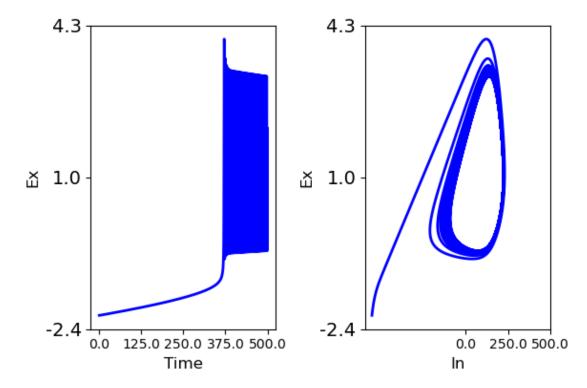
1.6 Transition to oscillation

```
print('End of run:', y.shape)
print('')
```

End of run: (500000, 2)

```
[137]: time_begin, time_end = 0, time_stop
fig, ax = plot_series_statespace(time, y, time_begin, time_end, sr)
if pars[1] == 1:
    title_chars = 'Figs/N=1/Onset_SNIC_h_ex.png'
elif pars[1] == 2:
    title_chars = 'Figs/N=1/Onset_Homoclinic_h_ex.png'
# fig.savefig(title_chars, dpi=300, format='png', bbox_inches='tight')
print(title_chars)
```

Figs/N=1/Onset_SNIC_h_ex.png



1.7 With added noise

```
[138]: # Set Initial conditions:
       \# y_i = y[-1, :]
       # Time array
       time_stop = 50
       time
                = linspace(start=0, stop=time_stop, num=time_stop*sr)
       # Initial parameter value
       h_ex_0
              = -4.2
       # Driving
       driving = linspace(0, 1, time.size)
       driving_strength = 1.0
       # Noise
       SEED = 123
       rng = default_rng(SEED)
       random_data = rng.normal(size=(time.size, y.shape[1]))
       order, band_low, band_high = 5, 1, 10
       sos = butter(order, (band_low, band_high), btype='bandpass', fs=sr,_
       →output='sos')
       random_data_filtered = zeros((time.size, y.shape[1]))
       for index, column in enumerate(random_data.transpose()):
           forward = sosfilt(sos, column)
           backwards = sosfilt(sos, forward[-1::-1])
           random_data_filtered[:, index] = backwards[-1::-1]
       random_strength = 10
       # Simulation /Add noise arrays to params
       y = odeint(func=single_oscillator_plus_driving_plus_noise, y0=y_ini, t=time,
                 args=(h_ex_0, h_in_0, pars, sr, time_stop, driving, driving_strength,
                  random_data_filtered, random_strength), hmax=0.1)
       print('End of run:')
       print('')
```

End of run:

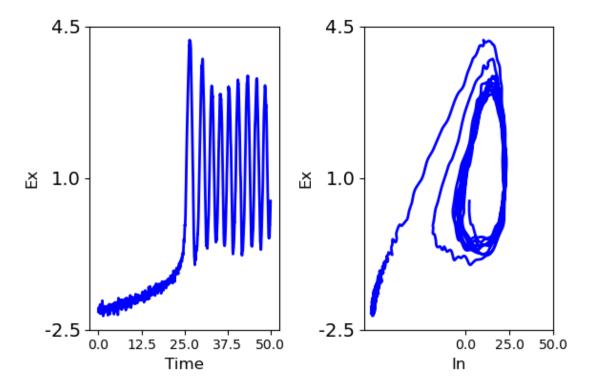
```
[139]: time_begin, time_end = 0, time_stop
fig, ax = plot_series_statespace(time, y, time_begin, time_end, sr)

if pars[1] == 1:
    title_chars = 'Figs/N=1/Onset_SNIC_h_ex.png'

elif pars[1] == 2:
    title_chars = 'Figs/N=1/Onset_Homoclinic_h_ex.png'

# fig.savefig(title_chars, dpi=300, format='png', bbox_inches='tight')
print(title_chars)
```

Figs/N=1/Onset_SNIC_h_ex.png



[]:

2 Try it Yourself

Re-run the above code with these parameter settings:

```
pars = (1.2, 0.1, 4, 6, 6, 0)
h_ex_0 = 0.2
driving_strength = -2
```

to simulate an onset with small fast oscillations that grow in amplitude. This is another common type of seizure onset in humans.

3 Notes on the Reading

3.1 A taxonomy of seizure dynamotypes, Maria Luisa Saggio et al

In this paper, transitions to epileptic seizures as recorded in invasive EEG are classified according to a small number of bifurcations in two-variable dynamical systems. Seizure onset is considered as a transition from fixed point to oscillations and there are four types of such transitions as a function of changes in a single parameter. Saddle-node in invariant cycle (SNIC) lead to sudden onset of large amplitude slow frequency oscillations. Supercritical Hopf bifurcations results in small amplitude fast oscillations with increasing amplitude. Figure 1 is a schematic of all possible types of this low-dimensional approach. Appendix 1, page 29, gives clinical examples of seizure onsets and their interpretation according to dynmical systems theory.