Replication / Computational Neuroscience

[Re] A circuit model of auditory cortex

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Introduction

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Methods

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In this replication, we focus on the rate models proposed in the original article. The firing rate model was an extensions of the traditional Wilson-Cowan model¹ and represented an iso-frequency unit of the auditory cortex. This iso-frequency unit consisted of one excitatory and two inhibitory populations. Building on this unit a more complex three-unit rate models was developed, to investigate stimulus-specific adaptation, forward suppression, tunig-curve adaptation and feedforward functional connectivity.

Iso-Frequency Unit Model The iso-frequency unit model was based on the Wilson-Cowan model¹ but was modified to include two different types of inihbitory interneurons. The two inhibitory population are meant to represent parvalbumin-psoitive (PV) and somatostatin (SST) cells. The single unit model is given by

$$\tau_u \frac{du(t)}{dt} = -u(t) + f(w_{ee}u(t) - w_{ep}p(t) - w_{es}s(t) + qg(t)i(t)), \tag{1}$$

$$\tau_p \frac{dp(t)}{dt} = -p(t) + f(w_{pe}u(t) - w_{pp}p(t) - w_{ps}s(t) + I_{Opt,PV}(t) + qg(t)i(t)),$$
 (2)

$$\tau_s \frac{ds(t)}{dt} = -s(t) + f(w_{se}u(t) - w_{sp}p(t) - w_{ss}s(t) + I_{Opt,SST}(t), \tag{3}$$

with u(t), p(t), and s(t) being the normalized firing rates (in [0,1]) of the pyramidal cell population, the PV population and the SST population, respectively. Furthermore, w_xy represents the strengths of connections from population y to population x. The two terms $I_{Opt,PV}$ and $I_{Opt,SST}$ describe the input current to cells due to optogenetic stimulation of the PV population and SST population, respectively. τ_i , $i \in u, p, s$ defines the time constants for the respective populaitons. The function f is realised as a threshold linear function given by

$$f(x) = \begin{cases} 0 & \text{if } x \le 0 \\ rx & \text{if } 0 \le x \le 1/r \\ 1 & \text{if } x > 1/r, \end{cases}$$

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The authors have declared that no competing interests exists.
Code is available at https://github.com/ChristophMetzner/Park-Geffen-Replication.

Table 1. Overview of the model parameters

а	b	С
1	2	3

(4)

which coarsely approximates a sigmoid function. Furthermore, the function f is thresholded by simply subtracting a constant u_i from the input x (i.e. $f(x-u_i)$) which varied for the different populations. Lastly, afferent auditory input is fed into the unit is given by qg(t)i(t), which is subdivided into the 'raw' input i(t) and a slow modulation g(t) mimicking synaptic depression at thalamic synapses. The input function i(t) is simply an instantaneous rise with amplitude q and an exponential decay with a time constant of τ_q . The synaptic depression g(t) is governed by the following equation

$$\frac{dg(t)}{dt} = \frac{g_0 - g(t)}{\tau_{d_1}} - \frac{g(t)i(t)}{\tau_{d_2}}.$$
 (5)

The parameter values can be found in Table 1

ToDo: Add parameter values to the table.

Three-Unit Model Building on the single unit a three-unit model was implemented, with each single unit representing a different input frequency, thus creating a simple tonotopic layout that allowed to explore more complex auditory inputs. Intra-unit connectivity was as described before for the single-uni model. Inter-unit connectivity was restricted to immediate neighbours and included the following connection types: Exc to exc, exc to PV and SST to exc. Together, the activity of the three populations of each unit was governed by

$$\tau_u \frac{du_i(t)}{dt} = -u_i(t) + f(w_{ee}u_i(t) - (w_{ep} - a(1 - D_i(t)))p_i(t) - w_{es}s_i(t) + J_{1,i}(t)), \quad (6)$$

$$\tau_p \frac{dp_i(t)}{dt} = -p_i(t) + f(w_{pe}u_i(t) - w_{pp}p_i(t) - w_{ps}s_i(t) + I_{Opt,PV}(t) + J_{2,i}(t)), \quad (7)$$

$$\tau_s \frac{ds_i(t)}{dt} = -s_i(t) + f(w_{se}u_i(t) - w_{sp}p_i(t) - w_{ss}s_i(t) + I_{Opt,SST}(t) + J_{3,i}(t)), \quad (8)$$

with

$$J_{1,i}(t) = \begin{cases} -F_i(t)s_2(t) + qI_i(t) + w_{ee}^* u_2(t) & \text{if } i = 1, 3 \\ -F_s(t)(s_1(t) + s_3(t)) + qI_2(t) + \frac{w_{ee}^*(u_1(t) + u_3(t))}{2} & \text{if } i = 2 \end{cases}$$

(9)

$$J_{2,i}(t) = \begin{cases} qI_i(t) + w_{pe}^* u_2(t) & \text{if } i = 1,3\\ qI_2(t) + \frac{w_{pe}^* (u_1(t) + u_3(t))}{2} & \text{if } i = 2 \end{cases}$$
(10)

and

$$J_{3,i}(t) = \begin{cases} qI_i(t) + w_{se}^* u_2(t) & \text{if } i = 1,3\\ qI_2(t) + \frac{w_{se}^* (u_1(t) + u_3(t))}{2} & \text{if } i = 2. \end{cases}$$
(11)

Here, $I_i(t)$ is described by

$$I_k(t) = g_k(t)i_k(t) + g_2(t)i_2(t)\alpha$$
 for $k = 1, 3$ (12)

and

$$I_2(t) = (g_1(t)i_1(t) + g_3(t)i_3(t))\alpha + g_2(t)i_2(t).$$
(13)

Here, $i_k(t)$ represents thalamic inputs to each of the three units. Taken together, the description of the three-unit model is the same than for the single-unit model, except for the addition of lateral inter-unit connectivity and short-term synaptic dynamics. Short-term facilitation is modelled by $F_i(t)$ and increases from 0 to positive values whereas depression is modelled by $D_i(t)$, whic decreases from 1 towards 0. Facilitating synapses were added to Exc to SST inputs and depressing terms to PV to Exc synapses (see²). The facilitating term $F_i(t)$ obeys

$$\frac{dF_j(t)}{dt} = -\frac{F_j(t)}{\tau_{D_1}} + \frac{i_j(t)}{\tau_{D_2}},\tag{14}$$

where τ_{D_1} and τ_{D_2} are again the depression time constants from the input functions of the single-unit model given in Equation 5. Analoguously, the depression term $D_j(t)$ follows

$$\frac{dD_j(t)}{dt} = \frac{1 - D_j(t)}{\tau_{D_1}} - \frac{D_j(t)i_j(t)}{\tau_{D_2}}.$$
 (15)

The choices for the parameters can again be found in Table 1 and were based on experimental studies^{3,4,5}.

Park and Geffen note that, when matching model behaviour to experimental findings, two distinct parameter sets emerged and a unified rate model description required a paradigm-dependent baseline inhibition, which reflected high thalamic activity (corresponding to weak baseline inhibition) versus low thalamic activity (corresponding to strong baseline inhibition). This was implemented using another variable \bar{F} governed by

$$\frac{d\bar{F}(t)}{dt} = \frac{\bar{F}^2(t)}{\tau_{F_1}} - \frac{\bar{I}(t)}{\tau_{F_2}}.$$
 (16)

ToDo: Add paragraph on the variable F

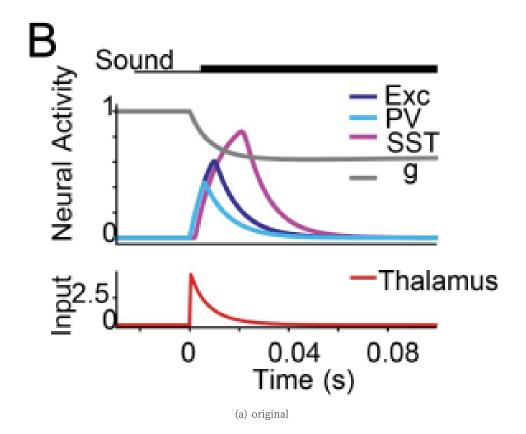
Reimplementation

The iso-frequency unit model and the three-unit model were both implemented in Python and integrated into the neurolib framework⁶.

Reproduction of experiments

Figure 1 compares the input and response profile for the original and the reproduced single-unit model. Qualitatively both models behave similarly, however, quantitative differences can be seen. The thalamic input in the reproduced model only reaches an amplitude of ≈ 1.4 while the original model reaches $\approx 4-5$. Increasing the amplitude of the input (in the reproduced model) did change the qualitative behaviour. **ToDo:** add subplots with changes to the input amplitude; describe the results for the changes; change colours to match original.

Figure 2 compares the input and response profile for the original and the reproduced three-unit model. Here, we see that the first panel (μ_1 activity) qualitatively matches, however, the amplitude of the response to the stimulation of f^* is too high. For the second panel (μ_2 activity), the activity in response to f_1 and f_2 is too high, almost as high as to the preferred frequency f^* . The same pattern can be observed for the third panel (μ_3 activity), where the response to f^* is too high. However, we can see that the input



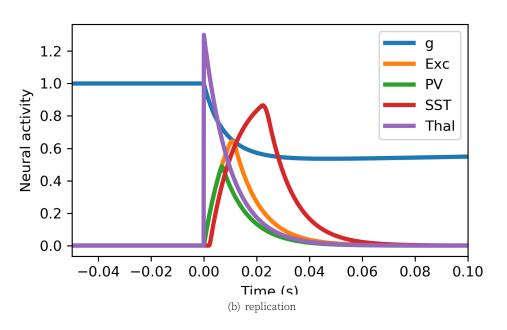
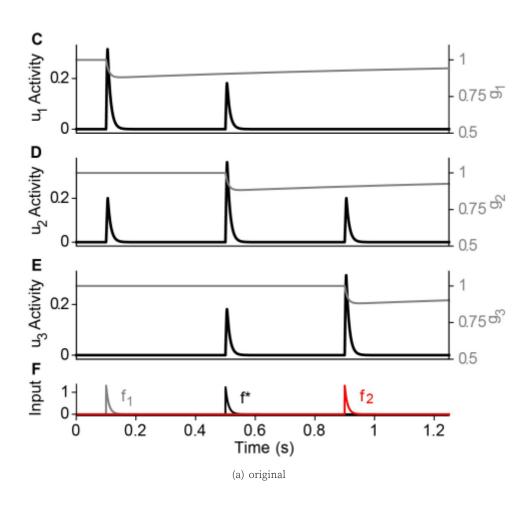


Figure 1. Replicates Figure 1B.

UNDER REVIEW

profile for the three units, original and reproduced model, looks almmost identical to the profile from the reproduced single-unit model (Figure 1 b). ToDo: inlcude g in the plots, see Figure 2C-E;



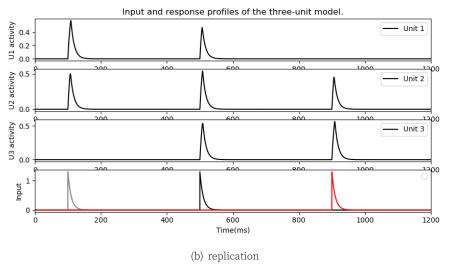


Figure 2. Replicates Figure 2C-F.

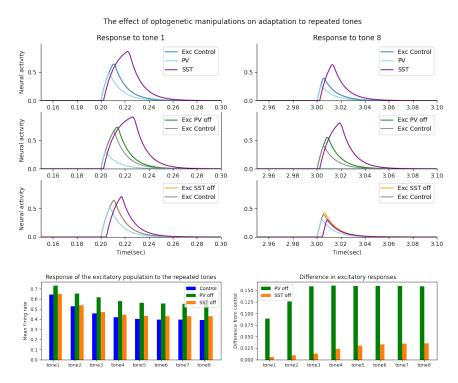
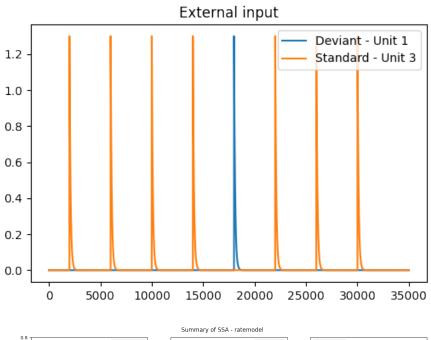


Figure 3. ReFig3



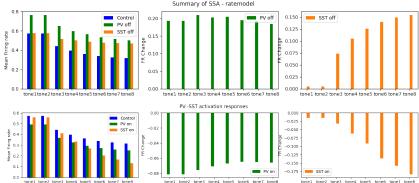


Figure 4. ReFig4

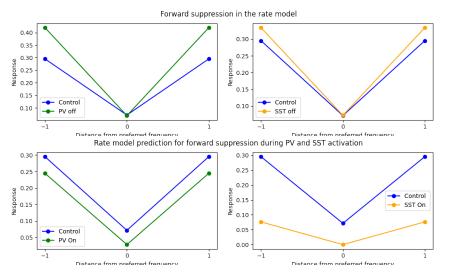


Figure 5. ReFig6

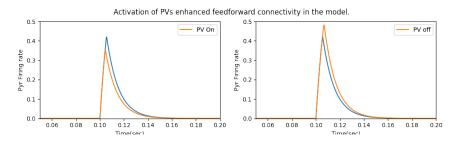


Figure 6. ReFig8

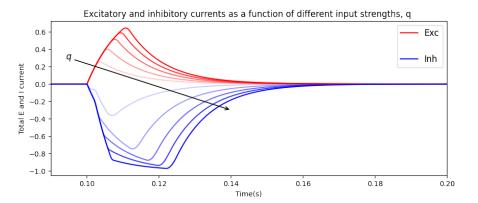


Figure 7. ReFig9

Discussion

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