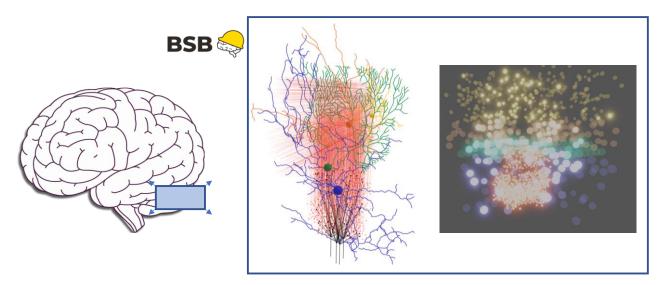
BSB design and usecases



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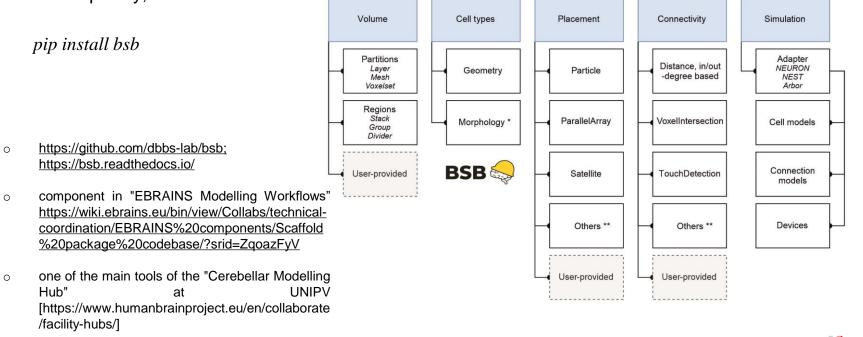


Human Brain Project



Network reconstruction and simulation

the **Brain Scaffold Builder**, a flexible software package to build and simulate brain models at different levels of complexity, constrained on data



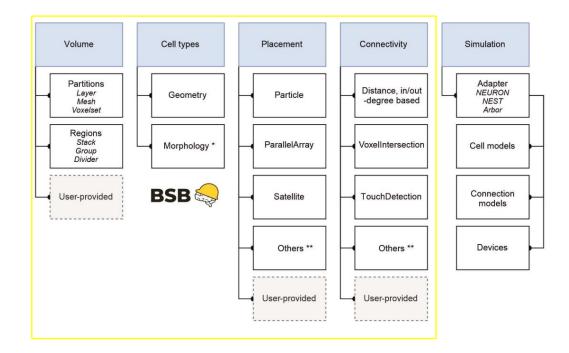
Cerebellar usecase: *De Schepper et al.,* <u>https://www.biorxiv.org/content/10.1101/2021.07.</u> <u>30.454314v1</u>

0

CSCS Centro Svizzero di Calcolo Scientifico Swiss National Supercomputing Centre



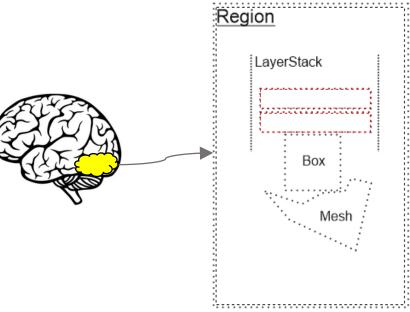
Network reconstruction



```
Volume
       },
       "network architecture": {
         "simulation volume x": 300.0,
         "simulation volume z": 200.0,
         "store kd trees": true,
         "store compound kd tree": true,
         "store pf kd trees": true
       },
       "layers": {
         "granular layer": {
           "thickness": 130.0,
           "stack": {
             "stack id": 0,
             "position in stack": 0,
             "position": [0.0, 0.0, 0.0]
         "purkinje layer": {
           "thickness": 15.0,
           "stack": {
             "stack id": 0,
             "position in stack": 1
         "molecular layer": {
           "thickness": 150.0,
           "stack": {
             "stack id": 0,
             "position in stack": 2
       },
```

[µm] x	[µm] z	[µm] y	[µm³]
300	200	295	17700000

x-y (sagittal plane), *x-z* (horizontal plane), *z-y* (coronal plane), *z*-axis (major lamellar axis, along which parallel fibers elongate)





Cell types

```
"cell types": {
  "granule cell": {
    "placement": {
     "class": "bsb.placement.ParticlePlacement",
      "layer": "granular layer",
      "soma radius": 2.5,
      "density": 3.9e-3
   },
    "morphology": {
      "class": "bsb.morphologies.NoGeometry",
      "dendrite length": 40,
      "detailed morphologies": {
        "names": ["GranuleCell"]
                                                11
      }
   },
                                                  },
                                                  },
```

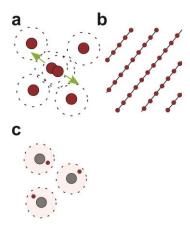
```
},
"purkinje_cell": {
    "placement": {
        "class": "bsb.placement.ParallelArrayPlacement",
        "layer": "purkinje_layer",
        "soma_radius": 7.5,
        "planar_density": 0.0017,
        "extension_x": 130.0,
        "extension_z": 3.5,
        "angle": 70
}
```

```
"morphology": {
    "class": "bsb.morphologies.NoGeometry",
    "detailed_morphologies": {
        "names": ["PurkinjeCell"]
    }
}
```

Placement

Neuron densitiesNeuron geometry

"placement": {
 "fill_base_layer": {
 "cls": "bsb.placement.ParticlePlacement",
 "cell_types": ["cell_type_A", "cell_type_B"],
 "partitions": ["base_layer"]
 },
 "load_atlas": {
 "cls": "bsb.placement.atlas.NRRD",
 "cell_types": ["cell_type_C"],
 "partitions": ["nucleus_voxelset", "deep_nucleus_voxelset"]
 }
}



Particle Placement algorithm for all cell types except for PCs (random placement, collisions/replacement, pruning)

Planar arrays placement algorithm for PCs

	N	
mossy fibers	117	
glomeruli	2336	
granule cells	28615	
golgi cells	70	
purkinje cells	99	
basket cells	147	
stellate cells	299	
tot cells	29230	
tot entities/relays	2453	
TOT elements	31683	

Ito et al., 1984

network_scene.html

Connectivity

d

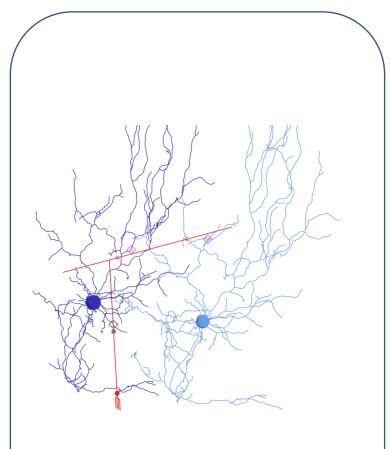
P(d)·P(I)·P(O)



"B_to_C": {
 "cls": "bsb.connectivity.TouchDetection",
 "from_type": "cell_type_B",
 "to_type": "cell_type_C",
 "intersection_radius": 2.0

},

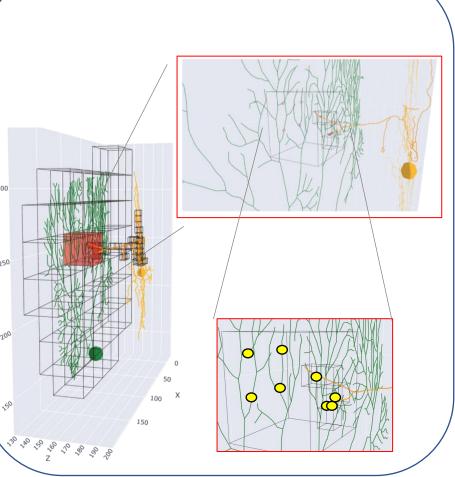
```
"A_to_C": {
    "cls": "bsb.connectivity.VoxelIntersection",
    "from_type": "cell_type_A",
    "to_type": "cell_type_C",
    "synapses": {
        "type": "norm",
        "loc": 4,
        "scale": 0.4
    }
}
```

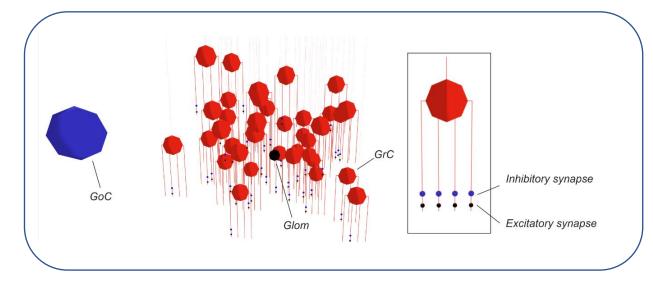


grc_golgi_touch_v2.html

Connectivity

```
"connectivity": {
 "A_to_B": {
   "cls": "bsb.connectivity.AllToAll",
   "from_type": "cell_type_A",
   "to_type": "cell_type_B"
 },
                                                     300
  "B_to_C": {
   "cls": "bsb.connectivity.TouchDetection",
   "from_type": "cell_type_B",
   "to_type": "cell_type_C",
   "intersection_radius": 2.0
                                                     250
  },
  "A to C": {
                                                    Υ
   "cls": "bsb.connectivity.VoxelIntersection",
   "from_type": "cell_type_A",
   "to_type": "cell_type_C",
                                                      200
   "synapses": {
     "type": "norm",
     "loc": 4,
     "scale": 0.4
                                                      150
```





Connectivity

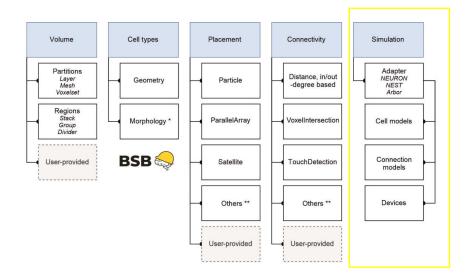
Ad-hoc algorithms, based on experimental evidences (divergence and convergence ratios, and geometrical neuronal features)

	Convergence	Divergence	Synapses	Synapses/pair
mf-Glom	1 ± 0	20 ± 8	2300	1 ± 0
Glom-GrC	4 ± 0	49 ± 26	110000	1 ± 0
Glom-GoC	56 ± 21	1.7 ± 1.4	3900	1 ± 0
GoC-GrC	2.4 ± 0.88	1000 ± 460	97000	1.4 ± 0.66
GoC-Glom	0.84 ± 0.37	28 ± 14	2000	1 ± 0
GoC-GoC	16 ± 6.5	16 ± 7.1	180000	160 ± 5
GrC (<i>aa</i>)-GoC	320 ± 230	0.78 ± 1.1	22000	1 ± 0
GrC (<i>aa</i>)-PC	82 ± 24	0.28 ± 0.52	20000	2.4 ± 1.1
GrC (<i>pf</i>)-GoC	910 ± 360	2.2 ± 1.6	64000	1 ± 0
GrC (<i>pf</i>)-PC	1500 ± 300	5.1 ± 2.6	140000	1 ± 0
GrC (<i>pf</i>)-SC	480 ± 160	5.1 ± 3.2	140000	1 ± 0
GrC (<i>pf</i>)-BC	740 ± 130	3.8 ± 2.1	110000	1 ± 0
SC-PC	5.4 ± 2.7	1.8 ± 1.5	530	1 ± 0
BC-PC	20 ± 9.9	14 ± 8	2000	1 ± 0
SC-SC	14 ± 6.1	14 ± 5.9	430000	100 ± 4
BC-BC	14 ± 6.7	14 ± 6.6	200000	100 ± 4
GoC-GoC (gap)	8.4 ± 3.5	8.4 ± 3.5	2100	3.5 ± 1.6

~ 1'500'000 chemical synapses and 2'100 electrical synapses

The **cerebellar connectome** was generated through appropriate connection rules, unifying a collection of scattered experimental data into a coherent construct and providing a **new model-based ground-truth** about circuit organization.

Network simulation



detailed or point-neuron networks





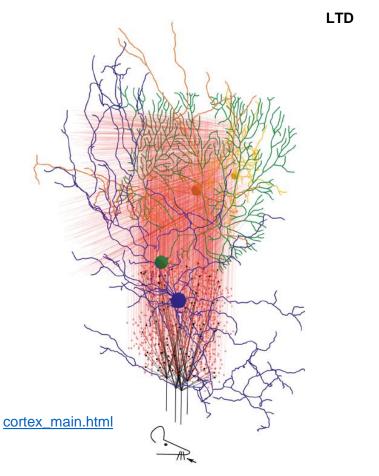
Release probabilities at mf-GrC synapses control

Time [ms]

LTP

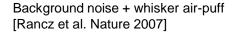
50 pA

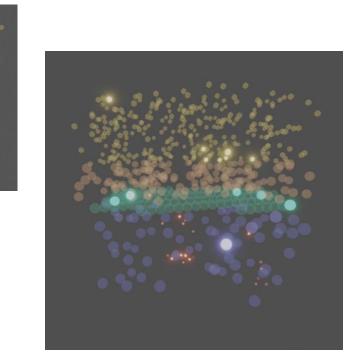
200 ms

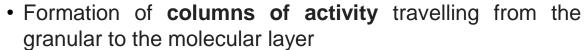


Functional validation against in vivo data, monitoring the impact of subcellular and cellular mechanisms on spatio-temporal signal processing

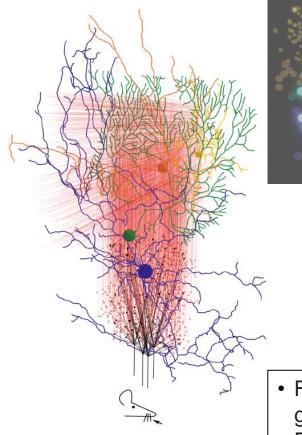
- ✓ Background frequency of all cerebellar neuron types
- ✓ Synchronous oscillatory behaviour in theta band of granular layer in resting state
- ✓ Impulsive responses of all cerebellar neuron types
- ✓ Burst-pause response of Purkinje Cells (PCs)
- ✓ Feedforward and lateral inhibition from Molecular Layer Interneurons (MLIs, i.e. Stellate Cells (SCs) and Basket Cells (BCs)) to Purkinje Cells

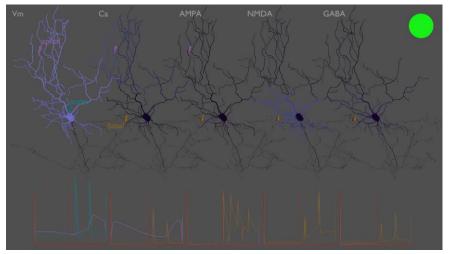


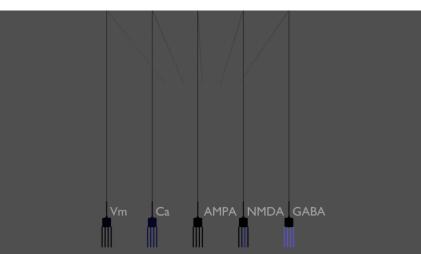


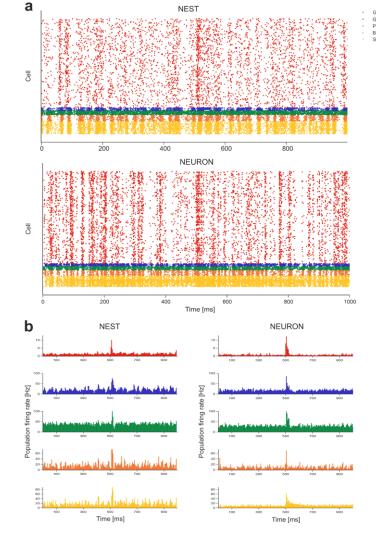


• Prediction of the impact of cellular mechanisms on signal propagation and **spatio-temporal processing**

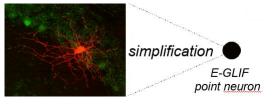








The reconstructed cerebellar network was simulated using the BSB NEST and NEURON Adapters. The simulation lasted 1 second, with background at 4Hz on all mfs and a burst on 4 adjacent mfs starting at 500 ms and lasting 20 ms. For NEST version, optimized E-GLIF neuron models and alpha-based conductance synapses (Geminiani et al., 2019a) were inserted. a) Raster plot of all cells; GrCs are undersampled (random 10%) for clarity. b) Peri-Stimulus-Time-Histogram (PSTH) of each population (number of spikes in 5 ms time bins, normalized on the total number of cells)



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Claudia Gandini Wheeler-Kingshott Fulvia Palesi Gloria Castellazzi (IRCCS Mondino) Giovanni Savini (IRCCS Mondino) Nick Rolandi Anita Monteverdi Roberta Lorenzi Marta Gaviraghi





Collaborations

UCLION

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