

# Robustness in TDMA Scheduling for Neuron-based Molecular Communication

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## ABSTRACT

This paper proposes and evaluates an optimizer for neuron-based body-area nanonetworks (BANNs). The proposed optimizer leverages an evolutionary algorithm to seek the optimal trade-off between communication latency and robustness in TDMA-based neuronal signaling. Simulation results demonstrate that the proposed optimizer efficiently obtains quality solutions and multiobjective analysis is critical in configuring neuron-based BANNs.

## Categories and Subject Descriptors

C.2.2 [Computer-Communication Networks]: Network Protocols; C.3 [Special-purpose and Application-based Systems]: *Signal processing systems*; I.2.8 [Artificial Intelligence]: Problem Solving, Control Methods, and Search—*Heuristic methods*

## General Terms

Algorithms

## Keywords

Molecular communication, Neuronal networks, TDMA scheduling, Evolutionary multiobjective optimization algorithms

## 1. INTRODUCTION

Molecular communication is a communication paradigm that utilizes molecules as a communication medium between nanomachines. Nanomachines are nanoscale devices that perform simple computation, sensing and/or actuation tasks [18]. They may be man-made devices built in the *top-down* approach, downscaling the current microelectronic and micro-electro-mechanical technologies or in the *bottom-up* approach, assembling synthesized nanomaterials such as graphene nano ribbons and carbon nanotubes [1]. Alternatively, nanomachines may be *bio-hybrid*, integrating man-made nanostructures with biological components such as DNA strands, antibodies and molecular motors [1]. Due to its advantages

such as inherent nanometer scale, biocompatibility and energy efficiency [6], a key application domain of molecular communication is body area nanonetworks (BANNs), where nanomachines are networked through molecular communication to perform their tasks in the body for biomedical and prosthetic purposes (e.g., vital information sensing, targeted drug release and neural signal augmentation) [3].

An approach to BANNs is to utilize neurons as a primary communication component [4, 19, 20]. A neuron-based BANN consists of a set of nanomachines and a network of neurons that are artificially formed into a particular topology. It allows nanomachines to interface (i.e., activate and deactivate) neurons and communicate to other nanomachines with electrochemical signals through a chain of neurons.

This paper focuses on a communication protocol framework, called Neuronal TDMA [20], which performs single-bit Time Division Multiple Access (TDMA) scheduling for neuron-based BANNs. Neuronal TDMA allows nanomachines to multiplex and parallelize neuronal signaling while avoiding signal interference to ensure that signals reach the destination. This paper proposes and evaluates a particular optimizer for Neuronal TDMA. The proposed optimizer seeks the optimal signaling schedules (i.e., which neurons to activate and when to activate them to trigger signal transmissions) for nanomachines with respect to communication latency and robustness. Communication robustness refers to the sensitivity of a signaling schedule to environmental noise that can cause signal interference.

Latency and robustness conflict with each other in neuronal signaling. For example, improving (i.e., decreasing) latency often means improving throughput in a neuronal network by increasing the degree of multiplexing. This can degrade robustness because a highly-multiplexed network has a higher risk of signal interference due to environmental noise. Conversely, improving robustness often means multiplexing neuronal signals less often. This can degrade latency because a rarely-multiplexed network tends to be low-throughput.

A goal of the proposed optimizer in Neuronal TDMA is to reveal the optimal trade-offs between communication latency and robustness subject to given constraints. Since there exists no single optimal solution (TDMA schedule) under conflicting objectives but rather a set of alternative solutions of equivalent quality, the proposed optimizer is designed to seek Pareto-optimal solutions that are distributed well in the objective space. Therefore, it can produce both *extreme* TDMA schedules (e.g., the one yielding low latency and low robustness) and *balanced* schedules (e.g., the one yielding moderate latency and robustness) at the same time.

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Given a set of heuristically-approximated Pareto-optimal TDMA schedules, the proposed optimizer allows the BANN designer to examine the trade-offs among them and make a well-informed decision to choose one of them, as the best TDMA schedule, according to his/her preferences and priorities. For example, the BANN designer can examine how he/she can trade (or sacrifice) latency for robustness and determine a particular TDMA schedule that yields a desirable/comfortable balance between latency and robustness.

Simulation evaluation results show that the proposed optimizer in Neuronal TDMA efficiently obtains quality solutions with acceptable computational costs and verify multiobjective analysis is critical in configuring and operating neuron-based BANNs.

## 2. BACKGROUND

This section describes the structural and behavioral properties of neurons. Neurons are a fundamental component of the nervous system, which includes the brain and the spinal cord. They are electrically excitable cells that process and transmit information via electrical and chemical signaling.

The structure of a neuron consists of a cell body (or soma), dendrites and an axon (Figure 1). The soma is the central part of a neuron. It can vary from 4 to 100 micrometers in diameter. Dendrites are thin structures that arise from the soma. They form a complex “dendritic tree” that extends the farthest branch a few hundred micrometers from the soma. Dendrites are where the majority of inputs to a neuron occur. An axon is a cellular extension that arises from the soma. It branches before it terminates and travels through the body in bundles called nerves. Its length can be over one meter in the human nerve that arises from the spinal cord to a toe.

Neurons are connected with each other to form a network(s). Neurons communicate with others via *synapses*, each of which is a membrane-to-membrane junction between two neurons. A synapse contains molecular machinery that allows a (presynaptic) neuron to transmit a chemical signal to another (postsynaptic) neuron. In general, signals are transmitted from the axon of a presynaptic neuron to a dendrite of a presynaptic neuron. An axon transmits an output signal to a postsynaptic neuron, and a dendrite receives an input signal from a presynaptic neuron.

Presynaptic and postsynaptic neurons maintain voltage gradients across their membranes by means of voltage-gated ion channels, which are embedded in the presynaptic membrane to generate the differences between intracellular and extracellular concentration of ions (e.g.,  $\text{Ca}^{2+}$ ) [17]. Changes in the cross-membrane ion concentration (i.e., voltage) can alter the function of ion channels. If the concentration (i.e., voltage) changes by a large enough amount (e.g., approximately 80 mV in a giant squid), ion channels initiate a voltage-dependent process; they pump extracellular ions inward. Upon the increase in intracellular ion concentration, the presynaptic neuron releases a chemical called a *neurotransmitter* (e.g., acetylcholine (ACh)), which travels through the synapse from the presynaptic neuron’s axon terminal to the postsynaptic neuron’s dendrite. The neurotransmitter electrically excites the postsynaptic neuron, and the neuron generates an electrical pulse called an *action potential*. This signal travels rapidly along the neuron’s axon and activates synaptic connections (i.e., opens ion channels) when it arrives at the axon’s terminals. This way, an action potential

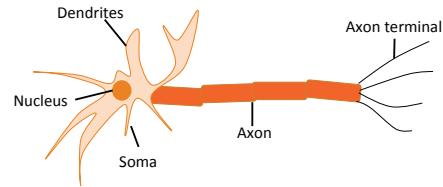


Figure 1: The structure of neurons

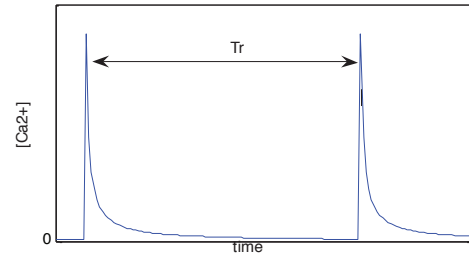


Figure 2: Intracellular  $\text{Ca}^{2+}$  concentration

triggers cascading neuron-to-neuron communication.

Figure 2 shows how  $\text{Ca}^{2+}$  concentration changes in a neuron. When the concentration peaks, the neuron releases a neurotransmitter to trigger an action potential. Upon a neurotransmitter release, the neuron goes into a *refractory period* ( $T_r$  in Figure 2), which is the time required for the neuron to replenish its internal  $\text{Ca}^{2+}$  store. During  $T_r$ , it cannot process any incoming signals. The refractory period is approximately two milliseconds in a giant squid.

## 3. RELATED WORK

There exist two major thrusts in communication research for body area nanonetworks (BANNs). One thrust is leveraging electromagnetic terahertz communication with, for example, graphene-based nanoscale antennas [2]. The other is molecular communication, which this paper focuses on. In general, molecular communication maintains several advantages over electromagnetic communication such as biocompatibility and energy efficiency [3, 6].

Compared with other molecular communication approaches (e.g., molecular motors [12], calcium signaling [14] and bacteria communication [10]), neuron-based communication has such advantages as long distance coverage, high speed signaling (up to 90 m/s [8]) and low attenuation in signaling [6].

Balasubramaniam et al. first examined TDMA communication for neuronal signaling [4]. Neuronal TDMA extended it with a multiobjective optimization algorithm that considers communication performance objectives such as signaling yield, fairness and latency [20]. This paper extends Neuronal TDMA by analyzing the optimal trade-offs between communication performance (latency) and robustness.

Tezcan et al. address communication robustness in TDMA-based neural signaling by proposing a signal buffering mechanism with neural delay lines, which parallel fiber delay lines in optical network switching. The proposed optimizer in this paper is similar to their mechanism in that both aim to avoid signal interference. However, this paper addresses robustness in the context of optimizing TDMA schedules while Tezcan et al. do not consider TDMA scheduling.

#### 4. NEURON-BASED BODY AREA NANONETWORKS (BANNs)

This paper assumes neural signaling in a network of natural neurons that are artificially grown and formed into particular topology patterns. This assumption is made upon numerous research efforts to grow neurons on substrates (e.g., [15]) and design topologically-specific neuronal networks (e.g., [9, 13, 21]).

Figure 3 illustrates an example neuron-based BANN. It contains an artificially-grown neuronal network and several nanomachines such as sensors and a sink. Sensors use neuronal signaling to transmit sensor data to the sink, which might work as an actuator or transducer. As potential applications, prosthetic devices and medical rehabilitation devices could leverage neuron-based BANNs to better perform sensing, transducing and actuation tasks in the body.

This paper assumes that nanomachines (e.g., sensors) interact with neuronal networks in a *non-invasive* manner. This means that it is not required to insert carbon nanotubes into neurons so that nanomachines can trigger signaling. Nanomachines may use a neurointerface based on chemical agents (e.g., acetylcholine and mecamylamine [4]) or light [7].

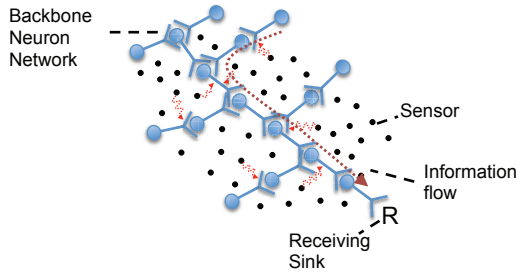


Figure 3: An Example Neuron-based BANN

#### 5. NEURONAL TDMA

Neuronal TDMA performs a single-bit TDMA communication that periodically assigns a *time slot* to each sensor. Sensors fire neurons, one after the other, each using its own time slot. This allows multiple sensors to transmit signals to the sink through the shared neuronal network. Each sensor transmits a single signal (a single bit) within a single time slot. This single-bit-per-slot design is based on two assumptions: (1) a signal (i.e., action potential) is interpreted with two levels of amplitudes, which represent 0 and 1, and (2) after a signal transmission, a neuron goes into a refractory period (waiting/sleeping period).

An important goal of Neuronal TDMA is to avoid signal interference, which occurs when multiple signals fire the same neuron at the same time and leads to corruption of transmitted sensor data at the sink. Signals can easily interfere with each other if sensors fire their neighboring neurons randomly. Neuronal TDMA is intended to eliminate signal interference by scheduling which sensors fire which neurons with respect to time. An optimizer in Neuronal TDMA seeks the optimal TDMA schedules for a set of sensors in a given neuronal network.

Figure 4 shows an example body area nanonetwork (BANN) that contains four nanomachines (three sensors and a sink) and a network of five neurons ( $n_1$  to  $n_5$ ). Figure 5 illustrates an example TDMA schedule for those sensors to fire neurons.

The scheduling cycle period lasts 6 time slots ( $T_s = 5$ ). The sensor  $s_1$  fires the neuron  $n_4$  to initiate signaling in the first time slot  $T_1$ . The signal travels through  $n_5$  in the next time slot  $T_2$  to reach the sink. The sensor  $s_2$  transmits a signal on  $n_3$  in  $T_2$ . During  $T_2$ , two signals travel in the neuronal network in parallel. The duration of each time slot must be equal to, or longer than, the refractory period  $T_r$  (Figure 2).

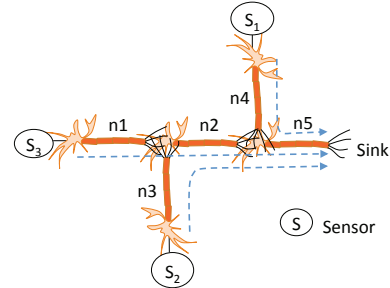


Figure 4: An Example Neuron-based BANN

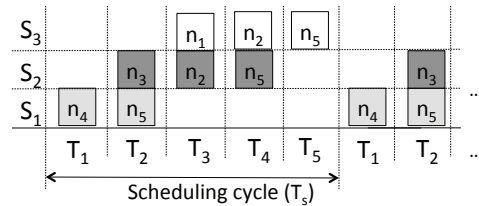


Figure 5: An Example TDMA Schedule

#### 5.1 Scheduling Problem in Neuronal TDMA

The scheduling problem in Neuronal TDMA is defined as an optimization problem where a neuron-based BANN contains  $M$  sensors,  $S = \{s_1, s_2, \dots, s_i, \dots, s_M\}$ , and  $N$  neurons,  $N = \{n_1, n_2, \dots, n_j, \dots, n_N\}$ .

The proposed optimizer considers two optimization constraints. The first constraint enforces that at most one signal can pass through each neuron in a single time slot. (Otherwise, signal interference occurs.) The second constraint enforces each sensor transmits one signal during the scheduling cycle  $T_s$ . A TDMA schedule is said to be feasible if it never violates constraints. On the contrary, it is said to be infeasible if it violates any of these two constraints. An example TDMA schedule in Figure 5 is feasible.

The proposed optimizer considers two optimization objectives: (1) communication latency and (2) communication robustness. Latency and robustness are to be minimized and maximized, respectively.

Communication latency ( $f_L$ ) indicates how soon the sink receives all signals from all of  $M$  sensors. It is computed as follows.

$$f_L = \max_{s_i \in S} t_a^{s_i} \quad (1)$$

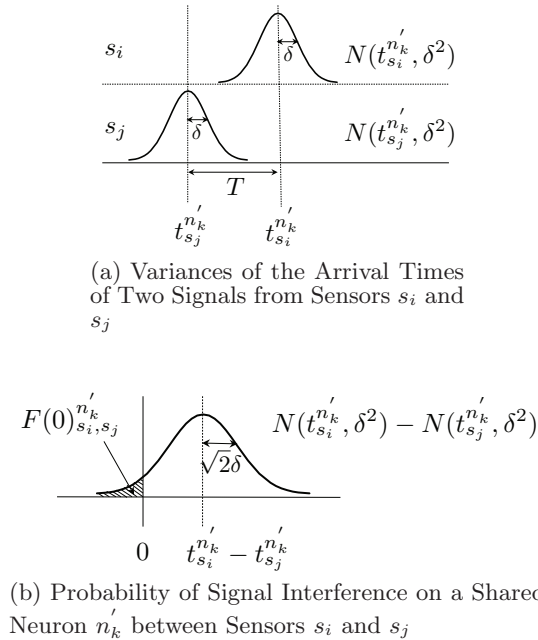
$t_a^{s_i}$  denotes the arrival time at which the sink receives a signal that  $s_i$  transmits.  $f_L$  determines the scheduling cycle period  $T_s$  ( $T_s = f_L$ ). In Figure 5,  $f_L = 5$ .

Communication robustness ( $f_R$ ) indicates the probability that signals do not interfere with each other on shared neurons  $N' = \{n'_1, n'_2, \dots, n'_k, \dots, n'_K\} \in N$ . Shared neurons are the neurons that sensors share to transmit their signals to the sink. In Figure 5,  $n_2$  and  $n_5$  are shared neurons. ( $s_2$

and  $s_3$  share  $n_2$ .  $s_1$ ,  $s_2$  and  $s_3$  share  $n_5$ .) Higher robustness means lower chances of signal interference that can occur due to variances of signal transmission speed in a neuron even if the *one-signal-per-slot* constraint (see above) is satisfied. In Figure 5,  $s_2$ 's signal travels on  $n_2$  in  $T_3$ , and  $s_3$ 's signal travels on  $n_2$  in  $T_4$ . This signaling schedule satisfies the *one-signal-per-slot* constraint; however, the two signals can interfere if  $s_2$ 's signal arrives at  $n_2$  during  $T_4$  due to variances signal transmission speed in  $n_2$  and/or if  $s_3$ 's signal arrives at  $n_2$  in  $T_3$  due to signaling speed variances in  $n_1$ .

A major contributor to the variances of signal transmission speed is *synaptic delay*, which is the interval between the arrival of an action potential at a presynaptic axon terminal and the start of a postsynaptic action potential. This interval is the time required for a neurotransmitter to be released from a presynaptic membrane, diffuse across the synaptic gap and bind to a receptor site on the post-synaptic membrane. It is known that a synaptic delay varies from 0.3–0.5 milliseconds to several milliseconds.

Figure 6 shows a statistical model to determine the probability of signal interference. Figure 6(a) shows two signals from  $s_1$  and  $s_2$  and their expected arrival times at  $n_k$  ( $t_{s_i}^{n_k}$  and  $t_{s_j}^{n_k}$ ).  $T$  represents the interval between the two arrival times. Arrival times follow a normal distribution with a standard deviation of  $\sigma$ . Figure 6(b) depicts  $N(t_{s_i}^{n_k}, \sigma^2) - N(t_{s_j}^{n_k}, \sigma^2)$ , which illustrates the probability that two distributions ( $N(t_{s_i}^{n_k}, \sigma^2)$  and  $N(t_{s_j}^{n_k}, \sigma^2)$ ) overlap.  $F$  is the cumulative distribution function for  $N(t_{s_i}^{n_k}, \sigma^2) - N(t_{s_j}^{n_k}, \sigma^2)$ .  $F(0)_{s_i, s_j}^{n_k}$  denotes the probability that two signals from  $s_1$  and  $s_2$  do not interfere.



**Figure 6: Robustness in Neural Signaling**

The communication robustness objective ( $f_R$ ) is computed as follows by generalizing the model described in Figure 6.

$$f_R = - \sum_{i=1}^M \sum_{j=1}^M \sum_{k=1}^K \left( F(0)_{s_i, s_j}^{n_k} \times I_s \right) \quad (2)$$

$$\text{where } F(0)_{s_i, s_j}^{n_k} = \frac{1}{2} \left( 1 + \frac{1}{\sqrt{\pi}} \int_{-\frac{T}{2\sigma}}^{\frac{T}{2\sigma}} e^{-x^2} dx \right)$$

Note that  $I_s = 1$  if  $t_{s_i}^{n_k} > t_{s_j}^{n_k}$ ; otherwise,  $I_s = 0$ .

## 5.2 The Proposed Optimizer in Neuronal TDMA

The proposed optimizer in Neuronal TDMA leverages an evolutionary multiobjective optimization algorithm (EMOA) to solve its scheduling optimization problem. The algorithm iteratively evolves the population of solution candidates, called *individuals*, through several operators (e.g., crossover, mutation and selection operators) toward the Pareto-optimal solutions in the objective space.

In order to seek Pareto optimality, the notion of *dominance* plays an important role [16]. An individual  $i$  is said to dominate an individual  $j$  if both of the following conditions are hold.

- $i$ 's objective values are superior than, or equal to,  $j$ 's in all objectives.
- $i$ 's objective values are superior than  $j$ 's in at least one objectives.

In Neuronal TDMA, each individual represents a particular TDMA schedule for  $M$  sensors. Figure 7 shows the structure of an example individual, which represents the schedule shown in Figure 5. In this example, the first sensor,  $s_1$ , fires its neighboring neuron,  $n_4$ , in the first time slot  $T_1$ . Similarly,  $s_2$  and  $s_3$  fire their neighboring neurons ( $n_2$  and  $n_1$ ) in  $T_2$  and  $T_3$ , respectively.

$S_3$	0	0	1	0	0
$S_2$	0	1	0	0	0
$S_1$	1	0	0	0	0
	$T_1$	$T_2$	$T_3$	$T_4$	$T_5$

**Figure 7: Individual Representation**

Algorithm 1 shows how the proposed optimizer works. It follows the algorithmic structure in NSGA-II [5]. At the  $\theta$ -th generation,  $N$  individuals are randomly generated as the initial population  $\mathcal{P}_0$  (Line 2).

In each generation ( $g$ ), two parent individuals ( $p_1$  and  $p_2$ ) are selected from the current population  $\mathcal{P}_g$  with binary tournaments (Lines 6 and 7). A binary tournament randomly takes two individuals from  $\mathcal{P}_g$ , compares them based on the notion of dominance, and chooses a superior one as a parent. With the crossover rate  $P_c$ , two parents reproduce two offspring through crossover (Lines 8 to 10). Then, mutation occurs on each offspring (Lines 11 to 16). It randomly alters a neuron-firing pattern with the mutation rate  $P_m$ . The binary tournament, crossover and mutation operators are executed repeatedly on  $\mathcal{P}_g$  to reproduce  $N$  offspring. The offspring ( $\mathcal{O}_g$ ) are combined with the parent population  $\mathcal{P}_g$  to form  $\mathcal{R}_g$  (Line 19).

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**Algorithm 1** Optimization Process

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```
1:  $g = 0$ ;  
2:  $\mathcal{P}_g =$  Randomly generated  $N$  individuals;  
3: while  $g < G_{max}$  do  
4:    $\mathcal{O}_g = \emptyset$ ;  
5:   while  $|\mathcal{O}_g| < N$  do  
6:      $p_1 =$  tournament( $\mathcal{P}_g$ )  
7:      $p_2 =$  tournament( $\mathcal{P}_g$ )  
8:     if random()  $\leq P_c$  then  
9:        $\{o_1, o_2\} =$  crossover( $p_1, p_2$ )  
10:    end if  
11:    if (random()  $\leq P_m$ ) then  
12:       $o_1 =$  mutation( $o_1$ )  
13:    end if  
14:    if random()  $\leq P_m$  then  
15:       $o_2 =$  mutation( $o_2$ )  
16:    end if  
17:     $\mathcal{O}_g = \{o_1, o_2\} \cup \mathcal{O}_g$   
18:  end while  
19:   $\mathcal{R}_g = \mathcal{P}_g \cup \mathcal{O}_g$   
20:   $\mathcal{F} =$  sortByDominationRanking( $\mathcal{R}_g$ )  
21:   $\mathcal{P}_{g+1} = \{\emptyset\}$   
22:   $i = 1$   
23:  while  $|\mathcal{P}_{g+1}| + |\mathcal{F}_i| \leq N$  do  
24:     $\mathcal{P}_{g+1} = \mathcal{P}_{g+1} \cup \mathcal{F}_i$   
25:     $i = i + 1$   
26:  end while  
27:  sortByCrowdingDistance( $\mathcal{F}_i$ )  
28:   $\mathcal{P}_{g+1} = \mathcal{P}_{g+1} \cup \mathcal{F}_i[1 : (N - |\mathcal{P}_{g+1}|)]$   
29:   $g = g + 1$   
30: end while
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Environmental selection follows reproduction. Best  $N$  individuals are selected from  $2N$  individuals in  $\mathcal{R}_g$  as the next generation population ( $\mathcal{P}_{g+1}$ ). First, the individuals in  $\mathcal{R}_g$  are ranked based on the dominance relationships among them. Non-dominated individuals are on the first rank. The  $i$ -th rank consists of the individuals dominated only by the individuals on the  $(i - 1)$ -th rank. Ranked individuals are stored in  $\mathcal{F}$  (Line 20).  $\mathcal{F}_i$  contains the  $i$ -th rank individuals.

Then, the individuals in  $\mathcal{F}$  move to  $\mathcal{P}_{g+1}$  on a rank by rank basis, starting with  $\mathcal{F}_1$  (Lines 23 to 26). If the number of individuals in  $\mathcal{P}_{g+1} \cup \mathcal{F}_i$  is less than  $N$ ,  $\mathcal{F}_i$  moves to  $\mathcal{P}_{g+1}$ . Otherwise, a subset of  $\mathcal{F}_i$  moves to  $\mathcal{P}_{g+1}$ . The subset is selected based on the crowding distance metric, which measures the distribution (or diversity) of individuals in the objective space [5] (Lines 27 and 28). The metric computes the distance between two closest neighbors of an individual in each objective and sums up the distances associated with all objectives. A higher crowding distance means that an individual in question is more distant from its neighboring individuals in the objective space. In Line 27, the individuals in  $\mathcal{F}_i$  are sorted from the one with the highest crowding distance to the one with the lowest crowding distance. The individuals with higher crowding distance measures have higher chances to be selected to  $\mathcal{P}_{g+1}$  (Line 28).

## 6. EVALUATION

This section evaluates the proposed optimizer in Neuronal TDMA through simulations.

### 6.1 Simulation Configurations

This paper simulates a neuronal network that contains 43 neurons (Figures 8). 11 sensors are evenly distributed in the network. The network topology is generated with a tree structure generation algorithm based on Diffusion Limited

Aggregation (DLA) [11].

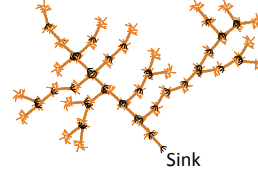


Figure 8: A Simulated Neuronal Network

The proposed optimizer is configured with a set of parameters shown in Table 1.  $Q$  denotes the total number of time slots in an individual ( $Q = 15$  in Figure 7).  $T_u$  denotes the amount of time allocated to each time slot.

Table 1: EMOA Configurations

Parameter	Value
Max. # of generations ( $g_{max}$ in Figure 1)	100
Population size ( $N$ in Figure 1)	100
Crossover rate ( $P_c$ in Figure 1)	0.9
Mutation rate ( $P_m$ in Figure 1)	$1/Q$
$\sigma$ in Equation 2	$T_u/3$

## 6.2 Simulation Results

Figure 9(a) shows how the proposed optimizer increases the union of the hypervolumes (HVs) that individuals dominate in the objective space as the number of generations grows. The HV metric quantifies the optimality and diversity of individuals [22]. A higher HV means that individuals are closer to the Pareto-optimal front and more diverse in the objective space. As Figure 9(a) shows, the proposed optimizer rapidly increases its HV measure in the first 20 generations and converges around the 60th generation. At the last generation, all individuals are non-dominated in the population. This demonstrates that the proposed optimizer allows individuals to efficiently evolve and improve their quality and diversity within 100 generation.

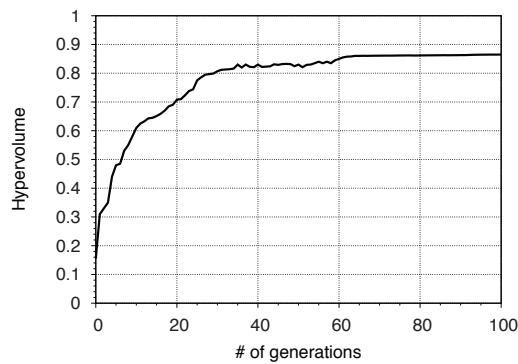
Figure 9(b) shows an objective space that plots the individuals obtained at the last generation. Those individuals approximate the Pareto front (i.e., the optimal trade-off) between latency and robustness. Note that the robustness of  $-20\%$  means the signal interference probability of  $20\%$ . Given objective value ranges ( $16 \leq T_L \leq 47$  and  $-40.77 \leq T_R \lesssim 0$ ), the BANN designer can examine how he/she can trade latency for robustness and choose a particular individual as the best TDMA schedule according to his/her preferences, priorities and/or constraints.

## 7. CONCLUSIONS

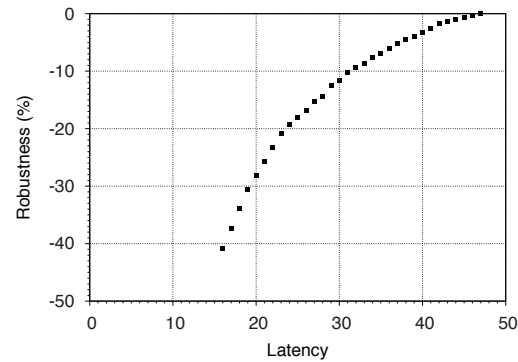
The proposed optimizer in Neuronal TDMA is designed to optimize TDMA signaling schedules for nanomachines in neuron-based BANNs. Simulation results demonstrate that the proposed optimizer efficiently obtains quality solutions and multiobjective analysis is critical in configuring neuron-based BANNs.

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(a) Hypervolume



(b) Trade-off between Latency and Robustness

Figure 9: Simulation Results

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