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## 2.1 Introduction

As previously mentioned, from the early days of the discipline, different approaches 6 have been followed to provide software tools assisting the P systems designers in 7 their design and verification tasks for a number of membrane system types and 8 variants. However, at the beginning, the most common case was the development 9 of specific-purpose tools devoted to the solution of a particular model based on 10 a P system or P systems family. While these early initiatives constituted relevant 11 achievements for membrane computing, their usefulness for the general community 12 was significant mostly in the context of the specific paper or scientific result they 13 were developed around. Surveys on the first generation of software tools related to 14 membrane computing can be found in [5, 49].

In order to move a step forward in this sense, aiming to provide some solution 16 for the P systems community in the form of a set of general tools for the software 17 implementation of P systems, P-Lingua framework emerged more than a decade 18 ago. As a first crucial element in the framework, a specification language, the socalled P-Lingua language, was defined, aiming to be a standard for the community 20 to speak the same language when defining P systems. One of the advantages of 21 having such a standard is to avoid ambiguities, and moreover to foster collaboration, 22 facilitating that researchers share their designs, even if they use different simulation 23 software—similarly as the Systems Biology Markup Language (SBML) format 24 works for the systems biology community. 25

The language started with some very general elements common to most P system <sup>26</sup> types, such as the membrane structure, objects, membrane labels, or rewriting rules. <sup>27</sup> Along with such general elements, each P system type or variant would admit <sup>28</sup> specific rules, and the framework would provide parsing tools to detect syntactic <sup>29</sup> or semantic errors. Along with the specification language, P-Lingua framework <sup>30</sup> provided from the beginning a number of built-in simulators, capturing the semantic <sup>31</sup> and dynamic aspects of each P system type. Such simulators were included for the <sup>32</sup> sake of completeness of the tool, but they were not intended to compete against 33 existing software. Actually, the framework included the functionality to compile P-34 Lingua code into something else so that one could provide such compiled result as 35 an input for an external simulator. Software implementation of P systems is further 36 explored in Chap. 3. 37

The chapter elaborates on some of the main capabilities of the framework and 38 is structured as follows. The main elements involved in P-Lingua language will be 39 introduced in Sect. 2.2, along with a classification of the main types and variants of 40 P systems supported by the framework, including the main references related with 41 them. Then, several simulation algorithms will be presented in Sect. 2.3, capturing 42 the dynamics of some especially relevant types of P systems used in the solution 43 of real-life problems. Finally, in Sect. 2.4, a higher-level tool will be presented, 44 MeCoSim, as a step forward to provide a visual virtual research environment. 45

#### 2.2 P-Lingua Language

P-Lingua is a domain-specific language started in 2008 [4] that has been continu- 47 ously evolving since then (technical details on the foundations can be found in [36] 48 and a survey together with some recent developments in [40]). The approach is to 49 keep the definitions as simple as possible, being a sort of "LaTeX-like" pseudocode, 50 in such a way that P systems designers can use similar notation to the one used in 51 the literature. A P system can thus be defined in a (plain text) .pli file, where the 52 designer indicates the model, structure, initial multisets, variables (if any), etc. The 53 elements of the definition will be further explained in what follows. 54

#### 2.2.1 P System Models

When designing a membrane system in P-Lingua, the instruction @model must 56 be present at the beginning of the .pli file, followed by a keyword identifying 57 the model used. Through the development of P-Lingua, several classes of P 58 systems have been included within the framework, while others have been discarded 59 due to the lack of their use. The latest stable version, pLinguaCore 4.0, was 60 released in 2013, covering only 10 model types. The P-Lingua framework has been 61 continuously expanding since then although the development efforts have been 62 focusing in the core distributed within MeCoSim. The following tables illustrate 63 the diversity of variants considered in the current version, with the corresponding 64 keywords and a reference introducing the model. For more details about the exact 65 pLinguaCore release where some models were included or discarded, we refer the 66 reader to [39]. 67

In the case of neural-like P systems, the model keyword spiking psystems 68 is slightly overloaded since it covers multiple subclasses. Each time the model has 69 been extended, special symbols and tokens were used so that the parser and the 70

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simulator are capable to identify which type of rules are being used and how they 71 should be interpreted. 72

#### 2.2.2 Membrane Structure

The topology of the membrane system to be simulated will depend on the model 74 selected in the file. If an invalid structure with respect to the model is defined, the 75 *parser* will show a message notifying it. The instruction @mu<sup>1</sup> is used to define the 76 architecture of the system. The syntax is similar to the one used in the literature. For 77 example, for cell-like membrane systems, the definition 78

would lead to a P system with a skin membrane labelled by 1 and 2 internal <sup>80</sup> membranes: an elementary membrane labelled by 3, and a membrane labelled by <sup>81</sup> 2 which contains an elementary membrane labelled by 4 inside. <sup>82</sup>

In the case of tissue P systems, membranes (called *cells*) are not hierarchically <sup>83</sup> arranged, but they can be connected by means of an arbitrary graph, which is not <sup>84</sup> required to be explicitly given in the definition. Typically, the set of directed arcs <sup>85</sup> connecting cells can be reconstructed from the implicit information provided by <sup>86</sup> the set of rules. However, in P-Lingua format, it is necessary to indicate the initial <sup>87</sup> cells in the system, formally considering them as elementary membranes located <sup>88</sup> within an external compartment labelled by 0, that will act as the environment of <sup>89</sup> the system. <sup>90</sup>

Spiking neural P systems need both the initial neurons and the *synapses* defined 91 in order to work. The former is defined as previously with the @mu instruction and 92 the later with @marcs indicating with pairs of labels which arcs will be present in 93 the underlying graph of the SNP system. 94

Note that not all the definitions of @mu must be in the same line, but instead 95 of the = symbol, it is possible to add more compartments to a specific region. For 96 instance, in cell-like membrane systems, it is possible to use 97

$$@mu = [[]'2]'1; @mu(1) + = []'3;$$
 98

in order to generate a structure identical to the one defined above. Note that a 99 semicolon indicates the end of an instruction.

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<sup>&</sup>lt;sup>1</sup>The usual notation for the structure of P systems is the Greek letter  $\mu$ .

#### 2.2.3 Initial Multisets

In order to describe the initial multisets of the different compartments, the command 102 @ms is used in a similar way to the literature, with braces {} as the delimiters of 103 the multisets. Like before, the + = symbols can be used to add new objects to a 104 predefined multiset. The multiplicity of a symbol is indicated by the \* symbol as in 105 a multiplication (e.g., c \* 5 indicates 5 copies of object c). 106

#### 2.2.4 P System Rules

Rules, like the structure of the P system, depend on the model of membrane system 108 being simulated. The P-Lingua parser was defined in such a way that P systems 109 researchers can use a very close language to the one used in literature, putting special 110 emphasis in the definition of rules. Therefore, brackets [] are used in P-Lingua 111 files as in the definition of rules in research papers. An evolution rule is defined 112 in the following way: + [ a1 - -> b, c ] ' 1. Note the differences between the 113 P-Lingua and the formal definitions: The subscripts are between braces, the arrow 114 is replaced by an ASCII version of an arrow, the label is preceded by a ' symbol 115 instead of being a subscript, and the polarization precedes the rule instead of being 116 a superscript. For tissue P systems, instead of using parentheses, a similar brackets 117 notation is used with a double arrow as follows: [a] ' 1 <- -> [b] ' 2 to denote 118 the rule (1, a/b, 2).

Usually, several rules with the same structure but with different subindexes 120 are defined in P systems, and it can be translated into a P-Lingua file with the 121 colon: symbol, followed by the corresponding limits. Let  $r \equiv [a_i \rightarrow a_{i+1}]_1$  122 for  $0 \leq i \leq n$  be a set of rules of a P system with active membranes 123 that can be defined in a P-Lingua file as follows:  $[a\{i\} - -> a\{i+1\}]_1$  124  $]'1 : 0 \leq i \leq n$ . If two or more variables have to be defined, they will 125 be declared from the right to the left; that is, if a variable j is limited by 126 i, then the range of i must be written "before" (to the right), for example, 127

 $[a\{i,j\} - -> a\{i,j+1\}]' 1 : 0 \le j \le i, 0 \le i \le n.$  128

The user must define all these parameters (except the model type) in a main 129 function. A function in the P-Lingua language is defined with the keyword def 130 followed by the name of the function. A function can have parameters whose names 131 will be indicated between parentheses and separated by commas. More than one 132 function can be defined in a single P-Lingua file, and they are widely used, for 133 instance, to construct the membrane system in a modular way. An example of this 134 would be the following code: 135

```
@model<membrane_division>
```

```
n = 3 /* A parameter n is defined to be used later */ m = 1000 /* A parameter m is defined to be used later */
```

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```
def main() {
    define_structure();
    define_initial_multisets(m);
    define_rules(n);
}
def define_structure() {
    @mu = []'1;
}
define_initial_multisets(number_objects) {
    @ms(1) += a{0}*{number_objects};
}
define_rules(number_steps) {
    [a{i} - -> a{i+1}]'1 : 0 <= i < {number_steps};
}</pre>
```

Note that it is allowed to insert comments in P-Lingua files, surrounded by the 156 symbols /\* and \*/. As indicated above, several examples of the different types of 157 P systems implemented in P-Lingua can be found in the websites of the P-Lingua 158 project [43] and MeCoSim [24].

### 2.3 Simulation Algorithms

P systems are bioinspired devices that work in a massively parallel and nondeterministic way. While there are preliminary studies analyzing the problems related to implementations in biological means, there is still a long way to reach this ultimate goal. That is why developing hardware/software implementations of P systems becomes a vital necessity for the advancement of scientific activities in membrane computing.

The P-Lingua framework includes a Java library called *pLinguaCore* that 167 provides at least one simulation algorithm for each P system variant. A simulation 168 algorithm for membrane computing can be described as an algorithm which 169 is able to reproduce P system computations on conventional software/hardware 170 architectures. Usually, only one branch of computation is considered, and it is 171 expected to display the sequence of configurations, including information on the 172 executed rules for each step of computation. Concerning the hardware used, the 173 simulation algorithms can be designed to run on sequential machines (singlethread CPU) or parallel architectures (multi-thread CPU, GPU, FPGA, etc.). The 175 simulation algorithms in pLinguaCore are designed for single-thread CPU, but it is possible to parse a P-Lingua file and compile it into an appropriate input for an external simulator.

All simulation algorithms in pLinguaCore share the same underlying implementation of a computation step as a loop divided into two stages: selection stage 180 and execution stage. The selection stage consists in searching for applicable rules 181 and selecting which ones will actually be executed in each membrane of a given 182 configuration, taking into account the restrictions dictated by the system semantics. 183

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Then, the execution stage actually implements the changes on the configuration 184 caused by the execution of the selected rules, and this completes the simulation of 185 the computation step. The input data for the selection stage contains the description 186 of the membranes with their multisets (strings over the working alphabet of objects, 187 labels associated with the membrane, etc.) and the set of defined rules. The output 188 data of this stage are the multisets of selected rules. Only the execution stage 190 needs synchronization when accessing to the membrane structure and the multisets. 191 At the end of the execution stage, the simulation process restarts the selection stage 192 in an iterative way until a halting configuration is reached (i.e., none of the rules 193 is applicable). Alternatively, a maximum number of iterations can be set at the 194 beginning of the simulation to avoid getting stuck on too long (or even infinite) 195 computations.

With the general design explained above, the pLinguaCore library includes <sup>197</sup> simulation algorithms for the cell-like, tissue-like, and neural-like P systems <sup>198</sup> enumerated in Sect. 2.2. For more information, see the corresponding references <sup>199</sup> in Tables 2.1, 2.2, and 2.3, respectively. There exist in the literature other P system <sup>200</sup> variants whose computations are not synchronized by a global clock in a step-bystep fashion (e.g., asynchronous, time-free, or stochastic models). Such variants are not currently supported under the P-Lingua framework, but there exist fully <sup>203</sup> functional alternative implementations available (see Chaps. 4 and 5). <sup>204</sup>

Other variants are also contemplated [2, 8, 10, 11]. A special mention should 205 be given to the simulation algorithms for population dynamics P systems (PDP 206 systems) which is a variant widely used for simulation of ecosystem dynamics 207 (see Chap. 6) in which each rule has a probability associated. The first description 208 of probabilistic semantics was quite ambiguous: "Rules should be applied in a 209 maximally parallel way, according to their probabilities." There are many ways of 210 interpreting this sentence, and each one could lead to different behaviors. While all 211 of them might be "correct" from a formal point of view, not all simulation algorithms 212 are acceptable when the goal is to reproduce the behavior of a complex system. 213 Since P-Lingua is a general-purpose framework, indicating which is the appropriate 214 choice should be a decision of the model designer (Table 2.4).

Three simulation algorithms have been designed for PDP systems and implemented in pLinguaCore [19]: 217

DNDP algorithm [21].	218
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BBB algorithm [19]. 219
 DCBA algorithm [22]. 220

In the algorithm DNDP, the rules are selected individually according to its 221 probabilities. On the other hand, algorithms BBB and DCBA work by grouping 222 rules in blocks by analyzing the left-hand side, each block has the same left-hand 223 side, and all the rule probabilities must sum 1. DCBA uses a refined definition of 224 block in which the charges of the right-hand side must be consistent. More about 225 simulation algorithms for PDP systems will be explained in Chap. 6 since this kind 226

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Variant of membrane systems	Model specification keyword	Ref.	t5.1
P systems with active membranes and membrane creation	membrane_creation	[25]	t5.2
P systems with active membranes and membrane division	membrane_division/dam	[33]	t5.3
P systems with symport/antiport rules	<pre>symport_antiport / infEnv_symport_antiport</pre>	[18]	t5.4
Polarizationless P systems with active membranes with minimal cooperation and membrane division	dam_wp	[48]	t5.5
Polarizationless P systems with active membranes with minimal cooperation, membrane division, and without dissolution	dam_wp_wd	[48]	t5.6
Polarizationless P systems with active membranes with minimal cooperation and membrane division only for elementary membranes and without dissolution	dam_wp_wd_wn	[48]	t5.7
P systems with active membranes with minimal cooperation and membrane separation	sam	[47]	t5.8
Polarizationless P systems with active membranes with minimal cooperation and membrane separation	sam_wp	[47]	t5.9
Polarizationless P systems with active membranes with minimal cooperation and membrane separation and without dissolution	sam_wp_wd	[47]	t5.10
Polarizationless P systems with active membranes with minimal cooperation and membrane separation only for elementary membranes and without dissolution	sam_wp_wd_wn	[47]	t5.1′
Transition P systems	transition/rewriting	[32]	t5.12

 Table 2.1
 Cell-like membrane systems implemented in P-Lingua

Table 2.2	Tissue-like membrane systems implemented in P-Lingua

Variant of membrane systems	Model specification keyword	Ref.	t8.
Tissue P systems with cell division	tissue_psystems/tpdc	[20, 34]	t8.2
Tissue P systems with cell division and antiport rules	tpda	[34]	t8.3
Tissue P systems with cell division and symport rules	tpds	[34]	
Tissue P systems with cell separation	TSCS	[27, 38]	t8.5
Tissue P systems with evolutional communication rules with cell division	evolution_communication/ ev_symport_antiport	[31]	
Tissue P systems with evolutional communication rules with cell separation	tsec	[31]	
Tissue P systems with promoters	tpdc/tpda/tpds	[45]	t8.8

			_
Variant of membrane systems	Model specification keywo	rd Ref.	t1
Asynchronous SN P systems	spiking_psystems	[3]	t1
Asynchronous SN P systems with local synchronization	spiking_psystems	[41]	t1
Cell-like spiking neural P systems	cell_like_snp	[46, 50]	t1
Dendrite P systems	dendrite	[26]	t1
Fuzzy reasoning spiking neural P systems	fuzzy_psystems	[13, 17]	t1
Limited asynchronous SN P systems	spiking_psystems	[29]	t1
Spiking neural P systems	spiking_psystems	[13, 15, 16]	- t1
Spiking neural P systems with anti-spikes	spiking_psystems	[28]	t1
Spiking neural P systems with hybrid astrocytes	spiking_psystems	[30]	- t1
Spiking neural P systems with structural plasticity	spiking_psystems	[1]	t1

Table 2.3 Neural-like membrane systems implemented in P-Lingua

Table 2.4 Other variants of membrane systems implemented in P-Lingua

Model specification keyword	Ref.	
enps	[35]	
probabilistic	[2]	
probabilistic_guarded_	[8, 10]	
_psystems		
regenerative_psystems	[11]	
simple_kernel_psystems	[12, 14]	
simple_regenerative_	[11]	
_psystems		
stochastic	[42]	
	Model specification keyword enps probabilistic probabilistic_guarded_ _psystems regenerative_psystems simple_kernel_psystems simple_regenerative_ _psystems stochastic	Model specification keywordRef.enps[35]probabilistic[2]probabilistic_guarded_[8,10]_psystems[11]regenerative_psystems[11]simple_kernel_psystems[12,14]simple_regenerative_[11]_psystemsstochastic

of algorithms requires a large amount of computational power being suitable for <sup>227</sup> high-performance computing platforms such as CUDA. <sup>228</sup>

As it was mentioned before, there are various approaches in the literature <sup>229</sup> where the standard semantics of P systems (namely, nondeterministic behavior <sup>230</sup> and maximally parallel application of the rules) is modified by adding different <sup>231</sup> regulation elements, which need to be carefully described in order to explain how the <sup>232</sup> system evolves. In particular, it is worth highlighting that the concept of "simulation <sup>233</sup> algorithm" is used in this section in a theoretical sense, that is, a formalization that <sup>234</sup> translates the specification of the semantics into a pseudocode capturing precisely <sup>235</sup> the routine that the system follows when deciding what rules to apply. It should not <sup>236</sup> be confused with an implementation of such algorithm in a programming language. <sup>237</sup> Some attempts trying to bring semantic elements explicitly into the description <sup>238</sup> of a P system in P-Lingua language have been already initiated, and it is being <sup>239</sup> considered for upcoming release of P-Lingua 5.0. <sup>240</sup>

### 2.4 Membrane Computing Simulator (MeCoSim)

The previous sections have presented the essential elements of P-Lingua framework: 242 the standard specification language and the simulation engines to run the computa-243 tions of the given P systems. These elements constitute the core features, the chassis 244 of our car. However, in order for this vehicle to move smoothly, several additional 245 pieces (as the external body but also others as the steering wheel, the pedals, or the 246 dashboard icons, among others) are needed to provide the users with the desired 247 driving experience, in order for them to sit down and enjoy while conducting their virtual experiments with models based on P systems. 249

With the metaphor introduced, we aim to present the main idea behind MeCoSim 250 (membrane computing simulator) [24, 37], conceived in a search for the gener-251 alization of certain high-level visual applications to manage population dynamics 252 models, known as EcoSim product family [36, 44]. Built on top of P-Lingua core, 253 this visual environment provides a higher level of abstraction, transforming the 254 solid set of tools of our internal chassis and engine given by P-Lingua core into 255 a whole car, complementing the previous elements with an external layer allowing 256 the drivers conduct their experiments through the proper sensors and actuators. 257

Thus, MeCoSim was devised with a manifold purpose, assisting in the design 258 of the heart of the cars (P system-based models), delivering the final cars (custom 259 apps based on the models), and helping users drive their vehicles (through the 260 tools coming with the apps). Firstly, the visual interface provides the expert users 261 (P system designers) with an interface where they can specify, debug, and run 262 (step-by-step or entire) computations of their P systems in a smoother way; this 263 is made easier with the tools provided by a friendly environment, aiding in the 264 task of designing and verifying the core part of our cars: the P systems modelling 265 certain case studies. Secondly, the environment provides certain tools to make the 266 technical pieces constituting the model become a final product, that is, bridging 267 the gaps to convert the engine, car axles, controls, or wheels into the final car. To 268 this purpose, MeCoSim provides some tools to define, through configuration files, a 269 final visual application using the core elements of the framework, plus the P system 270 (or P system family) specified, and the inputs and outputs to control and monitor, 271 respectively, each trip made with the car (i.e., each computation of the system, each 272 virtual experiment conducted). Finally, end users receive their car: the customized 273 application satisfying their needs. Probably, they will have no idea about the internal 274 specifications of the car, they are not car mechanics/technicians, but they will be 275 able to drive their specific car. In such car, the custom app, they will be able to sit, 276 introduce the details about each particular trip (experiment) they want to make (run), 277 decide about the speed, and control the steering wheel and pedals, enjoying the drive 278 and finally getting to their destiny (the end of the computation) while obtaining all 279 the desired additional information through the monitoring system provided by the 280 dashboard. 281

In this context, everything starts with the identification of a certain need, such 282 as solving a certain NP-complete problem or modelling a real-life system in 283

economy, ecology, medicine, or any other field. In this context, a P system (or P <sup>284</sup> system family) must be defined to satisfy such need. Thus, the definition of the <sup>285</sup> system requires the translation of the P system into a file using P-Lingua standard <sup>286</sup> specification language. Then, an iterative process of design, debug, and verification <sup>287</sup> starts, progressing with the problem until the model has been properly validated <sup>288</sup> according to the experts in the problem domain. <sup>289</sup>

Then, the central problem has been solved, but only the technicians could use the 290 tools to run computations of the system. Then, a new effort can be made by such 291 P system designers to set in a spreadsheet configuration file the specific elements 292 of the final car and the application where the end users (ecologists, economists, 293 etc.) will be able to conduct their visual experiments. The custom elements will 294 include the hierarchical structure arranging all the visual blocks of the final app, 295 the tables allowing the introduction of specific input data by the end user, and the 296 outputs to monitor the activity and the final results of the trip. Now, everything 297 is ready for the end user to drive, to analyze each particular scenario of interest 298 (each trip), introducing in the tables the specific parameter values and input data 299 for each scenario of interest and run each virtual experiments, getting the desired 300 results in their dashboard given by the custom output tables and charts defined in 301 the configuration file set by the P system designers during the building of the car 302 (the custom app).

The description above has been probably illustrative at a general level, but people 304 not familiarized with MeCoSim might find it difficult to figure out how this approach 305 look like at a deeper level. The following subsections will try to clarify those aspects 306 only outlined before, detailing the main goals achieved (Sect. 2.4.1) and the software 307 components involved (Sect. 2.4.2). In Ref. [44], a methodology is proposed based 308 for the solution of a problem through membrane-based systems making use of 309 P-Lingua framework and MeCoSim, where the corresponding tools described are employed in a systematic way. 311

#### 2.4.1 Primary goals

As it has just been depicted, MeCoSim's main intent is the provision of a high-level 313 visual interface to handle P system-based models. This is right, but what should we 314 exactly expect from this environment? Let us try to clarify this by analyzing the 315 origin and initial view of the tools involved. 316

To start with this overview, it is worth recalling that we are studying a paradigm, 317 membrane computing, where many computing models have been defined along 318 the years. As specifically addressed by this book, the implementation of these 319 computational devices is crucial in order to take advantage of all the theoretical 320 properties, the strengths, of such machines. However, our biologically inspired 321 models present certain features that are not easy to implement in certain biological or artificial substrate, and even if it can be done, it implies major efforts to apply 323 these machines to each particular problem. Nevertheless, there is a faster convenient 324 approach that can be applied in order to make this process more manageable in 325

(a) the study of theoretical aspects of the different types of devices (such as their 326 computational power, efficiency, etc.) or the practical use of models based on these 327 devices (its tasks such as the design, verification, or validation of properties and 328 virtual experimentation) and (b) the simulation of the computations for the given 329 models. 330

This volume is devoted to implementations of membrane computing models. <sup>331</sup> Consequently, we actually expect real devices that can capture all the features of the theoretical machines. Some of the solutions provided by later chapters will succeed, <sup>333</sup> addressing aspects such as capturing the inherent parallelism of such ideal machines. <sup>334</sup> However, those real machines will need to take many things into account at a very <sup>335</sup> technical level, it will be a very tough process, and this will make it very challenging <sup>336</sup> to validate the proper functioning of these devices according to all the properties of <sup>337</sup> the types of P systems used, along with all the properties of the models built for a <sup>338</sup> particular problem, based on such types of P systems. <sup>339</sup>

In order to avoid attacking all the problems at once, a different approach can 340 be followed: first facing the "soft" implementation of the theoretical devices, the 341 intended type of P systems, through sequential software simulators, not addressing 342 all the technical aspects required by the actual implementations but properly 343 simulating the computations of the theoretical systems, conducting to the very 344 same results. This allows us to validate a simpler implementation at a functional 345 level, in terms of the results of the computations, permitting a first step toward 346 more complex high-performance hardware, hybrid, or biological implementations. 347 These initial simulators could handle any solution or model solving a certain 348 problem (from SAT, 3-COL, or HAM-CYCLE to the population dynamics of an 349 ecosystem or an economy system, among others) by means of the types of P systems 350 implemented in the corresponding simulators. Such handling will involve helping 351 in the design, debug, and verification tasks, but of course also the computation of 352 the given model or solution according to the semantic and dynamic rules of the 353 theoretical devices. Naturally, for any problem of certain size, analyzing a manual 354 trace of the computation in a paper would be too tedious or practically unfeasible 355 for significantly big instances. Therefore, even if a real implementation with the 356 desirable parallelism is not available, it would be necessary to have at disposal a 357 machine where one could simulate computations to validate the model or analyze 358 the evolution of the system under certain scenarios. That would be the approach 359 followed by P-Lingua framework and MeCoSim so that we can focus on aspects 360 such as reliability or feasibility (to preserve the same evolution and results of the 361 theoretical systems), along with user-friendliness, over the efficiency of other later 362 implementations. 363

We have clarified the first goal of MeCoSim approach: providing an environment 364 for the design, verification, and simulation of the models based on P systems in 365 a reliable and user-friendly way, albeit not prioritizing efficiency. However, there 366 are more aspects to analyze in our approach. A major one is the search for the 367 generalization, *that is*, a definite purpose of providing general-purpose tools to be 368 applied to each particular membrane system type and each particular solution for a 369 problem based on them. Thus, the development of *ad hoc* simulators (for a specific 370 solution for a problem, a single instance/scenario, or different instances of that 371 problem) is definitely different from the development of a general-purpose simulator 372 for certain types of P systems, capturing the ingredients of the theoretical model of 373 computation so that this machine allows the provision of any P system and scenario 374 and performs its corresponding intended computations. 375

Many software developments in scientific research are focused on the first 376 approach, providing simulators for certain problems or even specific instances of 377 the problem only. Other studies address the development of tools for the analysis 378 or verification of a specific type or variant of membrane system, such that they can 379 handle any P system of the type. However, the approach of P-Lingua framework 380 and MeCoSim has been more ambitious from its origin: providing tools being as 381 general as possible, for as many types of P systems as possible (including many 382 variants of cell-like, tissue-like, and neuron-like P systems, among others), while 383 preserving the strict deep analysis of the syntactic, semantic, and dynamic aspects 384 of each P system variant, in order to control that the corresponding constraints are 385 met. The wide range of variants covered include computing models with different 386 global structures (hierarchical, plain graph, or graph with nodes containing trees— 387 as in multienvironment, PDP systems), a variety of ingredients in terms of rules or 388 other elements (dissolution, division, charges, stopping objects, etc.), and different 389 handling of semantic aspects (related with sequentiality, nondeterminism, priorities, 390 maximality, probabilistic or stochastic behaviors, among others). 391

As described above and in Sects. 2.2 and 2.3, many types and variants of 392 P systems have been covered by the tools developed within the framework. 393 Moreover, as detailed in Sect. 2.2, along with the specification language and its 394 corresponding parsers for each computing models, many simulators were developed 395 inside P-Lingua project. This infrastructure provided from the beginning [36] 396 a complete programming environment for *membrane computing* and has kept 397 incorporating new elements along the years, staying as an alive project, including a 398 living version inside MeCoSim. 399

In Sect. 3.4, the basic steps of the approach followed with these tools are 400 depicted, illustrating their use for real applications. 401

#### 2.4.2 Main Functional Components

As previously mentioned, a clear separation of the roles involved in modeling and 403 simulation process is stated (apart from the software developers in charge of P- 404 Lingua and MeCoSim development): (a) P systems designer and (b) end users of a 405 simulation app. What does MeCoSim provide within this scope? 406

As shown in Fig. 2.1, the software developer releases different versions of 407 MeCoSim (and certain plug-ins) available for any potential users. In contrast, 408 P systems designer, possibly unrelated with software development, defines a 409 simulation app based on MeCoSim, customized for its particular problem. Then, 410 he can debug its solution and analyze the underlying P system. Finally, the end user 411



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Pandemic	- 🗆 X	Pandemic	- 0
enario Edit Model Simulation Plugins Help		Scenario Edit Model Simulation Plugins Help	
Input Output Debug console		Input Output Debug console	
ParsingInfo SimulationInfo Errors Warnings		ParsingInfo SimulationInfo Errors Warnings	
$ \begin{array}{l} \label{eq:result} Inter. Adding rule #101799 N[25,51]- III \rightarrow IA2(25,21, A(25,720), X) \\ Inter. Adding rule #101709 N[25,51]- III \rightarrow IA2(3,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,51]- III \rightarrow IA2(3,52), A(35,720), X) \\ Inter. Adding rule #101781 N[35,51]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,51]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,51]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,11]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,11]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[35,11]- III \rightarrow IA2(5,52), A(25,720), X) \\ Inter. Adding rule #101781 N[15,11]- III \rightarrow IA2(5,52), A(15,570), X) \\ Inter. Adding rule #101781 N[15,11]- III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,11]- III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #1017781 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,22), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,52), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,52), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,52), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+ III \rightarrow IA2(15,52), A(15,570), X) \\ Inter. Adding rule #101778 N[15,51]+$	M(= 0.035 M(= 0.035) M(= 0.035 M(= 0.035 M(= 0.035) M(= 0.035 M(= 0.035) M(= 0.035 M(= 0.035) M(=	1 * 415 [023,30] +(1] → X3(3,32] \$5(3)?20[1 = 0.075 1 * 4800 [0213,30] +(0] → X4[13,31] XM, 41(3)?20[1 = 0. 1 * 6800 [0213,01] +(0] → X4[13,31] XM, 41(3)?20[1 = 0. 1 * 6910 [0213,01] +(0] → X4[13,31] XM, 41(3)?20[1 = 0. 1 * 6910 [013,01] +(0] → X4[13,1] XM, 41(3)?20[1 = 0. Fulles selected for MEXMBRANE ID: 6, Labelt 1,102, Charge 0 1 * 4112 [R[0] → R[1][1 = 1.0 CONFIGURATION: 1 MEMBRANE ID: 3, Labelt 1,101, Charge 0 <	275 275 213 213 213 213 213 213 213 213 213 213
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Fig. 2.2 Model debugging

employs the custom simulation app to study different scenarios of interest involving	412
specific instances of the problem.	413
In summary, the main functionalities of MeCoSim are the following:	414

In summary, the main functionalities of MeCoSim are the following:

General environment to simulate computations of P systems

With the default custom application, any P-Lingua file not requiring additional 416 inputs can be edited, while detecting aspects to modify; parsed and debugged (see 417 Fig. 2.2), to find possible warnings or errors, both at a syntactic and a semantic 418 level, alerting the P systems designer if some rules of the intended model type 419 are violated; and simulated (through the algorithm selected in the interface, as 420 shown in Fig. 2.3), generating the initial structure and multisets of the system 421 and then running the computation either step-by-step or until its end (after a fixed 422 number of steps or when a halting configuration is reached, where no rules can 423 be applied). Besides, the default output is given in the form of a flat table (with a 424 row for each object symbol present in each computation step inside each region, 425 with a certain multiplicity), and also some of the main internal elements of the 426 P system can be visualized at any moment (membrane structure, multisets, and 427 alphabet). 428

- Mechanism for the definition of custom simulation apps Any custom app consists of:
  - A hierarchical structure for the visual arrangement of the information (inputs 431 and outputs) in the app for the end user, according to the setting introduced 432 by the designer in an .xls spreadsheet file, as illustrated by the first table in 433 Fig. 2.4. 434
  - A definition of input tables, and output tables and charts, to respectively 435 introduce data and visualize results. More details about the definition of such 436

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Fig. 2.3 Available simulators for a loaded P systems variant

		Tab Id	Ta	ib Name		Tal	Parent	ld		
			1 Tis	sue Exa	mple			0		
			2 Inp	out				1		
			3 Si	ze (n)				2		
Table Id		Table Name		Tab Id	Columns	Init Rows	Save T	o File	Input /	Output
1		Size (n)		3	1	1	TRI	JE	Inp	out
Colu	mn Id	Column Nam	e l	Default V	/alue E	ditable	Tooltip	Grap	nicRole	
	1	Size	3		TRU	E	n			
			Par	am Nam	e Param \	/alue				
			n		<5,1,1>					

Fig. 2.4 Custom app definition—input

components can be found at Refs. [24, 37, 44], and a basic example of tables 437 (header and columns) configuration is given by the second and third tables 438 present in Fig. 2.4.

The configuration of parameters, establishing which parameters and input data 440 for the model, should be generated from the input tables, either directly taking 441 the value from the table at the beginning of the simulation or applying some 442 processing from the input data to generate calculated derived values. To this 443 purpose, a specific parameters generation language was defined, as described 444 in detail in [37, 44]. The generation of a basic parameter *n* from the first row 445

and column of a table with id = 5 is illustrated in the last table of Fig. 2.4. 446 Much more complex parameters can be generated, as described in [37,44]. 447

Results: In order for the output tables and charts to show some information 448 about the simulation, the custom configuration must define which elements 449 from the computation should be taken into account when extracting informa- 450 tion from all the computation trace data. An additional language is used in the 451 *.xls* spreadsheet file to provide a flexible mechanism to express the retrieval of 452 information from the computation. Internally, for every simulation performed, 453 from the previous definition, a database query is generated, being executed 454 against the given *on-memory* database containing the flat structure with the 455 computation.

Apart from this core functionalities, additional features and abilities are provided 457 in the form of MeCoSim plug-ins following a certain architecture proposed [24, 37, 458 44]. These *add-ons* can be given either as Java-based packages (as a graph viewer 459 [see Fig. 2.5], a window for the introduction and encoding of logical formulas or a 460 tool to define and detect invariants in the models based on Daikon [7]) or as external 461 programs being called from MeCoSim and properly connected (using the so-called 462 *processes plug-in*). A detailed description of the underlying mechanisms and the 463 plug-ins developed is given in [44].

Additionally to the features of MeCoSim software and its plug-ins, a system of 465 repositories was made available, manageable from MeCoSim environment, having 466 access to repositories of four types: apps (*.xls*), models (*.pli*), scenarios (*.ec2*), and 467 plug-ins (*.jar*). Besides the official repositories, any user can provide additional 468 ones (through the definition of the corresponding *.xml* file for the desired type of 469 repository), providing the corresponding URL to the resource.



Fig. 2.5 GraphsPlugin-trees of graphs

# 2.5 Conclusion

This chapter presented the very representative and widely used P-Lingua language 472 for a variety of P systems such as cell-like P systems, tissue-like P systems, spiking 473 neural P systems, and kernel P systems. The description of P-Lingua language with 474 pLinguaCore consists of P systems models, membrane structure, initial multisets, 475 and P system rules. The simulation algorithms in P-Lingua and MeCoSim on top 476 of pLinguaCore with primary goals and main functional components were also 477 discussed. 478

### References

- F.G.C. Cabarle, H.N. Adorna, N. Ibo, Spiking neural P systems with structural plasticity, in 480 Pre-proceedings of 2nd Asian Conference on Membrane Computing, Chengdu, China (2013), 481 pp. 13–26
   M.A. Cohene, A. Manulida, A. Dalan, L. Dúna, L. Dúna, M.J. Dúna, J. Dúna, J.
- M. Cardona, M.A. Colomer, A. Margalida, A. Palau, I. Pérez-Hurtado, M.J. Pérez-Jiménez, 483 D. Sanuy, A computational modeling for real ecosystems based on P systems. Nat. Comput. 484 10(1), 39–53 (2011). https://doi.org/10.1007/s11047-010-9191-3
- M. Cavaliere, O. Egecioglu, O.H. Ibarra, M. Ionescu, Gh. Păun, S. Woodworth, Asynchronous 486 spiking neural P systems: decidability and undecidability, in *DNA Computing. Lecture Notes* 487 *in Computer Science*, ed. by M. Garzon, H. Yan, vol. 4848 (2008), 246–255. https://doi.org/10. 488 1007/978-3-540-77962-9\_26 489
- D. Díaz-Pernil, I. Pérez-Hurtado, M.J. Pérez-Jiménez, A. Riscos-Núñez, P-Lingua: a programming language for Membrane Computing, in *Proceedings of the Sixth Brainstorming Week on Membrane Computing*, Fénix Editora, D. Díaz-Pernil, C. Graciani, M.A. Gutiérrez-Naranjo, Gh. Păun, I. Pérez-Hurtado, A. Riscos-Núñez (2008), pp. 135–155
- D. Díaz-Pernil, C. Graciani, M.A. Gutiérrez-Naranjo, I. Pérez-Hurtado, M.J. Pérez-Jiménez, 494
   Software for P systems, in *The Oxford Handbook of Membrane Computing*, ed. by Gh. Păun, 495
   G. Rozenberg, A. Salomaa (Oxford University, Oxford, 2009), pp. 437–454. Chapter 17 496
- D. Díaz-Pernil, I. Pérez-Hurtado, M.J. Pérez-Jiménez, A. Riscos-Núñez, A P-Lingua programming environment for Membrane Computing, in *Membrane Computing (WMC 2008)*, ed. by
   D.W. Corne, P. Frisco, Gh. Păun, G. Rozenberg, A. Salomaa. Lecture Notes in Computer
   Science, vol. 5391 (2009), pp. 187–203. https://doi.org/10.1007/978-3-540-95885-7\_14
- M.D. Ernst, J.H. Perkins, P.J. Guo, S. McCamant, C. Pacheco, M.S. Tschantz, C. Xiao, The 501 Daikon system for dynamic detection of likely invariants. Sci. Comput. Program. 69(1–3), 35– 45 (2007). https://doi.org/10.1016/j.scico.2007.01.015
- M. García-Quismondo, Modelling and Simulation of Real-life Phenomena in Membrane 504 Computing. Ph.D. Thesis (Universidad de Sevilla, Sevilla, 2014). http://hdl.handle.net/11441/ 506
- M. García-Quismondo, R. Gutiérrez-Escudero, I. Pérez-Hurtado, M.J. Pérez-Jiménez, A. 507 Riscos-Núñez, An overview of P-Lingua 2.0, in *Membrane Computing. WMC 2009. Lecture* 508 *Notes in Computer Science*, vol. 5957, ed. by Gh. Păun, M.J. Pérez-Jiménez, A. Riscos, G. 509 Rozenberg, A. Salomaa (2010), pp. 264–288. https://doi.org/10.1007/978-3-642-11467-0\_20 510
- M. García-Quismondo, M.A. Martínez-del-Amor, M.J. Pérez-Jiménez, Probabilistic guarded P 511 systems: a formal definition, in *Proceedings of the Twelfth Brainstorming Week on Membrane* 512 *Computing*, Fénix Editora, ed. by L.F. Macías-Ramos, M.A. Martínez-del-Amor, Gh. Păun, A. Riscos-Núñez, L. Valencia-Cabrera (2014), pp. 183–206 514
- M. García-Quismondo, M. Levin, D. Lobo, Modeling regenerative processes with membrane 515 computing. Inf. Sci. 381, 229–249 (2017). https://doi.org/10.1016/j.ins.2016.11.017 516

27

471

- M. Gheorghe, F. Ipate, R. Lefticaru, M.J. Pérez-Jiménez, A. Turcanu, L. Valencia, M. García-517 Quismondo, F. Mierla, 3-COL problem modelling using simple kernel P systems. Int. J. 518 Comput. Math. 90(4), 816–830 (2013). https://doi.org/10.1080/00207160.2012.743712 519
- M. Ionescu, Gh. Păun, T. Yokomori, Spiking Neural P systems. Fundam. Inform., 71(2–3), 520 279–308 (2006)
- F. Ipate, R. Lefticaru, L. Mierla, L. Valencia, H. Hang, G. Zhang, C. Dragomir, M.J. Pérez-Jiménez, M. Gheorghe, Kernel P systems: applications and implementations. Adv. Intell. Syst. Comput. 212, 1081–1089 (2013). https://doi.org/10.1007/978-3-642-37502-6\_126
- L.F. Macías-Ramos, *Developing Efficient Simulators for Cell Machines*. Ph.D. Thesis (Universidad de Sevilla, Seville, 2016). http://hdl.handle.net/11441/36828
- 16. L.F. Macías-Ramos, I. Pérez-Hurtado, M. García-Quismondo, L. Valencia-Cabrera, M.J. 527
   Pérez-Jiménez, A. Riscos-Núñez, A P-Lingua based simulator for Spiking Neural P systems, 528
   in *Membrane Computing (CMC 2011)*, ed. by M. Gheorghe, Gh. Păun, G. Rozenberg, A. 529
   Salomaa, S. Verlan. Lecture Notes in Computer Science, vol. 7184 (2012), pp. 257–281. https:// 530
   doi.org/10.1007/978-3-642-28024-5\_18
- 17. L.F. Macías-Ramos, M.A. Martínez-del-Amor, M.J. Pérez-Jiménez, Simulating FRSN P 532 systems with real numbers in P-Lingua on sequential and CUDA platforms, in *Membrane* 533 *Computing (CMC 2015)*, ed. by G. Rozenberg, A. Salomaa, J.M. Sempere, C. Zandron. Lecture 534 Notes in Computer Science, vol. 9504, pp. 262–276 (2015). https://doi.org/10.1007/978-3-536 319-28475-0\_18 536
- L.F. Macías-Ramos, L. Valencia-Cabrera, B. Song, T. Song, L. Pan, M.J. Pérez-Jiménez, A 537
   P-lingua based simulator for P systems with symport/antiport rules. Fundam. Inform. 139(2), 538
   211–227 (2015). https://doi.org/10.3233/FI-2015-1232
- M.A. Martínez-del-Amor, Accelerating Membrane Systems Simulators using High Performance Computing with GPU. Ph.D. Thesis (Universidad de Sevilla, Sevilla, 2013). http://hdl.
   handle.net/11441/15644
- M.A. Martínez-del-Amor, I. Pérez-Hurtado, M.J. Pérez-Jiménez, A. Riscos-Núñez, A P-Lingua based simulator for Tissue P systems. J. Logic Algebraic Program. **79**(6), 374–382 544 (2010). https://doi.org/10.1016/j.jlap.2010.03.009 545
- M.A. Martínez-del-Amor, I. Pérez-Hurtado, M.J. Pérez-Jiménez, A. Riscos-Núñez, M.A. 546 Colomer, A new simulation algorithm for multienvironment probabilistic P systems, in 2010 547 *IEEE Fifth International Conference on Bio-Inspired Computing: Theories and Applications* 548 (*BIC-TA*), Changsha, 2010, vol. 1 (2010), pp. 59–68. https://doi.org/10.1109/BICTA.2010. 549 5645352 550
- M.A. Martínez-del-Amor, I. Pérez-Hurtado, M. García-Quismondo, L.F. Macías-Ramos, L. 551 Valencia-Cabrera, A. Romero-Jiménez, C. Graciani-Díaz, A. Riscos-Núñez., M.A. Colomer, 552 M.J. Pérez-Jiménez, DCBA: simulating Population Dynamics P Systems with proportional 553 object distribution, in *Membrane Computing. CMC 2012*, ed. by E. Csuhaj-Varjú, M. Gheorghe, G. Rozenberg, A. Salomaa, G. Vaszil. Lecture Notes in Computer Science, vol. 7762 (2012), pp. 291–310. https://doi.org/10.1007/978-3-642-36751-9\_18
- M.A. Martínez-del-Amor, I. Pérez-Hurtado, M. García-Quismondo, L.F. Macías-Ramos, L. 557 Valencia-Cabrera, A. Romero-Jiménez, C. Graciani, A. Riscos-Núñez, M.A. Colomer, M.J. 558 Pérez-Jiménez, DCBA: simulating population dynamics P systems with proportional objects 559 distribution, in *Membrane Computing (CMC 2012)*, ed. by E. Csuhaj-Varjú, M. Gheorghe, G. 560 Rozenberg, A. Salomaa, G. Vaszil. Lecture Notes in Computer Science, vol. 7762 (2013), pp. 561 257–276. https://doi.org/10.1007/978-3-642-36751-9\_18 562
- 24. MeCoSim website. http://www.p-lingua.org/mecosim
- 25. M. Mutyam, K. Krithivasan, P systems with membrane creation: universality and efficiency, in 564 *Machines, Computations, and Universality (MCU 2001)*, ed. by M. Margenstern, Y. Rogozhin. 565 Lecture Notes in Computer Science, vol. 2055 (2001), pp. 276–287. https://doi.org/10.1007/3-566 540-45132-3\_19 567

563

AQ2

- 26. D. Orellana-Martín, M.A. Martínez-del-Amor, L. Valencia-Cabrera, I. Pérez-Hurtado, Agustín Riscos-Núñez, M.J. Pérez-Jiménez, Dendrite P Systems toolbox: representation, algorithms and simulators. Int. J. Neural Syst.. Available online 30 September 2020. https://doi.org/10.570 1142/S0129065720500719 571
- L. Pan, T.-O. Ishdorj, P systems with active membranes and separation rules. J. Universal 572 Comput. Sci. 10(5), 630–64 (2004). https://doi.org/10.3217/jucs-010-05-0630 573
- 28. L. Pan, Gh. Păun, Spiking neural P systems with anti-spikes. Int. J. Comput. Commun. Control 574
   4(3), 273–282 (2009). https://doi.org/10.15837/ijccc.2009.3.2435
   575
- 29. L. Pan, J. Wang, H.J. Hoogeboom, Limited asynchronous spiking neural P systems. Fundam. 576 Inform. 110(1–4), 271–293 (2011). https://doi.org/10.3233/FI-2011-543 577
- L. Pan, J. Wang, H.J. Hoogeboom, Asynchronous extended spiking neural Psystems with 578 astrocytes, in *Membrane Computing (CMC 2011)*, ed. by M. Gheorghe, Gh. Păun, G. 579 Rozenberg, A. Salomaa, S. Verlan. Lecture Notes in Computer Science, vol. 7184 (2012), pp. 243–256. https://doi.org/10.1007/978-3-642-28024-5\_17 581
- L. Pan, B. Song, L. Valencia-Cabrera, M.J. Pérez-Jiménez, The computational complexity 582 of tissue P systems with evolutional symport/antiport rules. Complexity 2018, Article ID 583 3745210, 21 (2018). https://doi.org/10.1155/2018/3745210 584
- 32. Gh. Păun, Computing with membranes. J. Comput. Syst. Sci. 61(1), 108–143 (2000). https:// 585 doi.org/10.1006/jcss.1999.1693. First circulated at TUCS Research Report No. 208, November 586 1998. http://www.tucs.fi
- 33. Gh. Păun, P systems with active membranes: attacking NP complete problems. J. Autom. Lang. 588
   Comb. 6(1), 75–90 (2000). Auckland University, CDMTCS Report No 102 (1999) 589
- Gh. Păun, M.J. Pérez-Jiménez, A. Riscos-Núñez, Tissue P systems with cell division. Int. J. 590 Comput. Commun. Control 3(3), 295–303 (2008). https://doi.org/10.15837/ijccc.2008.3.2397 591
- 35. A.B. Pavel, O. Arsene, C. Buiu, Enzymatic numerical P systems: a new class of Membrane 592
   Computing systems, in *Proceedings of the 2010 IEEE Fifth International Conference on Bio-*593
   *Inspired Computing: Theories and Applications (BIC-TA 2010), Changsha, China, September*594
   23–26 (2010), pp. 1331–1336. https://doi.org/10.1109/BICTA.2010.5645071
- 36. I. Pérez-Hurtado, Desarrollo y Aplicaciones de un Entorno de Programación para Computación Celular: P-Lingua. Ph.D. Thesis (Universidad de Sevilla, Sevilla, 2010, in Spanish).
   597 http://hdl.handle.net/11441/66241
   598
- 37. I. Pérez-Hurtado, L. Valencia-Cabrera, M.J. Pérez-Jiménez, M.A. Colomer, A. Riscos-Núñez, 599 MeCoSim: a general purpose software tool for simulating biological phenomena by means 600 of P systems, in *Proceedings of the IEEE Fifth International Conference on Bio-inspired* 601 *Computing: Theories and Applications (BIC-TA 2010)*, vol. I, ed. by K. Li, Z. Tang, R. Li, A.K. 602 Nagar, R. Thamburaj (2010), pp. 637–643. https://doi.org/10.1109/BICTA.2010.5645199 603
- 38. I. Pérez-Hurtado, L. Valencia-Cabrera, J.M. Chacón, A. Riscos-Núñez, M.J. Pérez-Jiménez, A 604
   P-lingua based simulator for tissue P systems with cell separation. Rom. J. Inf. Sci. Technol. 605
   17(1), 89–102 (2014) 606
- I. Pérez-Hurtado, D. Orellana-Martín, M.A. Martínez-del-Amor, L. Valencia-Cabrera, A. 607 Riscos-Núñez, M.J. Pérez-Jiménez, 11 years of P-Lingua: a backward glance, in *Proceedings* 608 of the 20th International Conference on Membrane Computing (CMC20), ed. by Gh. Păun 609 (2019), pp. 451–462
- 40. I. Pérez-Hurtado, D. Orellana-Martín, G. Zhang, M.J. Pérez-Jiménez, P-Lingua in two steps: 611 flexibility and efficiency. J. Membr. Comput. 1(2), 93–102 (2019). https://doi.org/10.1007/ 612 s41965-019-00014-1 613
- 41. T. Song, L. Pan, Gh. Păun. Asynchronous spiking neural P systems with local synchronization.
   614 Inf. Sci. 219, 197–207 (2013). https://doi.org/10.1016/j.ins.2012.07.023
   615
- 42. A. Spicher, O. Michel, M. Cieslak, J.-L. Giavitto, P. Prusinkiewicz, Stochastic P systems and 616 the simulation of biochemical processes with dynamic compartments. Biosystems, 91(3), 458–472 (2008). https://doi.org/10.1016/j.biosystems.2006.12.009
- 43. The P-Lingua website. http://www.p-lingua.org

29

- 44. L. Valencia-Cabrera, An Environment for Virtual Experimentation with Computational Models 620 Based on P Systems. Ph.D. Thesis (Universidad de Sevilla, Sevilla, 2015). http://hdl.handle. 621 net/11441/45362 622
- 45. L. Valencia-Cabrera, B. Song, Tissue P systems with promoter simulation with MeCoSim 623 and P-Lingua framework. J. Membr. Comput. 2(2), 95–107 (2020). https://doi.org/10.1007/ 624 s41965-020-00037-z 625
- 46. L. Valencia-Cabrera, T. Wu, Z. Zhang, L. Pan, M.J. Pérez-Jiménez, A simulation software tool for cell-like spiking neural P systems. Rom. J. Inf. Sci. Technol. 20(1), 71–84 (2017) 627
- 47. L. Valencia-Cabrera, D. Orellana-Martín, M.A. Martínez-del-Amor, A. Riscos-Núñez, M.J. 628
   Pérez-Jiménez, Computational efficiency of minimal cooperation and distribution in polarizationless P systems with active membranes. Fundam. Inform. 153(1–2), 147–172 (2017). https:// doi.org/10.3233/FI-2017-1535
- L. Valencia-Cabrera, D. Orellana-Martín, M.A. Martínez-del-Amor, A. Riscos-Núñez, M.J. 632 Pérez-Jiménez, Reaching efficiency through collaboration in membrane systems: dissolution, 633 polarization and cooperation. Theor. Comput. Sci. **701**, 226–234 (2017). https://doi.org/10. 634 1016/j.tcs.2017.04.015
- 49. L. Valencia-Cabrera, D. Orellana-Martín, M.A. Martínez-del-Amor, M.J. Pérez-Jiménez, An 636 interactive timeline of simulators in Membrane Computing. J. Membr. Comput. 1(3), 209–222 637 (2019). https://doi.org/10.1007/s41965-019-00016-z 638
- T. Wu, Z. Zhang, Gh. Păun, L. Pan, Cell-like spiking neural P systems. Theor. Comput. Sci. 639
   623, 180–189 (2016). https://doi.org/10.1016/j.tcs.2015.12.038

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