

# HAL (Hybrid Automata Library) Manual

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## Abstract

The presented Hybrid Automata Library (HAL) is a Java Library made of simple, efficient, generic components which can be used to model complex spatial systems. HAL's components can broadly be classified into: on and off lattice agent containers, finite difference diffusion fields, a gui building system, and additional tools and utilities for computation and collecting data. These components were designed to operate indepenently, but are standardized to make them easy to interface with one another. A complete example of how to build a hybrid model (a spatial model with an interacting agent based component and PDE component, commonly used for oncology modeling) using HAL is included to showcase how modeling can be simplified using our approach. HAL is a useful asset for researchers who wish to build efficient 2D and 3D hybrid models in Java, while not starting entirely from scratch. It is available on github at <https://github.com/torococo/HAL> under the MIT License. HAL requires at least Java 8 or later to run, and the java jdk version 1.8 or later to compile the source code.

## Contents

<b>1</b>	<b>Setup</b>	<b>3</b>
1.1	Getting Java . . . . .	3
1.2	Getting the Framework Source Code . . . . .	3
1.3	Getting IntelliJ IDEA . . . . .	3
1.4	Setting up the Project . . . . .	4
<b>2</b>	<b>Introduction</b>	<b>4</b>
2.1	Learning Java . . . . .	4
2.2	Using IntelliJ IDEA . . . . .	4
2.3	Understanding HAL . . . . .	5
2.4	Source Code Organization . . . . .	5
2.4.1	Examples . . . . .	5
2.4.2	LEARN_HERE . . . . .	5
2.4.3	Framework . . . . .	5
2.5	Framework organization . . . . .	5

<b>3</b>	<b>Grids and Agents</b>	<b>6</b>
3.1	Types of Grid	6
3.2	Types of Agent	6
3.3	Grid and Agent Class Definition	7
3.4	Grid Constructors	7
3.5	Agent Initialization Functions	8
3.6	Grid Indexing	8
3.6.1	Single Indexing	8
3.6.2	Square Indexing	8
3.6.3	Point Indexing	9
3.7	Typical Grid Loop	9
3.8	Type Heirarchy	9
3.9	Agent Functions and Properties	10
3.10	SphericalAgent Functions and Properties	11
3.11	Universal Grid Functions and Properties	12
3.12	AgentGrid Method Descriptions	13
3.12.1	AgentGrid Agent Search Functions	14
3.13	PDEGrid Method Descriptions	15
3.14	Griddouble/Gridint/Gridlong Method Descriptions	16
3.15	AgentList Method Descriptions	16
<b>4</b>	<b>Util.java</b>	<b>17</b>
4.1	Util Array Functions	17
4.2	Util Neighborhood Functions	17
4.3	Util Math Functions	18
4.4	Util Misc Functions	19
4.5	Util Color Functions	19
4.6	Util MultiThread Function	20
4.7	Util Save and Load Functions	20
<b>5</b>	<b>Rand.java</b>	<b>21</b>
<b>6</b>	<b>Gui</b>	<b>21</b>
6.1	Types of Gui and Method Descriptions	21
6.1.1	GridWindow	22
6.1.2	UIWindow	23
6.1.3	OpenGL2DWindow	23
6.1.4	OpenGL3DWindow	24
6.2	Types of GuiComponent and Method Descriptions	25
6.2.1	UIGrid	25
6.2.2	UILabel	25
6.2.3	UIButton	25
6.2.4	UIBoolInput	25
6.2.5	UIIntInput	25
6.2.6	UIDoubleInput	26
6.2.7	UIStringInput	26
6.2.8	UIComboBoxInput	26
6.2.9	UIFileChooserInput	26
6.3	GifMaker	26

<b>7</b>	<b>Tools</b>	<b>26</b>
7.1	Tools/ FileIO . . . . .	26
7.2	Tools/ MultiWellExperiment . . . . .	27
<b>8</b>	<b>Example: Competitive Release Model</b>	<b>28</b>
8.1	Competitive Release Introduction . . . . .	28
8.2	Main Function . . . . .	31
8.3	ExampleModel Constructor and Properties . . . . .	32
8.4	InitTumor Function . . . . .	33
8.5	ModelStep Function . . . . .	34
8.6	CellStep Function and Cell Properties . . . . .	35
8.7	DrawModel Function . . . . .	36
8.8	Imports . . . . .	37
8.9	Model Results . . . . .	37

# 1 Setup

## 1.1 Getting Java

As pretty as HAL's source code is, your going to need at least the Java8 JDK (Java Development Kit) installed to do anything with it.

To check if the JDK is installed, open a command line/terminal/cmd window and enter

---

```
1 java -version
2 javac -version
```

---

If both of these commands print 1.8.anything or later, you're good to go. If not, get the latest Java JDK by googling it or from the oracle website.

<http://www.oracle.com/technetwork/java/javase/downloads/jdk9-downloads-3848520.html>

You don't need to download the demos and samples.

## 1.2 Getting the Framework Source Code

To download the framework java files, go to

<https://github.com/torococo/HAL>

and click the clone or download button to get a zip file containing the source code along with the included examples.

unzip this folder and put it somewhere easily accessible.

## 1.3 Getting IntelliJ IDEA

IntelliJ idea is my favorite ide for programming in Java, and it probably should be yours too (<https://dzone.com/articles/why-idea-better-eclipse>). It can be downloaded here:

<https://www.jetbrains.com/idea/download/>

make sure you download the community edition, unless you really don't know what to do with your grant money.

## 1.4 Setting up the Project

1. Open IntelliJ Idea and click “create project from existing sources” (“file/ new/ project from existing sources” from the main gui) and direct it to the unzipped AgentFramework Source code directory.
2. Continue through the rest of the setup, you can basically click next until it asks for the Java sdk:
  - “/Library/ Java/ JavaVirtualMachines/” on mac.
  - “C:\ Program Files\ Java\” on windows.
1. Once the setup is complete we will need to do one more step and add some libraries that allow for 2D and 3D OpenGL visualization:
2. open the IntelliJ IDEA main gui
3. go to “file/ project structure”
4. click the “libraries” tab
5. use the minus button to remove any pre-existing library setup
6. click the plus button, and direct the file browser to the “Framework/ lib” folder.
7. click apply or ok

This will setup the classes, sources, and native library locations that OpenGL needs. Try running the “[Examples/Example3D](#)” program in the examples folder to see that everything is working properly

If you think that the colorscheme that comes with IntelliJ leaves something to be desired, you’re not alone. try “[File/ import settings](#)” and give it the HalColorSchemes.jar file in the top level folder.

## 2 Introduction

### 2.1 Learning Java

If your already pretty familiar with programming, and want a quick, shallow description of the language syntax, the first video on this list may be enough to get up to speed and jump into the framework:

[https://www.youtube.com/results?search\\_query=learn+java](https://www.youtube.com/results?search_query=learn+java)

The resources are listed in order by popularity, just try some until you find one that gives well. For a more textual perspective,

<https://www.codecademy.com/learn/learn-java>

<http://www.learnjavaonline.org/>

are some potential places to start.

### 2.2 Using IntelliJ IDEA

Some of the ways in which IntelliJ may make your life easier include:

- automatically importing classes as they are first mentioned in the code (right click on the class name and intelliJ should offer to import it)

- debugging with the debugger (shocker) when necessary rather than relying on print statements alone (<https://www.youtube.com/watch?v=1bCgzjatcr4>), HINT: while paused in the debugger right click on anything and click “evaluate expression”
- using refactoring to rename variables, change function signatures, etc. without having to go hunting for every place that the variable/function/class is mentioned. (right click on some code and check out the “refactor” submenu)
- using “find usages” and “Go To Declaration” to move fluidly around your code base and to see how its pieces connect together. (right clicking on things will get you there, see “find usages” and the “go to” submenu)
- using Shift-F10 to quickly run the currently open file, and learning other hotkeys to speed up your workflow.
- tapping Shift twice allows you to search the entire codebase for anything
- using fori, iter, and sout shortcuts to create a for loop, foreach loop, and print statement respectively

Don't worry too much about learning all of these up front, but I suggest that you do check them out as you become more comfortable with the platform.

## 2.3 Understanding HAL

Once you have a basic grasp of Java, I recommend skimming this manual as well as checking out the [LEARN\\_HERE](#) folder. then look at the examples

## 2.4 Source Code Organization

At the top level, there are 3 folders which hold different framework parts. These are...

### 2.4.1 Examples

contains models that use the framework as a basis

### 2.4.2 LEARN\_HERE

contains small examples that serve as tests of framework components

### 2.4.3 Framework

the top level framework source code folder, contains all of the following subfolders that hold various framework components. Descriptions of the 7 subfolders contained within the framework folder are detailed below.

## 2.5 Framework organization

**Grids** this folder contains all of the Agent and Grid types that the framework supports. There are 2D and 3D, stackable and unstackable, on-lattice and off-lattice agents and grids to contain them. The base classes that the grids and agents extend are also in this folder.

**Gui** this folder contains the main UIWindow class, as well as many component classes that can be added to the UI.

**Tools** this folder contains many useful tool classes, such as a FileIO wrapper, a genetic algorithm class, a multiwell experiment runner, etc. as well as a Util class that is a container of generic static methods. It also contains classes that are used internally by these tools.

**Interfaces** this folder contains interfaces that are used by the framework internally. It is useful to reference this folder if a framework function takes a function interface argument, or implements an interface.

**Lib** contains outside libraries that have been integrated with the framework.

**Util** The only file directly on the top level, it's full of goodies that can be broadly categorized into: color functions, array functions, cellular automata neighborhood functions, and math functions.

The manual from here turns into a glossary that looks at the framework components in some detail (to the extent that I bothered to write about them) and after that a simple but complete model example is provided.

Those looking to dive headfirst into modeling may want to first go to the [LEARN\\_HERE](#) folder for a hands on approach, or check out the last section for an in depth explanation of a model and use the other chapters as a reference. If you like reading dictionaries in your spare time then read this from end to end.

## 3 Grids and Agents

The bread and butter of the framework consists of different types of Grids and Agents. these are presented below.

### 3.1 Types of Grid

**AgentGrid0D** Holds nonspatial agents

**AgentGrid1D** Holds all 1D agents

**AgentGrid2D** Holds all 2D agents

**AgentGrid3D** Holds all 3D agents

**PDEGrid1D** facilitates modeling a single diffusible field in 1D

**PDEGrid2D** facilitates modeling a single diffusible field in 2D

**PDEGrid3D** facilitates modeling a single diffusible field in 3D

### 3.2 Types of Agent

**Agent0D** Nonspatial agents. this agent type is a member of AgentGrid0D grid type.

**AgentSQ2Dunstackable** "Square" agents in 2D. this agent type is bound to the AgentGrid2D lattice, and only one AgentSQ2Dunstackable can occupy a given lattice position at a time.

**AgentSQ2D** "Square" agents in 2D. this agent type is bound to the AgentGrid2D lattice, however multiple AgentSQ2Ds can occupy the same lattice position

**AgentPT2D** "Point" agents in 2D. this agent type is not bound to the AgentGrid2D lattice, and is free to move continuously within the Grid, provided it does not try to cross the Grid boundaries.

**AgentSQ2Dunstackable** "Square" agents in 2D. this agent type is bound to the AgentGrid2D lattice, and only one AgentSQ2Dunstackable can occupy a given lattice position at a time.

**AgentSQ2D** "Square" agents in 2D. this agent type is bound to the AgentGrid2D lattice, however multiple AgentSQ2Ds can occupy the same lattice position

**AgentPT2D** "Point" agents in 2D. this agent type is not bound to the AgentGrid2D lattice, and is free to move continuously within the Grid, provided it does not try to cross the Grid boundaries.

**AgentSQ3Dunstackable** "Square" agents in 3D (cube?). this agent type is bound to the AgentGrid3D lattice, and only one AgentSQ3Dunstackable can occupy a given lattice position at a time (gotta love copy paste)

**AgentSQ3D** "Square" agents in 3D. this agent type is bound to the AgentGrid3D lattice, however multiple AgentSQ3Ds can occupy the same lattice position

**AgentPT3D** "Point" agents in 3D. this agent type is not bound to the AgentGrid3D lattice, and is free to move continuously within the Grid, provided it does not try to cross the Grid boundaries.

### 3.3 Grid and Agent Class Definition

for demonstration we will use bits of code from the CompetitiveReleaseModel example. It is a fairly simple yet complete model, so it is a good place to begin learning the syntax of HAL. Grid classes and Agent classes used in models will usually be created as extensions of the Grid and Agent classes shown above. This extension is done with the following syntax:

Project specific AgentGrid definition syntax:

---

```
1 public class ExampleModel extends Grid2D<ExCell> {
```

---

Project specific Agent class definition syntax:

---

```
1 class ExampleCell extends AgentSQ2Dunstackable<ExModel> {
```

---

Note the <> after the base class name, in java this is called a generic type argument. This is how we tell the Grid what kinds of Agent it will store, and how we tell the Agents what kind of Grid will store them. It is used by the ExampleModel and ExampleCell to identify each other, so that their constituent functions can return the proper type, and access each other's variables and methods.

### 3.4 Grid Constructors

In order to create a class that extends any of the Grids, you must provide a constructor. let's look at part of the ExampleGrid constructor as an example

---

```
1 public ExampleModel(int x, int y, Rand generator) {
2     super(x, y, ExampleCell.class);
```

---

the first line declares the constructor and arguments, the second line calls super, which is required since our class extends a class with a constructor. Super is used to call the constructor of the base class. Into super we pass what the AgentGrid2D needs for initialization: an x and y dimension, which define the size of the grid, and ExampleCell.class, which is the class object of the ExampleCell. We pass the class object so that the ExampleGrid can create ExampleCells for us, as described in the next section.

### 3.5 Agent Initialization Functions

a word of caution: **DO NOT DEFINE A CONSTRUCTOR FOR YOUR AGENT CLASSES.**

The AgentGrid that houses the agent will act as a “factory” for agents, and produce them with the NewAgent() function. This will return an agent that is either newly constructed, or an agent that has died and is being recycled. This returning of dead agents for reuse as new ones allows the model to run without tasking the garbage collector with removing all of the dead agents. This will increase the speed and decrease the memory footprint of your model. Instead of a constructor you should define some sort of initialization for your agents, which you do directly after the call to NewAgent. An example from CompRelModel.java:

```
1 for (int i = 0; i < hoodSize; i++) {  
2     if (rng.Double() < resistantProb) {  
3         NewAgentSQ(tumorNeighborhood[i]).type = RESISTANT;  
4     } else {  
5         NewAgentSQ(tumorNeighborhood[i]).type = SENSITIVE;  
6     }  
7 }
```

These lines of code come from the InitTumor function that the ExampleModel calls once at the beginning of a simulation. We use a random number generator and an if-else statement to decide whether to create a sensitive or resistant cell. since this type property is the only information individual cells store in this model, setting it is all that is needed to initialize a new cell. We call the NewAgentSQ() function, and pass in an index (“tumorNeighborhood” is an array of starting indices to setup the tumor) that marks where to place the new agent.

### 3.6 Grid Indexing

There are 3 different ways to describe or index locations on framework grids:

#### 3.6.1 Single Indexing

since the x dimension, y dimension, and possibly the z dimension values are Grid constants, every square or voxel can be uniquely identified with a single integer index. Functions using this kind of indexing typically end with the SQ (abbreviating Square) phrase at the end of the function name. Single indexing is done with the following mappings:

In 2D:  $I(x, y) = x * yDim + y$

In 3D:  $I(x, y, z) = x * yDim * zDim + y * zDim + z$

the agents/values in the grids are stored as a single array, so single indexing is actually the most efficient as it requires no conversion.

#### 3.6.2 Square Indexing

Similar to single indexing, square indexing uses a set of integers to refer to a specific square or voxel, as an (x,y) or an (x,y,z) set. Functions using this kind of indexing typically end with the SQ (abbreviating Square) phrase at the end of the function name.



### 3.6.3 Point Indexing

Uses a set of double values, to define continuous coordinates. Functions using this kind of indexing typically end with the PT (abbreviating Point) phrase at the end of the function name. The integer flooring of a coordinate set corresponds to the Square or Voxel that contains the point.

## 3.7 Typical Grid Loop

here we look at an example Loop or Run function. This is taken from the GOLGrid class, but all models will usually follow a similar pattern.

```
1 public void Run(){
2     for (int i = 0; i < runTicks; i++) {
3         for (GOLAgent a : this) {
4             a.Step();
5         }
6     }
7 }
```

The outer for loop counts the ticks, meaning that we will run for a total of runTicks steps. The inner for loop iterates over all the agents in the grid ("this" here is the grid that calls the Run function), and inside the loop we call that agent's Step function, which is defined in the GOLAgent class.

## 3.8 Type Heirarchy

in order to navigate the source code and see the full set of functions and properties of the framework components, it is important to become familiar with the type heirarchy that the framework uses. Figure 1 summarizes this heirarchy for 2D agents.

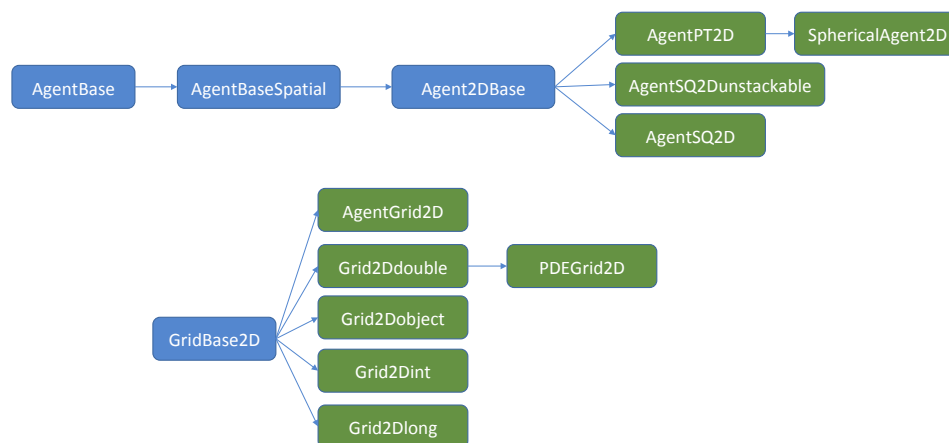


Figure 1:

each node in the heirarchy names a class. each arrow denotes an extends relationship, eg. AgentBaseSpatial extends AgentBase. The blue classes are abstract and cannot be used directly. The green classes are fully implemented and can be used/extended further. classes later in the heirarchy keep all properties and methods of the classes that they extend, which means that to see the full set of functions a class implements, one has to look at all of the extended classes beneath that class as well. the method summaries included in this manual can simplify this process somewhat as they show useful methods of the extended classes regardless of where they come from in the class heirarchy.

### 3.9 Agent Functions and Properties

here we describe the builtin functions that the Agents expose to the user. (3D adds a z)

**G:** returns the grid that the agent belongs to (this is a permanent agent property, not a function)

**Age():** returns the age of the agent, in ticks. Be sure to use IncTick on the AgentGrid appropriately for this function to work.

**BirthTick():** returns the tick on which the agent was born

**IsAlive():** returns whether or not the agent currently exists on the grid

**Isq():** returns the index of the square that the agent is currently on

**Xsq(),Ysq():** returns the X or Y indices of the square that the agent is currently on.

**Xpt(),Ypt():** returns the X or Y coordinates of the agent. If the Agent is on-lattice, these functions will return the coordinates of the middle of the square that the agent is on.

**MoveSQ(x,y),MoveSQ(i):** moves the agent to the middle of the square at the indices/index specified

**MovePT(x,y):** moves the agent to the coordinates specified

**MoveSafeSQ(x,y),MoveSafePT(x,y):** Similar to the move functions, only it will automatically either apply wraparound, or prevent moving along a particular axis if movement would cause the agent to go out of bounds.

**Dispose():** removes the agent from the grid

**SwapPoosition(otherAgent):** swaps the positions of two agents. useful mostly for the AgentSQ2unstackable and AgentSQ3unstackable classes, which don't allow stacking of agents, making this maneuver otherwise difficult.

**MapHood(int[]neighborhood):** This function takes a neighborhood (As generated by the neighborhood Util functions, or using GenHood2D/GenHood3D), translates the neighborhood coordinates to be centered around the agent, and computes the set of indices that the translated coordinates map to. The function returns the number of valid locations it set, which if wraparound is disabled may be less than the full set of coordinate pairs. this means that the passed in ret list will not be completely filled with valid entries. See the CellStep function in the Complete Model example for more information.

**MapEmptyHood(int[]neighborhood),MapOccupiedHood(int[]neighborhood):** These functions are similar to the the MapHood function, but they will only include valid empty or occupied indices, and skip the others.

**MapHood(int[]neighborhood,CoordsToBool):** This function takes a neighborhood and another mapping function as argument, the mapping function should return a boolean that specifies whether coordinates map to a valid location. the function will only include valid indices, and skip the others.

**Xdisp(x),Ydisp(y):** returns the displacement from the agent to a given x or y position

**Dist(x,y):** returns the distance from this agent to the given position

### 3.10 SphericalAgent Functions and Properties

an extension of the AgentPT2D and AgentPT3D agent types, spherical agents have the additional property of a radius and x and y velocities. These properties allow spherical agents to behave like spheres that repel each other. (3D adds a z)

**radius:** the radius property is used during SumForces to determine collisions

**xVel,yVel:** the x and y velocity properties are added to by SumForces and applied to the agent's position by calling ForceMove. adding to these properties can cause agents to move in a particular direction

**Init(radius):** a default initialization function that sets the radius based on the argument, and the x and y velocities to 0

**SumForces(interactionRad,OverlapForceResponse):** The interactionRad argument is a double that specifies how far apart to check for other agent centers to interact with, and should be set to the maximum distance apart that two interacting agent centers can be. the OverlapForceResponse argument must be a function that takes in an overlap and an agent, and returns a force response. aka. (double,Agent) -> double. the double argument is the extent of the overlap. if this value is positive, then the two agents are "overlapping" by the distance specified. if the value is negative, then the two agents are separated by the distance specified. the OverlapForceResponse should return a double which indicates the force to apply to the agent as a result of the overlap. if the force is positive, it will repel the agent away from the overlap direction, if it is negative it will pull it towards that direction. SumForces alters the xVel and yVel properties of the agent by calling OverlapForceResponse using every other agent within the interactionRad.

**ForceMove():** adds xVel and yVel property values to the x,y position of the agent.

**Divide(divRadius,scratchCoordArr,Rand):** Facilitates modeling cell division. The divRadius specifies how far apart from the center of the parent agent the daughters should be separated. the scratchCoordArr will store the randomly calculated axis of division. The axis is calculated using the Rand argument (HAL's random number generator object) if no Rand argument is provided, the values currently in the scratchCoordArr will be used to determine the axis of division. The first entry of scratchCoordArr is the x component of the axis, the second entry is the y component. division is achieved by placing the newly generated daughter cell divRadius away from the parent cell using divCoordArr for the x and y components of the direction, and placing the parent divRadius away in the negative direction. Divide returns the newly created daughter cell.

**CapVelocity(maxVel):** caps the xVel and yVel variables such that their norm is not greater than maxVel

**ApplyFriction(frictionConst):** multiplies xVel and yVel by frictionConst. if frictionConst = 1, then no friction force will be applied, if frictionConst = 0, then the cell won't move.

### 3.11 Universal Grid Functions and Properties

These functions and properties are shared by all of the different types of grids (3D adds a z):

**xDim,yDim:** the x or y dimension of the Grid

**length:** the total number of squares in the grid, equivalent to  $xDim*yDim$

**I(x,y):** converts a set of coordinates to the index of the square at those coordinates.

**ItoX(i),ItoY(i):** converts an index of a square to that square's X or Y coordinate

**WrapI(x,y):** returns the index of the square at the provided x,y coordinates, with wraparound.

**In(x,y):** returns whether the x and y coordinates provided are inside the Grid.

**ConvXsq(x,otherGrid),ConvYsq(y,otherGrid):** returns the index of the center of the square in otherGrid that the coordinate maps to.

**ConvXpt(x,otherGrid),ConvYpt(y,otherGrid):** returns the position that the x/y argument rescales to in otherGrid

**DispX(x1,x2),DispY(y1,y2):** gets the displacement from the first coordinate to the second. using wraparound if allowed over the given axis to find the shortest displacement.

**Dist(x1,y1,x2,y2):** gets the distance between two positions with or without grid wrap around (if wraparound is enabled, the shortest distance taking this into account will be returned)

**DistSquared(x1,y1,x2,y2):** gets the distance squared between two positions with or without grid wrap around (if wraparound is enabled, the shortest distance taking this into account will be returned) more efficient than the Dist function above as it skips a square-root calculation.

**MapHood(int[]hood,centerX,centerY):** This function takes a neighborhood centered around the origin, translates the set of coordinates to be centered around a particular central location, and computes which indices the translated coordinates map to. The function returns the number of valid locations it set. this function differs from HoodTols and CoordsTols in that it takes no ret[], MapHood instead puts the result of the mapping back into the hood array.

**MapHood(int[]hood,centerX,centerY,EvaluationFunction):** This function is very similar to the previous definition of MapHood, only it additionally takes as argument an EvaluationFunction. this function should take as argument (i,x,y) of a location and return a boolean that decides whether that location should be included as a valid one.

**IncTick():** increments the internal grid tick counter by 1, used with the Age() and BirthTick() functions to get age information about the agents on an AgentGrid. can otherwise be used as a counter with the other grid types.

**GetTick():** gets the current grid timestep.

**ResetTick():** sets the tick to 0.

**ApplyRectangle(startX,startY,width,height>ActionFunction):** applies the action function to all positions in the rectangle, will use wraparound if appropriate

**ApplyHood(int[]hood,centerX,centerY,ActionFunction):** applies the action function to all position in the neighborhood

**ApplyHoodMapped(int[]hood,validCount,ActionFunction):** applies the action function to all positions in the neighborhood up to validCount, assumes the neighborhood is already mapped

**ContainsValid(int[]hood,centerX,centerY,EvaluationFunction):** returns whether a valid index exists in the neighborhood

**BoundaryIs():** returns a list of indices, where each index maps to one square on the boundary of the grid

**AlongLineIs(x1,y1,x2,y2,int[]writeHere):** returns the set of indices of squares that the line between (x1,y1) and (x2,y2) touches.

### 3.12 AgentGrid Method Descriptions

here we describe the builtin functions that the agent containing Grids expose to the user. Most functions that take x,y arguments can alternatively take a single index argument. The Grid will use the index to find the appropriate square. (3D adds a z)

**NewAgentSQ(x,y),NewAgentSQ(i):** returns a new agent, which will be placed at the center of the indicated square. x,y, and i are assumed to be integers

**NewAgentPT(x,y):** returns a new agent, which will be placed at the coordinates specified. x and y are assumed to be doubles

**Pop():** returns the number of agents that are alive on the entire grid.

**CleanAgents():** reorders the list of agents so that dead agents will no longer have to be iterated over. don't call this during the middle of iteration!

**ShuffleAgents():** shuffles the list of agents, so that they will no longer be iterated over in the same order. don't call this during the middle of iteration!

**CleanShuffle():** Cleans the agent list and shuffles it. This function is often called at the end of a timestep. don't call this during the middle of iteration!

**Reset():** disposes all agents, and resets the tick counter.

**MapHoodEmpty(int[]coords,centerX,centerY):** similar to the MapHood function, but will only include indices of locations that are empty

**MapHoodOccupied(int[]coords,centerX,centerY):** similar to the MapHood function, but will only include indices of locations that are occupied

**RandomAgent(rand):** returns a random living agent

### 3.12.1 AgentGrid Agent Search Functions

Several convenience methods have been added to make searching for agents easier.

**GetAgent(x,y)GetAgent(i):** Gets a single agent at the specified grid square, beware using this function with stackable agents, as it will only return one of the stack of agents. This function is recommended for the Unstackable Agents, as it tends to perform better than the other methods for single agent accesses.

**GetAgentSafe(x,y):** Same as GetAgent above, but if x or y are outside the domain, it will apply wrap around if wrapping is enabled, or return null.

**IterAgents(x,y),IterAgents(i):** use in a foreach loop to iterate over all agents at a location. example: `for(AGENT_TYPE a : IterAgents(5,6))` will run a for loop over all agents at grid square (5,6) with the agent being stored as the "a" variable. be sure to set AGENT\_TYPE to the type of agent that lives of the grid that you are iterating over.

**IterAgentsSafe(x,y):** Same as IterAgents above, but will apply wraparound if x,y fall outside the grid dimensions.

**IterAgentsRad(xPT,yPT,rad):** will iterate over all agents around the given coordinate pair that fall within radius rad.

**IterAgentsRadApprox(xPT,yPT,rad):** will iterate over all agents around the given coordinate pair that at least fall within radius rad, it is more efficient than IterAgentsRad, but it will also include some agents that fall outside rad, so be sure to do an additional distance check inside the function.

**IterAgentsHood(int[]hood,centerX,centerY):** will iterate over all agents in the neighborhood as though it were mapped to centerX,centerY position. note that this function won't technically do the mapping. if the neighborhood is already mapped, use IterAgentsHoodMapped instead.

**IterAgentsHoodMapped(int[]hood,hoodLen):** will iterate over all agents in the already mapped neighborhood. be sure to supply the proper hood length.

**IterAgentsRect(x,y,width,height):** iterates over all agents in the rectangle defined by (x,y) as the lower left corner, and (x+width,y+height) as the top right corner.

**GetAgents(ArrayList<AgentType>,x,y)GetAgentsHood...:** A variation of this function exists that matches every IterAgents variant mentioned above. puts into the argument arraylist all agents found at the specified location. call the clear function on the arraylist before passing it to GetAgents if you don't want to append to whatever agents were already added there.

**RandomAgent(x,y,rand,EvalAgent)RandomAgentHood...:** A variation of this function exists that matches every IterAgents variant mentioned above. these functions are used to query a location and choose a random agent from that area, an optional EvalAgent function (a function that takes an Agent and returns a boolean) can be provided to ensure that the random agent satisfies the condition

**PopAt(x,y),PopAt(i):** returns the number of agents that occupy the specified position. this is more efficient than using GetAgents and taking the length of the resulting arraylist.

### 3.13 PDEGrid Method Descriptions

The PDEGrid classes do not store agents. They are meant to be used to model Diffusible substances. PDEGrids contain two fields (arrays) of double values: a current field, and a next field. The intended usage of these fields is that the values from the current timestep are stored in the current field while changes due to transformations are accumulated in the next field. Central to the function of the PDEGrid is the Update() function, which updates the current field with the values in the next field. The functions included in the PDEGrid class are the following: (3D adds a z, 1D removes y)

**Get(x,y):** returns the “current field” double array which is usually used as the main field that agents interact with

**Add(x,y,val),Mul(x,y,val):** adds or multiplies a value in the “current field” and adds the change to the “next field”

**Set(x,y,val):** sets a value in the “next field”. any previous changes will be overwritten.

**GetAll(val),SetAll(val[]),AddAll(val),MulAll(val):** applies the Set/Get/Mul operations to all entries of the current field

**MaxDifRecord():** returns the maximum difference in any single lattice position between the current field and the current field the last time MaxDifRecord was called. (the function saves a record of the state for future comparison whenever it is called)

**MaxDifNext():** returns the maximum difference in any single lattice position between the current field and the next field.

**MaxDifNextScaled(denomOffset):** returns

**Update():** adds the next field into the current field.

**Diffusion(diffCoef):** runs diffusion on the current field, adding the deltas to the next field. This form of the function assumes either a reflective or wrapping boundary (depending on how the PDEGrid was specified). the diffCoef variable is the nondimensionalized diffusion coefficient. If the dimensionalized diffusion coefficient is  $x$  then diffCoef can be found by computing  $\frac{x * SpaceStep}{TimeStep^2}$ . Note that if the diffCoef exceeds 0.25, this diffusion method will become numerically unstable.

**Diffusion(diffCoef,boundaryValue):** has the same effect as the above diffusion function without the boundary value argument, except rather than assuming zero flux, the boundary condition is set to either the boundaryValue, or wrap around

**DiffusionADI(diffCoef):** runs diffusion on the current field using the ADI (alternating direction implicit) method. without a boundaryValue argument, a zero flux boundary is imposed. wraparound will not work with ADI. ADI is numerically stable at any diffusion rate.

**DiffusionADI(diffCoef,boundaryValue):** runs diffusion on the current field using the ADI (alternating direction implicit) method. ADI is numerically stable at any diffusion rate. Adding a boundary value to the function call will cause boundary conditions to be imposed.

**Advection(xVel,yVel):** runs advection, which moves the concentrations using a constant flow with the x and y velocities passed. this signature of the function assumes wrap-around, so there can be no net flux of concentrations.

**Advection(xVel,yVel,boundaryValue):** runs advection as described above with a boundary value, meaning that the boundary value will advect in from the upwind direction, and the concentration will disappear in the downwind direction.

**GradientX(x,y),GradientY(x,y):** returns the gradient of the diffusible in a direction at the coordinates specified

**GetAvg():** returns the mean value of the field

**GetMax():** returns the max value in the field

**GetMin():** returns the min value in the field

**SetAll(val),SetAll(val[])AddAll(val)MulAll(val):** applies the Set/Get/Mul operations to all entries of the field, adding the deltas to the next field

### 3.14 Griddouble/Gridint/Gridlong Method Descriptions

As an alternative to this class, it may be useful to simply employ a double/int/long array whose length is equal to the length of the other associated grids. The I() function of any associated grids can be used to access values in the double array with x,y or x,y,z coordinates.

**GetField():** returns the field double array that the grid stores

**Get(i),Get(x,y),Set(i),Set(x,y,val),Add(i),Add(x,y,val),Mul(i,val),Mul(x,y,val):** gets, sets, adds, or multiplies with a single value in the field at the specified coordinates

**SetAll(val),SetAll(val[])AddAll(val)MulAll(val):** applies the Set/Get/Mul operations to all entries of the field

**BoundAll(min,max),BoundAllSwap(min,max):** sets all values in the field so that they are between min and max

**GradientX(x,y),GradientY(x,y):** returns the gradient of the field in a direction at the coordinates specified

**GetAvg():** returns mean value of the grid

**GetMax():** returns the max value in the grid

**GetMin():** returns the min value in the grid

### 3.15 AgentList Method Descriptions

AgentLists allow for keeping track of agent sub-populations. AgentLists must be instantiated with a type argument which specifies the type of agent that they will hold. When an agent dies, it is automatically removed from any AgentLists that contain it, so the AgentList will only contain living agents. It is not necessarily ordered. Use the java foreach loop syntax to iterate over the AgentList, just like the AgentGrids.

**AddAgent(agent):** adds an agent to the AgentList, agents can be added multiple times.

**RemoveAgent(agent):** removes all instances of the agent from the AgentList, returns the number of instances removed

**GetPop():** returns the size of the AgentList, duplicate agents will be counted multiple times



**InList(agent):** returns whether a given agent is already in the AgentList

**ShuffleAgents(rng):** Shuffles the agentlist order

**CleanAgents():** may speed up AgentList iteration if many agents from the AgentList have died recently

**RandomAgent(rng):** returns a random agent from the AgentList

## 4 Util.java

The Util class is one of the most ubiquitous classes in the framework, and contains all of the generic functions that wouldn't make sense to add to any particular object in the framework.

The list of utilities functions in the framework has only grown with time. the list presented here is not exhaustive, so I recommend looking at the file itself if you feel something is missing, on top of that feel free to ask for a new feature or better yet send me an implementation and I'll most likely add it to the repository for everyone to share.

### 4.1 Util Array Functions

A set of utilities for making array manipulation easier.

**ArrToString(arr,delim):** useful for collecting data or print statement debugging, returns the contents of an array as a single string. entries are separated by the delimiter argument

**IndicesArray(numEntries):** generates an array of ascending indices starting with 0, up to nEntries.

**Mean(arr):** returns the mean of an array

**Sum(arr):** returns the sum of an array

**Norm(arr):** returns the euclidean norm of an array

**NormSq(arr):** returns the squared euclidian norm of an array, somewhat more efficient than Norm

**SumTo1(arr):** scales all entries in an array so that their sum is 1.

**Normalize(arr):** scales all entries in an array so that their norm is 1.

### 4.2 Util Neighborhood Functions

a set of utilities for generating neighborhoods. neighborhoods are lists of x,y index pairs, of the form  $[0_1, 0_2, \dots 0_n, x_1, y_1, x_2, y_2, \dots x_n, y_n]$  in the third dimension these are  $[0_1, 0_2, \dots 0_n, x_1, y_1, z_1, x_2, y_2, z_2, \dots x_n, y_n, z_n]$  these arrays are useful when finding the indices of the locations that make up the neighborhood around an agent when using Grid/Agent functions such as MapHood. Neighborhood arrays are always padded with 0s at the beginning, for use with the MapHood functions.

These functions should not be called over and over, as this would wastefully create arrays over and over. instead the function should be called once and be stored by the grid.

**GenHood2D(int[]coords), GenHood3D(int[]coords):** generates a 2D/3D neighborhood array for use with the MapHood functions. input should be an integer array of the form  $[x_1, y_1, x_2, y_2, \dots x_n, y_n]$  in 2D, or  $[x_1, y_1, z_1, x_2, y_2, z_2, \dots x_n, y_n, z_n]$  in 3D.

**VonNeumannHood(origin?),MooreHood(origin?):** returns an array that contains the coordinates of the VonNeumann or Moore Neighborhood in 2D. the boolean argument specifies whether the center of these neighborhood (0,0) should be included as part of the set of coordinates.

**VonNeumannHood3D(origin?),MooreHood3D(origin?):** returns an array that contains the coordinates of the VonNeumann or Moore Neighborhood in 3D. the boolean argument specifies whether the center of these neighborhood (0,0) should be included as part of the set of coordinates.

**HexHoodEvenY(origin?),HexHoodOddY(origin?):** to simulate a hex lattice on a Grid2D, the neighborhood used for a given agent changes depending on the position of the agent. The hex hood changes depending on whether the agent is on an even or odd Y position. These functions return an array that contains the proper coordinates for a given case.

**TriangleHoodSameParity(origin?),TriangleHoodDifParity(origin?):** to simulate a triangle lattice on a Grid2D, the neighborhood used for a given agent changes depending on the position of the agent. The Triangle hood changes depending on whether the parity ("even-ness") of the agents X and Y position match. These functions return an array that contains the proper coordinates for a given case.

**CircleHood(origin?,radius):** generates a neighborhood of all squares within the radius of a starting position at the middle of the (0,0) origin square. the boolean argument specifies whether (0,0) should be included in the return array.

**RectangleHood(origin?,radX,radY):** generates a neighborhood of all squares whose x displacement is within radX, and whose y displacement is within radY of a starting position at the middle of the (0,0) origin square. the boolean argument specifies whether (0,0) should be included in the return array.

**AlongLineCoords(x1,y1,x2,y2):** returns an array of all squares that touch a line between the two starting positions.

### 4.3 Util Math Functions

For all of the following functions, and throughout the framework, rn refers to a random number generator.

**InfiniteLinesIntersection2D(x1,y1,x2,y2,x3,y3,x4,y4,double[]ret):** computes the intersection of lines between points 1 and 2, and points 3 and 4. puts the coordinates of the intersection point in ret. returns true if the infinite lines intersect (if they are not parallel). returns false if the lines are parallel.

**Bound(val,min,max):** returns the value bounded by the min and max.

**Rescale(val,min,max):** assumes the starting value is in the 0-1 scale, and rescales it to be in the min-max scale.

**ModWrap(val,max):** wraps the value provided so that it must be between 0 and max. used to implement wraparound by the framework.

**ProtonsToPh(protonConc):** converts proton concentration to ph

**PhToProtons(ph):** converts ph to proton concentration

**ProbScale(prob,duration):** converts the probability that an event happens in unit time to the probability that that same event happens in duration time.

**Sigmoid(val,stretch,inflectionValue,minCap,maxCap):** val is the value that the sigmoid function is being applied to, the stretch argument stretches or shrinks the sigmoid function along the x axis, the inflectionValue governs where the inflection point of the sigmoid is, minCap and maxCap bound the sigmoid along the y axis.

#### 4.4 Util Misc Functions

**TimeStamp():** returns a timestamp string with format "YYYY\_MM\_DD\_HH\_MM\_SS"

**PWD():** returns the current working directory as a string

**MemoryUsageStr():** returns a string with information about the current memory usage of the program.

**QuickSort(sortMe,greatestToLeast):** requires the passed sortMe class to implement the Sortable interface. the passed boolean indicates whether the array should be sorted from greatest to least or least to greatest.

#### 4.5 Util Color Functions

These functions generate integers that store RGBA (Red,Green,Blue,Alpha) color channels internally, 8 bits per channel (integer values 0-255). These so-called "ColorInts" are intended to be used as arguments for the UIGrid, GridWindow, Vis2DOpenGL, and Vis3DOpenGL to set the color of the pixels or objects being displayed. the RGB components set the color, and the alpha component adds transparency.

*All of these functions (except the "Getters") return a new ColorInt*

**RGB(r,g,b):** sets the rgb color channels using the continuous 0-1 range mapping. the alpha is always set to 1.

**RGBA(r,g,b,a):** sets the rgb and alpha channels using the continuous 0-1 range mapping.

**RGB256(r,g,b):** sets the rgb color channels using the discrete 0-255 range mapping. the alpha is always set to 1.

**RGBA256(r,g,b,a):** sets the rgb and alpha channels using the 0-255 range mapping.

**GetRed(color),GetBlue(color),GetGreen(color),GetAlpha(color):** returns the value of a single channel using the continuous 0-1 range mapping

**GetRed256(color),GetBlue256(color),GetGreen256(color),GetAlpha256(color):** returns the value of a single channel using the discrete 0-255 range mapping

**SetRed(color),SetGreen(color),SetBlue(color),SetAlpha(color):** returns a new ColorInt with the one of its channels changed compared to the argument passed in, uses the continuous 0-1 range mapping

**SetRed256(color),SetGreen256(color),SetBlue256(color),SetAlpha256(color):** returns a new ColorInt with the one of its channels changed compared to the argument passed in, uses the discrete 0-255 range mapping

**CategoricalColor(index):** returns a categorical color from a nice mutually distinct set. Valid indices are 0-19. the color order is (blue,red,green,yellow,purple,orange,cyan,pink,brown,light blue,light red,light green,light yellow,light purple, light orange, light cyan,light pink, light brown, light gray, dark gray

**HeatMapRGB(val),HeatMap???(val):** returns a new colorInt using the heatmap color scale. values are distinguished in the 0-1 range. the heatmap color scale is black at 0, white at 1, and transitions between these by changing one color channel at a time from none to full. the order that the channels are changed is dictated by the order of the 3 letters at the end of the function name. the possible orders are RGB, RBG, GRB, GBR, GRB and GBR

**HeatMapRGB(val,min,max),HeatMap???(val,min,max):** same as HeatMapRGB with fewer args, but works to distinguish values in the min-max range.

**HSBColor(hue,saturation,brightness):** returns a new colorInt using the HSB colorspace. values are distinguished in the continuous 0-1 range.

**YCbCrColor(y,cb,cr):** returns a new colorInt using the YCbCr colorspace. values are distinguished in the continuous 0-1 range.

**CbCrPlaneColor(x,y):** returns a new colorInt using the CbCr plane at Y=0.5. a very nice colormap for distinguishing position in a 2 dimensional space. x,y are expected to be in the continuous 0-1 range.

## 4.6 Util MultiThread Function

**Multithread(nRuns,nThreads,RunFun):** A function so useful that it deserves its own section, the multithread function creates a thread pool and launches a total of nRun threads, with nThreads running simultaneously at a time. the RunFun that is passed in must be a void function that takes an integer argument. when the function is called, this integer will be the index of that particular run in the lineup. This can be used to assign the result of many runs to a single array, for example, if the array is written to once by each RunFun at its run index. If you want to run many simulations simultaneously, this function is for you.

## 4.7 Util Save and Load Functions

**NOTE:SerializableModel\_Interface** In order to save and load models, you must first have the model extend the SerializableModel interface (this interface is defined in the Interfaces folder). this interface has one method, called SetupConstructors. all you have to do in this method is call the AgentGrid function \_PassAgentConstructor(AGENT.cass) once for each AgentGrid in your model, where AGENT is the type of agent that the AgentGrid holds. see LEARN\_HERE/Agents/SaveLoadModel for an example.

**SaveState(model):** Saves a model state to a byte array and returns it. The model must implement the SerializableModel interface

**SaveState(model,fileName):** Saves a model state to a file with the name specified. creates a new file or overwrites one if the file already exists. The model must implement the SerializableModel interface

**LoadState(byte[]state):** Loads a model form a byte array created with SaveState. The model must implement the SerializableModel interface

**LoadState(fileName):** Loads a model from a file array created with SaveState. The model must implement the SerializableModel interface

## 5 Rand.java

**Int(bound):** generates an integer in the uniformly distributed range 0 up to but not including the bound value.

**Double():** generates a double in the range 0 to 1

**Double(bound):** generates a double in the uniformly distributed range 0 up to the bound value.

**Long(bound):** generates a long in the uniformly distributed range 0 up to but not including the bound value.

**Bool():** generates a random boolean value, with equal probability of true and false.

**Binomial(n,p):** samples the binomial distribution, returns the number of heads with n weighted coin flips and probability p of heads

**Multinomial(double[]probs,n,Binomial,int[]ret,rn):** fills the return array with the number of occurrences of each event. n is the total number of occurrences to bin. Binomial is a class in the Tools folder.

**Shuffle(arr,sampleSize,numberOfShuffles,rn):** shuffles an array, sampleSize is how much of the complete array should be involved in shuffling, and numberOfShuffles is the number of entries of the array that will be shuffled. the shuffling results will always start from the beginning of the array up to numberOfShuffles.

**Gaussian(mean,stdDev,rn):** samples a gaussian with the mean and standard deviation given.

**RandomVariable(double[]probs,rn):** samples the distribution of probabilities (which should sum to 1, the SumTo1 function comes in handy here) and returns the index of the probability bin that was randomly chosen.

**RandomPointOnSphereEdge(radius,double[]ret,rn):** writes into ret the coordinates of a random point on a sphere with given radius centered on (0,0,0)

**RandomPointInSphere(radius,double[]ret,rn):** writes into ret the coordinates of a random point of inside a sphere with given radius centered on (0,0,0)

**RandomPointOnCircleEdge(radius,double[]ret,rn):** writes into ret the coordinates of a random point on the edge of a circle with given radius centered on (0,0)

**RandomPointInCircle(radius,double[]ret,rn):** writes into ret the coordinates of a random point on the edge of a circle with given radius centered on (0,0)

## 6 Gui

What fun is a model without being able to see and play with it in real time? The Gui classes allow you to easily do this and works on top of the Java Swing gui system. (except the Vis2DOpenGL and Vis3DOpenGL, which are built on lwjgl)

### 6.1 Types of Gui and Method Descriptions

Here we list the different guis that are provided by the framework and provide a summary of their functions

for a more complex look at how to use the gui system, check out the CSCCA and Polyp3D example models in the ManualModels folder.

### 6.1.1 GridWindow

The simplest built-in Gui, it is nothing more than a UIGrid imbedded in a UIWindow. Recommended for first-time users. All functions after `Close()` are also shared with the UIGrid class

**GridWindow(title,xDim,yDim,scaleFactor,main?,active?):** sets up a GridWindow, the pixel dimensions of the created window will be:  $Width = xDim * scaleFactor$  and  $Height = yDim * scaleFactor$  the main boolean specifies whether the program should exit when the window is closed. the active boolean allows easily toggling the objects on and off.

**TickPause(milliseconds):** call this once per step of your model, and the function will ensure that your model runs at the rate provided in milliseconds. the function will take the amount time between calls into account to ensure a consistent tick rate.

**IsKeyDown(char):** returns whether the given key is currently pressed

**AddKeyResponse(OnKeyDown,OnKeyUp):** takes 2 key response functions that will be called whenever a key is pressed or released

**Close():** disposes of the GridWindow.

**Clear(color):** sets all pixels to a single color.

**SetPix(x,y,color),SetPix(i,color):** sets an individual pixel on the GridWindow. in the visualization the pixel will take up  $scaleFactor * scaleFactor$  screen pixels.

**SetPix(x,y,ColorIntGenerator),SetPix(i,ColorIntGenerator):** same functionality as SetPix with a color argument, but instead takes a ColorIntGenerator function (a function that takes no arguments and returns an int). the reason to use this method is that when the gui is inactivated the ColorIntGenerator function will not be called, which saves the computation time of generating the color.

**SetString(string,xLeft,yTop,color,bkColor):** draws a string to the GridWindow. the characters in the string have length 3 pixels, and height 5 pixels, with one pixel width between characters. all characters will be drawn to the same line.

**GetPix(x,y),GetPix(i):** returns the pixel color at that location as a colorInt

**PlotSegment(x1,y1,x2,y2,color,scaleX,scaleY):** plots a line segment, connecting all pixels between the points defined by  $(x1, y1)$  and  $(x2, y2)$  with the provided color. If you are using this function on a per-timestep basis, I recommend setting individual pixels with SetPix, as it is more performant. the scaling variables adjust the spatial scale of the points.

**PlotLine(double[]xs,double[]ys,color,startPoint,endPoint,scaleX,scaleY):** plots a line by drawing segments between consecutive points. point  $i$  is defined by  $(xs[i], ys[i])$ . points are drawn starting at index startPoint, and ending at index endPoint. the scaling variables adjust the spatial scale of the points.

**PlotLine(double[]xys,color,startPoint,endPoint,scaleX,scaleY):** plots a line by drawing segments between consecutive points. points are expected in the coords format  $(x1, y1, x2, y2, x3, y3...)$ . point  $i$  is defined by  $(xys[i * 2], xys[i * 2 + 1])$ . points are drawn starting at index startPoint, and ending at index endPoint. the scaling variables adjust the spatial scale of the points.

**DrawStringSingleLine(string,xLeft,yTop,color,bkColor):** draws a string onto the GuiWindow. each string

**AddAlphaGrid(overlay):** adds another UIGrid as an overlay to compose with the main UIGrid. alpha blending will be used to combine them.

### 6.1.2 UIWindow

a container for Gui Components, which will be detailed below. the most supported of the gui types.

**UIWindow(title,main?,CloseAction,active?):** the title string is displayed in the top bar, the main boolean specifies whether the program should exit when the window is closed. the active boolean allows easily toggling the objects on and off.

**AddCol(column,component):** components are added to the gui from top to bottom in columns. when the window is displayed, the rows and columns expand to fit the largest element in each.

**RunGui():** once all components have been added, the RunGui function runs the gui and displays it to the screen.

**GetBool(label):** attempts to pull a boolean from the Param with the cooresponding label, works with the UIBoolInput

**GetInt(label):** attempts to pull an integer from the Param with the cooresponding label, works with the UIIntInput and UIComboBoxInput (returns the index of the chosen option)

**GetDouble(label):** attempts to pull a double from the Param with the cooresponding label, works with the UIIntInput and UIDoubleInput

**GetString(label):** attempts to pull a string from the Param with the cooresponding label, works with all Param types

**Dispose():** disposes of the GridWindow.

**TickPause(millis):** call this once per step of your model, and the function will ensure that your model runs at the rate provided in millis. the function will take the amount time between calls into account to ensure a consistent tick rate.

**IsKeyDown(char):** returns whether the given key is currently pressed

**AddKeyResponse(OnKeyDown,OnKeyUp):** takes 2 key response functions that will be called whenever a key is pressed or released

### 6.1.3 OpenGL2DWindow

A window for visualizing 2D models, especially off lattice ones. I usually recommend using a UIGrid instead for 2D models.

**OpenGL2DWindow(xPix,yPix,xDim,yDim,title,active?):** creates a Vis2DOpenGL window. The dimensions on screen are xPix by yPix. the xDim and yDim dimensions should match the model being drawn. the title string is displayed on the top of the window, the active boolean allows easily disabling the Vis2DOpenGL

**Update():** renders all draw commands to the window

**IsClosed():** returns true if the close button has been clicked in the Gui

**Close():** closes the Vis2DOpenGL. happens automatically when the main function finishes.

**Clear(clearColor):** usually called first, sets the screen to a color.

**Circle(x,y,z,radius,color):** Draws a circle. Currently this is the only builtin draw functionality along with the FanShape function from which it is derived.

**Line(x1,y1,x2,y2,color):** Draws a line between 2 points

**LineStrip(double[]xs,double[]ys,color):** draws a set of connected line segments, xs and ys are expected to be the same length.

**LineStrip(double[]coords,color):** draws a set of connected line segments, coords is expected to consist of [x1,y1,x2,y2...] pairs of point coordinates.

**TickPause(millis):** call this once per step of your model, and the function will ensure that your model runs at the rate provided in millis. the function will take the amount time between calls into account to ensure a consistent tick rate.

#### 6.1.4 OpenGL3DWindow

A window for visualizing 3D models.

**Vis3DOpenGL(xPix,yPix,xDim,yDim,title,active?):** creates a Vis3DOpenGL window. the dimensions on screen are xPix by yPix. the xDim, yDim, and zDim dimensions should match the model being drawn. the title string is displayed on the top of the window, the active boolean allows easily disabling the Vis3DOpenGL

**CONTROLS:** to move around inside an active Vis3DOpenGL window, click on the window, after which the following controls apply:

- **Esc Key:** Get the mouse back
- **Mouse Move:** Change look direction
- **W Key:** Move forward
- **S Key:** Move backward
- **D Key:** Move right
- **A Key:** Move left
- **Shift Key:** Move up
- **Space Key:** Move down
- **Q Key:** Temporarily increase move speed
- **E Key:** Temporarily decrease move speed

**Clear(color):** usually called before anything else: clears the gui

**Show():** push the OpenGL display to the main gui



**CheckClosed():** returns true if the close button has been clicked in the Gui

**Close():** closes the Vis3DOpenGL. happens automatically when the main function finishes.

**Circle(x,y,z,radius,color):** Draws a circle. Currently this is the only builtin draw functionality along with the FanShape function from which it is derived.

**CelSphere(x,y,z,radius,color):** Draws a cool looking cel-shaded sphere, really several Circle function calls in a row internally.

**TickPause(millis):** call this once per step of your model, and the function will ensure that your model runs at the rate provided in millis. the function will take the amount time between calls into account to ensure a consistent tick rate.

**Line(x1,y1,z1,x2,y2,z2,color):** Draws a line between 2 points

**LineStrip(double[]xs,double[]ys,double[]zs,color):** draws a set of connected line segments, xs and ys are expected to be the same length.

**LineStrip(double[]coords,color):** draws a set of connected line segments, coords is expected to consist of [x1,y1,z1,x2,y2,z2...] triplets of point coordinates.

## 6.2 Types of GuiComponent and Method Descriptions

### 6.2.1 UIGrid

a grid of pixels that are each set individually. very fast and useful for displaying the contents of grids

**UIGrid(gridW,gridH,scaleFactor,active?):** sets up a UIGrid, the pixel dimensions of the created area will be:  $Width = xDim * scaleFactor$  and  $Height = yDim * scaleFactor$ . The active boolean allows easily toggling the objects on and off. The functions that the UIGrid can execute are included above. see the GridWindow for the list of UIGrid functions

### 6.2.2 UILabel

a label that displays text on the Gui and can be continuously updated. the UILabel's on-screen size will remain fixed at whatever size is needed to render the string first passed to it.

### 6.2.3 UIButton

a button that when clicked triggers an interrupting function

### 6.2.4 UIBoolInput

a button that can be set and unset, must be labeled. use the UIWindow Param functions to interact

### 6.2.5 UIIntInput

an input line that expects an integer, must be labeled. use the UIWindow Param functions to interact

### 6.2.6 UIDoubleInput

an input line that expects a double, must be labeled. use the UIWindow Param functions to interact

### 6.2.7 UIStringInput

an input line that takes any string, must be labeled. use the UIWindow Param functions to interact

### 6.2.8 UIComboBoxInput

a dropdown menu of text options, must be labeled. use the UIWindow Param functions to interact

### 6.2.9 UIFileChooserInput

a button that when clicked triggers a gui that facilitates choosing an existing file or creating one, must be labeled. use the UIWindow Param functions to interact

## 6.3 GifMaker

the GifMaker object is used in combination with a UIGrid or GridWindow to make GIF videos of model runs.

**GifMaker(outputPath,timeBetweenFramesMS,loopContinuously?):** creates a GifMaker object. the outputPath argument specifies the name of the file that will be output. timeBetweenFramesMS specifies how many milliseconds the GIF should pause between frames. loopContinuously specifies whether the GIF should automatically restart when it finishes.

**AddFrame(visualization):** adds the current UIGrid state to the GIF as a single frame

**Close():** Close this GifMaker object

## 7 Tools

### 7.1 Tools/ FileIO

An essential piece of the framework, the FileIO class facilitates easily writing to and reading from files. this is important for collecting data from your models as well as systematically parameterizing them. the API for the FileIO object is discussed.

**FileIO(filename,mode):** the FileIO constructor expects a filename or path as a string, and a mode string, of which there are 6 options:

- **“r”** creates a FileIO in read mode, this FileIO is able to read text files
- **“w”** creates a FileIO in write mode, this FileIO is able to write to a new text file
- **“a”** creates a FileIO in append mode, this FileIO is able to append to an existing text file or write to a new file.
- **“rb”** creates a FileIO in read binary mode, this FileIO is able to read binary files

- **“wb”** creates a FileIO in write binary mode, this FileIO is able to write to binary files.
- **“ab”** creates a FileIO in append binary mode, this FileIO is able to append to an existing binary file or write to a new file.

The functions in the next sections are split up based on which mode was used to open the FileIO

**Close():** make sure to call this function when finished with the FileIO. the FileIO uses buffers internally to make writing more efficient. without calling close the buffers may never be fully written out.

### Read Mode Functions

**Read():** returns an arraylist of Strings. each string is one line from the file

**ReadLine():** returns the next line from the file as a string

**ReadLineDelimit(delimiter),ReadLineIntDelimit(delimiter):** returns an array of strings,ints,or doubles, etc. each entry is parsed using the delimiter.

**ReadDelimit(delimiter),ReadIntDelimit(delimiter):** returns an arraylist of arrays of strings,ints,or doubles, etc. each entry is parsed using the delimiter. each array is a line.

### Write Mode Functions

**Write(string):** writes the string argument to a file

**WriteDelimit(arr,delimit):** writes the contents of the provided array to a file, entries are separated using the delimiter.

### ReadBinary Mode Functions

**ReadBinBool(bool),ReadBinInt(int),ReadBinDouble(double):** read the next single value from the binary file.

**ReadBinBools(bool[]),ReadBinInts(int[]),ReadBinDoubles(double[]):** fills the array arugment with values read from the binary file.

### WriteBinary Mode Functions

**WriteBinBool(bool),WriteBinInt(int),WriteBinDouble(double):** writes a single value to the binary file.

**WriteBinBools(bool[]),WriteBinInts(int[]),WriteBinDouble(double[]):** writes every entry in the array to the binary file.

## 7.2 Tools/ MultiWellExperiment

the multiwell experiment class will visualize many models simultaneously on a single GridWindow. It can also run the models in parallel for better throughput.

**MultiWellExperiment(numWellsX,numWellsY,models[],visXdim,visYdim,borderColor,scaleFactor,StepFn,ColorFn):**

the constructor for the MultiWellExperiment class. numWellsX, numWellsY define the number of well (model) rows, models[] is an array of the starting conditions of the models. visXdim, visYdim, scaleFactor define the x pixels, y pixels, and scaling of the visualization of each model. borderColor defines the color of the separator between models, StepFn is a function argument. it takes a model, a well index, and a timestep as argument, and should update the model argument for one timestep. ColorFn is a function argument that takes a model, x, and y, and is used to set one pixel of the visualization.

**Run(numTicks,multiThread?,tickPause):** runs a multiwell experiment for a numTicks duration. if the multiThread boolean is set to true, the model execution will be multithreaded.

**Step():** runs a single timestep

**LoadWells(models[]):** sets up the multiwell experiment with a new array of models

**RunGif(numTicks,outFileName,recordPeriod,multithread?):** runs a multiwell experiment for a numTicks duration. if the multiThread boolean is set to true, the model execution will be multithreaded. saves every recordPeriod frames to a gif.

## 8 Example: Competitive Release Model

To demonstrate how the aforementioned principles and components of HAL are applied, we now look at a simple but complete example of a hybrid modeling experiment. We will be designing a model of pulsed therapy based on a publication from our lab [1]. We also showcase the flexibility that the modular component approach brings by displaying 3 different parameterizations of the same model side by side in a “multiwell experiment”.

### 8.1 Competitive Release Introduction

As in [1], the presented model assumes two competing tumor-cell phenotypes: a rapidly dividing, drug-sensitive phenotype and a slower dividing, drug-resistant phenotype. There is also a drug diffusible that enters the system through the model edges and is consumed over time by the tumor cells.

Every tick (timestep), each cell has a probability of death and a probability of division. The division probability is affected by phenotype and the availability of space. The death probability is affected by phenotype and the local drug concentration.

An interesting outcome of the experiment is that pulsed therapy is better at managing the tumor than constant therapy. The modular design of HAL allows us to test 3 different treatment conditions, each with an identical starting tumor (No drug, constant drug, and pulsed drug) Under pulsed therapy the sensitive population is kept in check, while still competing spatially with the resistant phenotype and preventing its expansion. The rest of the section describes in detail how this abstract model is generated.

Figure 2 provides a high level look at the structure of the code. Table 1 provides a quick reference for the built-in HAL functions in this example. Any functions that are used by the example but do not exist in the table are defined within the example itself and explained in detail below the code. Those fluent in Java may be able to understand the example just by reading the code and using the reference table.

Built-in framework functions and classes used in the code are highlighted in green to make identifying framework components easier

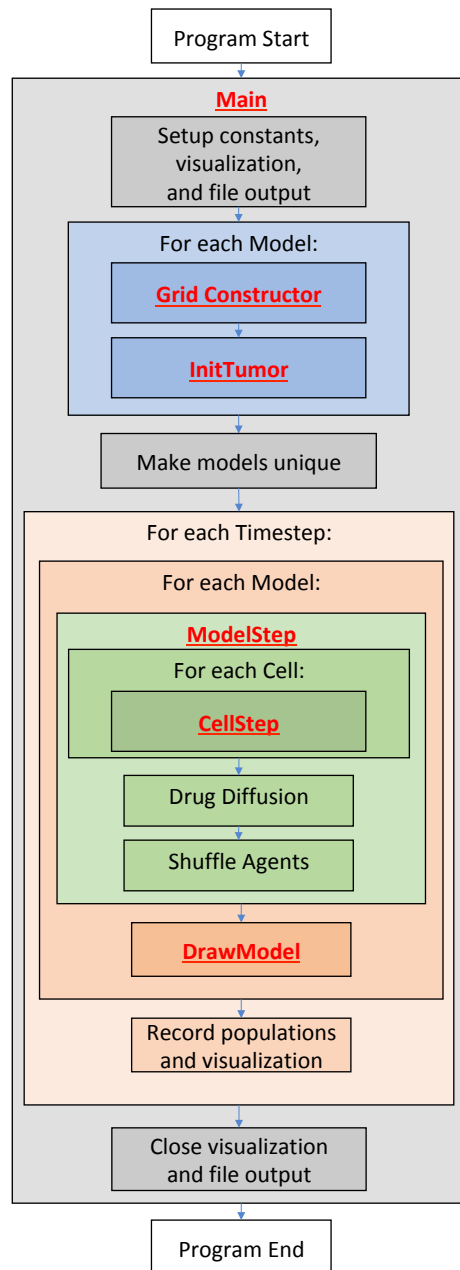


Figure 2: Example program flowchart. Yellow font indicates where functions are first called.

Function	Object	Action
MapHood( NEIGHBORHOOD, X, Y)	AgentGrid2D	Finds all indices in the provided neighborhood, centered around X,Y on the AgentGrid2D. Writes these indices into the NEIGHBORHOOD argument, and returns the number that were found.
NewAgentSQ( INDEX)	AgentGrid2D	Returns a new agent, placed at the center of the of the square at the provided INDEX.
ShuffleAgents( RNG)	AgentGrid2D	Usually called after every timestep to: 1) remove dead agents. 2) shuffle order of agents so next iteration is over a new random order. 3) increment the grid timestep.
GetTick()	AgentGrid2D	Returns the current grid timestep.
ItoX( INDEX), ItoY( INDEX)	AgentGrid2D	Converts from a grid position INDEX to the x and y components that point to the same grid position.
G	AgentSQ2D	Gets the grid that the agent occupies.
Isq()	AgentSQ2D	Gets the index of the grid square that the agent occupies.
MapEmptyHood( NEIGHBORHOOD)	AgentSQ2D	Finds all indices in the provided neighborhood, centered around the agent, that do not have an agent occupying them. Writes these indices into the HOOD argument, and returns the number that were found.
Dispose()	AgentSQ2D	Removes the agent from the grid and from iteration.
Get( INDEX)	PDEGrid2D	Returns the concentration of the PDE field at the given index.
Mul( INDEX, VALUE)	PDEGrid2D	Multiplies the concentration at the given INDEX by VALUE.
DiffusionADI( RATE)	PDEGrid2D	Applies diffusion using the ADI method with the rate constant provided. a reflective boundary is assumed.
DiffusionADI( RATE, BOUNDARY_COND)	PDEGrid2D	Applies diffusion using the ADI method with the RATE constant provided. the BOUNDARY_COND value diffuses from the grid borders.
Update()	PDEGrid2D	Applies all changes to the PDEGrid simultaneously
SetPix( INDEX, COLOR)	GridWindow	Sets the color of a pixel.
TickPause( MILLISECONDS)	GridWindow	Pauses the program between calls to TickPause. The function automatically subtracts the time between calls from MILLISECONDS to ensure a consistent framerate.
ToPNG( FILENAME)	GridWindow	Writes out the current state of the UIWindow to a PNG image file.
Close()	GridWindow	Closes the GridWindow.
RGB( RED, GREEN, BLUE)	Util	Returns an integer with the requested color in RGB format. This value can be used for visualization.
HeatMapRGB( VALUE)	Util	Maps the VALUE argument (assumed to be between 0 and 1) to a color in the heat colormap.
CircleHood( INCLUDE_ORIGIN, RADIUS)	Util	Returns a set of coordinate pairs that define the neighborhood of all squares whose centers are within the RADIUS distance of the center (0,0) origin square. The INCLUDE_ORIGIN argument specifies whether to include the origin in this set of coordinates.
MooreHood( INCLUDE_ORIGIN)	Util	Returns a set of coordinate pairs that define a Moore neighborhood around the (0,0) origin square. The INCLUDE_ORIGIN boolean specifies whether we intend to include the origin in this set of coordinates.
Write( STRING)	FileIO	Writes the STRING to the output file.
Close()	FileIO	Closes the output file.

Table 1: HAL functions used in the example. Each function is a method of a particular object, meaning that when the function is called it can readily access properties that pertain to the object that it is called from.

## 8.2 Main Function

```
1 public static void main(String[] args) {
2     //setting up starting constants and data collection
3     int x = 100, y = 100, visScale = 5, tumorRad = 10, msPause = 0;
4     double resistantProb = 0.5;
5     GridWindow win = new GridWindow("Competitive Release", x * 3, y, visScale);
6     FileIO popsOut = new FileIO("populations.csv", "w");
7     //setting up models
8     ExampleModel[] models = new ExampleModel[3];
9     for (int i = 0; i < models.length; i++) {
10         models[i] = new ExampleModel(x, y, new Rand());
11         models[i].InitTumor(tumorRad, resistantProb);
12     }
13     models[0].DRUG_DURATION = 0; //no drug
14     models[1].DRUG_DURATION = 200; //constant drug
15     //Main run loop
16     for (int tick = 0; tick < 10000; tick++) {
17         win.TickPause(msPause);
18         for (int i = 0; i < models.length; i++) {
19             models[i].ModelStep(tick);
20             models[i].DrawModel(win, i);
21         }
22         //data recording
23         popsOut.Write(models[0].Pop() + "," + models[1].Pop() + "," + models[2].Pop() +
24             "\n");
25         if ((tick) % 100 == 0) {
26             win.ToPNG("ModelsTick" + tick + ".png");
27         }
28     }
29     //closing data collection
30     popsOut.Close();
31     win.Close();
32 }
```

We first look at the main function for a bird's-eye view of how the program is structured.

Note: Source code elements highlighted in green are already built into HAL.

- 3-4:** Defines all of the constants that will be needed to setup the model and display.
- 5:** Creates a GridWindow of RGB pixels for visualization and for generating timestep PNG images.  $x*3$ ,  $y$  define the dimensions of the pixel grid.  $x$  is multiplied by 3 so that 3 models can be visualized side by side in the same window. The last argument is a scaling factor that specifies that each pixel on the grid will be viewed as a 5x5 square of pixels on the screen.
- 6:** Creates a file output object to write to a file called populations.csv
- 8:** Creates an array with 3 entries to fill in with Models.
- 9-12:** Fills the model list with models that are initialized identically. Each model will hold and update its own cells and diffusible drug. See the Grid Definition and Constructor section and the InitTumor Function section for more details.
- 13-14:** Setting the DRUG\_DURATION constant creates the only difference in the 3 models being compared. In models[0] no drug is administered (the default value of DRUG\_DURATION is 0). In models[1] drug is

administered constantly (DRUG\_DURATION is set to equal DRUG\_CYCLE). In models[2] drug will be administered periodically. See the ExampleModel Constructor and Properties section for the default values.

- 16: Executes the main loop for 10000 timesteps. See the ModelStep Function for where the Model tick is incremented.
- 17: Requires every iteration of the loop to take a minimum number of milliseconds. This slows down the execution and display of the model and makes it easier for the viewer to follow.
- 18: Loops over all models to update them.
- 19: Advances the state of the agents and diffusibles in each model by one timestep. See the Model Step Function for more details.
- 20: Draws the current state of each model to the window. See the Draw Model Function for more details.
- 23: Writes the population sizes of each model every timestep to allow the models to be compared.
- 24: Every 100 ticks, writes the state of the model as captured by the GridWindow to a PNG file.
- 29-30: After the main for loop has finished, the FileIO object and the visualization window are closed, and the program ends.

### 8.3 ExampleModel Constructor and Properties

```
1 public class ExampleModel extends AgentGrid2D<ExampleCell> {  
2     //model constants  
3     public final static int RESISTANT = RGB(0, 1, 0), SENSITIVE = RGB(0, 0, 1);  
4     public double DIV_PROB_SEN = 0.025, DIV_PROB_RES = 0.01, DEATH_PROB = 0.001,  
5         DRUG_DIFF_RATE = 2, DRUG_UPTAKE = 0.91, DRUG_TOXICITY = 0.2, DRUG_BOUNDARY_VAL  
6         = 1.0;  
7     public int DRUG_START = 400, DRUG_CYCLE = 200, DRUG_DURATION = 40;  
8     //internal model objects  
9     public PDEGrid2D drug;  
10    public Rand rng;  
11    public int[] divHood = MooreHood(false);  
12  
13    public ExampleModel(int x, int y, Rand generator) {  
14        super(x, y, ExampleCell.class);  
15        rng = generator;  
16        drug = new PDEGrid2D(x, y);  
    }
```

This section covers how the grid is defined and instantiated.

- 1: The ExampleModel class, which is user defined and specific to this example, is built by extending the generic AgentGrid2D class. The extended grid class requires an agent type parameter, which is the type of agent that will live on the grid. To meet this requirement, the <ExampleCell> type parameter is added to the declaration.
- 3: Defines RESISTANT and SENSITIVE constants, which are created by the Util RGB function. These constants serve as both colors for drawing and as labels for the different cell types.



- 4-5: Defines all constants that will be needed during the model run. These values can be reassigned after model creation to facilitate testing different parameter settings. In the main function, the DRUG\_DURATION variable is modified for the Constant-Drug, and Pulsed Therapy experiment cases.
- 8: Declares that the model will contain a PDEGrid2D, which will hold the drug concentrations. The PDEGrid2D can only be initialized when the x and y dimensions of the model are known, which is why we do not define them until the constructor function.
- 9: Declares that the Grid will contain a Random number generator, but take it in as a constructor argument to allow the modeler to seed it if desired.
- 10: Defines an array that will store the coordinates of a neighborhood generated by the MooreHood function. The MooreHood function generates a set of coordinates that define the Moore Neighborhood, centered around the (0,0) origin. The neighborhood is stored in the format  $[0_1 0_2, \dots, 0_n, x_1, y_1, x_2, y_2, \dots, x_n, y_n]$ . The leading zeros are written to when MapHood is called, and will store the indices that the neighborhood maps to. See the CellStep function for more information.
- 12: Defines the model constructor, which takes as arguments the x and y dimensions of the world and a Random number generator.
- 13: Calls the AgentGrid2D constructor with super, passing it the x and y dimensions of the world, and the ExampleCell Class. This Class is used by the Grid to generate a new cell when the NewAgentSQ function is called.
- 14-15: The random number generator argument is assigned and the drug PDEGrid2D is defined with matching dimensions.

## 8.4 InitTumor Function

---

```

1  public void InitTumor(int radius, double resistantProb) {
2      //get a list of indices that fill a circle at the center of the grid
3      int[] tumorNeighborhood = CircleHood(true, radius);
4      int hoodSize = MapHood(tumorNeighborhood, xDim / 2, yDim / 2);
5      for (int i = 0; i < hoodSize; i++) {
6          if (rng.Double() < resistantProb) {
7              NewAgentSQ(tumorNeighborhood[i]).type = RESISTANT;
8          } else {
9              NewAgentSQ(tumorNeighborhood[i]).type = SENSITIVE;
10         }
11     }
12 }

```

---

The next segment of code is a function from the ExampleModel class that defines how the tumor is first seeded after the ExampleModel is created.

- 1: The arguments passed to InitTumor function are the approximate radius of the circular tumor being created and the probability that each created cell will be of the resistant phenotype.
- 3: Sets the circleCoords array using the built-in CircleHood function, which stores coordinates in the form  $[0_1, 0_2, \dots, 0_n, x_1, y_1, x_2, y_2, \dots, x_n, y_n]$ . These coordinate pairs define a neighborhood of all squares whose centers are within the radius distance of the center (0,0) origin square. The leading 0s are used by the

MapHood function to store the mapping indices. The boolean argument specifies that the origin will be included in this set of squares, thus making a completely filled circle of squares.

- 4: Uses the built-in MapHood function to map the neighborhood defined above to be centered around  $xDim/2, yDim/2$  (the dimensions of the AgentGrid). The results of the mapping are written as indices to the beginning of the tumorNeighborhood array. MapHood returns the number of valid indices found, and this will be the size of the starting population.
- 5: Loops from 0 to hoodSize, allowing access to each mapped index in the tumorNeighborhood.
- 6: Samples a random number in the range  $(0 - 1]$  and compares to the resistantProb argument to set whether the cell should have the resistant phenotype or the sensitive phenotype.
- 7-9: Uses the built-in NewAgentSQ function to place a new cell at each tumorNeighborhood position. In the same line we also specify that the type should be either resistant or sensitive, depending on the result of the rng.Double() call.

## 8.5 ModelStep Function

---

```

1  public void ModelStep(int tick) {
2      ShuffleAgents(rng);
3      for (ExampleCell cell : this) {
4          cell.CellStep();
5      }
6      int periodTick = (tick - DRUG_START) % DRUG_CYCLE;
7      if (periodTick > 0 && periodTick < DRUG_DURATION) {
8          //drug will enter through boundaries
9          drug.DiffusionADI(DRUG_DIFF_RATE, DRUG_BOUNDARY_VAL);
10     } else {
11         //drug will not enter through boundaries
12         drug.DiffusionADI(DRUG_DIFF_RATE);
13     }
14     drug.Update();
15 }

```

---

This section looks at the main step function which is executed once per tick by the Model.

- 2: The ShuffleAgents function randomizes the order of iteration so that the agents are always looped through in random order.
- 3-4: Iterates over every cell on the grid, and calls the CellStep function on every cell.
- 6: The GetTick function is a built-in function that returns the current Grid tick. The If statement logic checks if the tick is past the drug start and if the tick is in the right part of the drug cycle to apply drug. (See the Grid Definition and Constructor section for the values of the constants involved, the DRUG\_DURATION variable is set differently for each model in the Main Function)
- 7-9: If it is time to add drug to the model, the built-in DiffusionADI function is called. The default Diffusion function uses the standard 2D Laplacian and is of the form:  $\frac{\delta C}{\delta t} = D \nabla^2 C$ , where D in this case is the DRUG\_DIFF\_RATE. DiffusionADI uses the ADI method which is more stable and allows us to take larger steps than the 2D Laplacian can support. The additional argument to the DiffusionADI equation specifies the boundary condition value DRUG\_BOUNDARY\_VAL. This causes the drug to diffuse into the PDEGrid2D from the boundary.

12: Without the second argument the DiffusionADI function assumes a reflective boundary, meaning that drug concentration cannot escape or enter through the sides. Therefore the only way for the drug concentration to decrease is via consumption by the Cells. See the CellStep function section, line 6, for more information.

14: Update is called to apply the reaction and diffusion changes to the PDEGrid

## 8.6 CellStep Function and Cell Properties

---

```

1  class ExampleCell extends AgentSQ2Dunstackable<ExampleModel> {
2      public int type;
3
4      public void CellStep() {
5          //Consumption of Drug
6          G.drug.Mul(Isq(), G.DRUG_UPTAKE);
7          double deathProb, divProb;
8          //Chance of Death, depends on resistance and drug concentration
9          if (this.type == RESISTANT) {
10             deathProb = G.DEATH_PROB;
11         } else {
12             deathProb = G.DEATH_PROB + G.drug.Get(Isq()) * G.DRUG_TOXICITY;
13         }
14         if (G.rng.Double() < deathProb) {
15             Dispose();
16             return;
17         }
18         //Chance of Division, depends on resistance
19         if (this.type == RESISTANT) {
20             divProb = G.DIV_PROB_RES;
21         } else {
22             divProb = G.DIV_PROB_SEN;
23         }
24         if (G.rng.Double() < divProb) {
25             int options = MapEmptyHood(G.divHood);
26             if (options > 0) {
27                 G.NewAgentSQ(G.divHood[G.rng.Int(options)]).type = this.type;
28             }
29         }
30     }
31 }

```

---

We next look at how the Agent is defined and at the CellStep function that runs once per Cell per tick.

- 1: The ExampleCell class is built by extending the generic AgentSQ2Dunstackable class. The extended Agent class requires the ExampleModel class as a type argument, which is the type of Grid that the Agent will live on. To meet this requirement, we add the <ExampleModel> type parameter to the extension.
- 2: Defines an internal int called type. each Cell holds a value for this field. If the value is RESISTANT, the Cell is of the resistant phenotype, if the value is SENSITIVE, the cell is of the sensitive phenotype. The RESISTANT and SENSITIVE constants are defined in the ExampleGrid as constants.
- 6: The G property is used to access the ExampleGrid object that the Cell lives on. G is used often with Agent functions as the AgentGrid is expected to contain any information that is not local to the Cell itself. Here it is used to get the drug PDEGrid2D. The drug concentration at the index that the Cell is currently occupying (Isq()) is then multiplied by the drug uptake constant, thus modeling local drug consumption by the Cell.

- 7:** Defines deathProb and divProb variables, these will be assigned different values depending on whether the ExampleCell is RESISTANT or SENSITIVE.
- 9-12:** If the cell is resistant the deathProb variable is set to the DEATH\_PROB value alone, if the cell is sensitive, an additional term is added to account for the probability of the cell dying from drug exposure, using the concentration of drug at the cell's position (lsq())
- 14-16:** Samples a random number in the range (0 – 1] and compares to deathProb to determine whether the cell will die. If so, the built-in agent Dispose() function is called, which removes the agent from the grid, and then return is called so that the dead cell will not divide.
- 19-22:** Sets the divProb variable to either DIV\_PROB\_RES for resistant cells, or DIV\_PROB\_SEN for sensitive cells.
- 24:** Samples a random number in the range (0 – 1] and compare to divProb to determine whether the cell will divide.
- 25:** If the cell will divide, the built-in MapEmptyHood function is used which displaces the divHood (the moore neighborhood as defined in the Grid Definition and Constructor section) to be centered around the x and y coordinates of the Cell, and writes the empty indices into the neighborhood. The MapEmptyHood function will only map indices in the neighborhood that are empty. MapEmptyHood returns the number of valid division options found.
- 26-27:** If there are one or more valid division options, a new daughter cell is created using the builtin NewAgentSQ function and its starting location is chosen by randomly sampling the divHood array to pull out one if its valid locations. Finally with the .type=this.type statement, the phenotype of the new daughter cell is set to the phenotype of the pre-existing daughter that remains in place, thus maintaining phenotypic inheritance.

## 8.7 DrawModel Function

---

```

1  public void DrawModel(GridWindow vis, int iModel) {
2      for (int x = 0; x < xDim; x++) {
3          for (int y = 0; y < yDim; y++) {
4              ExampleCell drawMe = GetAgent(x, y);
5              if (drawMe != null) {
6                  vis.SetPix(x + iModel * xDim, y, drawMe.type);
7              } else {
8                  vis.SetPix(x + iModel * xDim, y, HeatMapRGB(drug.Get(x, y)));
9              }
10         }
11     }
12 }

```

---

We next look at the DrawModel Function, which is used to display a summary of the model state on a GridWindow object. DrawModel is called once for each model per timestep, see the Main Function section for more information.

- 2-3:** Loops over every lattice position of the grid being drawn, xDim and yDim refer to the dimensions of the model.
- 4:** Uses the built-in GetAgent function to get the Cell that is at the x,y position.

**5-6:** If a cell exists at the requested position, the corresponding pixel on the GridWindow is set to the cell's phenotype color. To draw the models side by side, the pixel being drawn is displaced to the right by the model index.

**7-8:** If there is no cell to draw, then the pixel color is set based on the drug concentration at the same index, using the built-in heat colormap.

## 8.8 Imports

---

```
1 package Examples._6CompetitiveRelease;
2 import Framework.GridsAndAgents.AgentGrid2D;
3 import Framework.GridsAndAgents.PDEGrid2D;
4 import Framework.Gui.GridWindow;
5 import Framework.GridsAndAgents.AgentSQ2Dunstackable;
6 import Framework.Gui.UIGrid;
7 import Framework.Tools.FileIO;
8 import Framework.Rand;
9 import static Examples._6CompetitiveRelease.ExampleModel.*;
10 import static Framework.Util.*;
```

---

The final code snippet looks at the imports that are needed. Any modern Java IDE should generate import statements automatically.

**1:** The package statement is always needed and specifies where the file exists in the larger project structure

**2-8:** Imports all of the classes that we will need for the program.

**9:** Imports the static fields of the model so that we can use the type names defined there in the Agent class.

**10:** Imports the static functions of the Util file, which adds all of the Util functions to the current namespace, so we can natively call them. Statically importing Util is recommended for every project.

## 8.9 Model Results

Table 2 displays the model visualization at tick 0, tick 400, tick 1100, tick 5500, and tick 10,000. The Figure caption explores the notable trends visible in each image. Figure 3 displays the population sizes as recorded by the FileIO object at the end of every timestep.

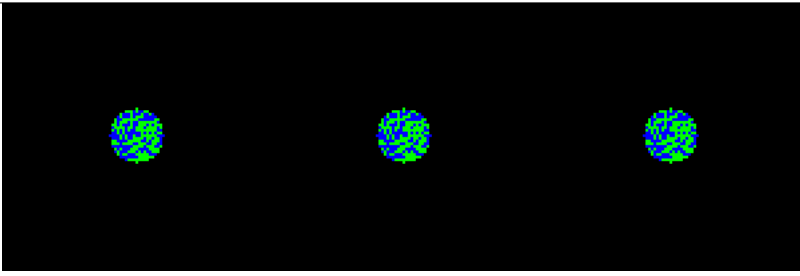
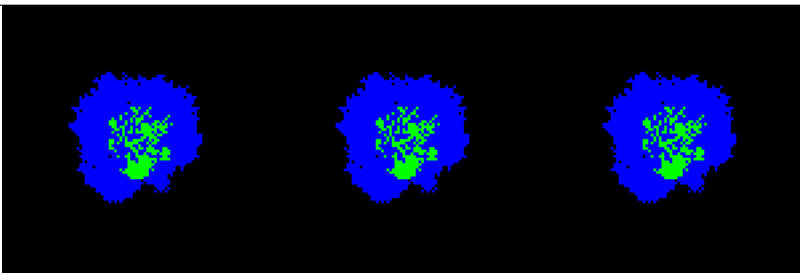
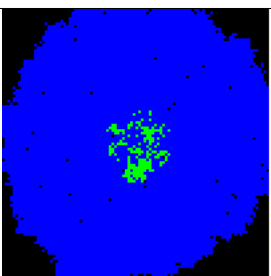
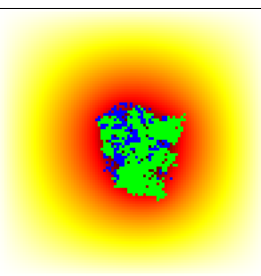
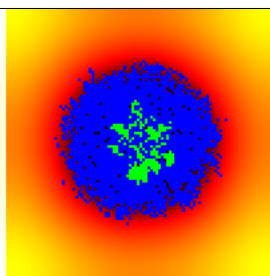
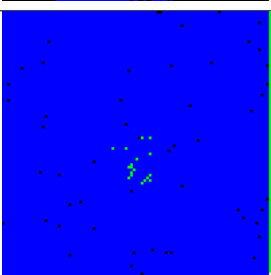
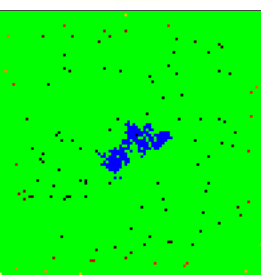
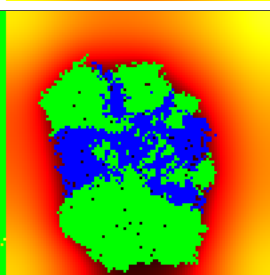
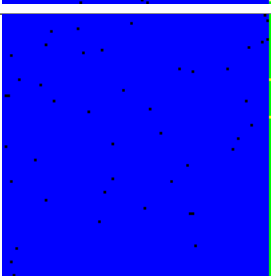
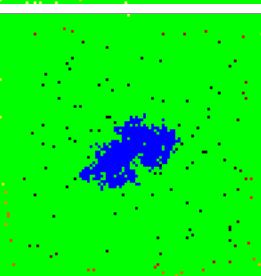
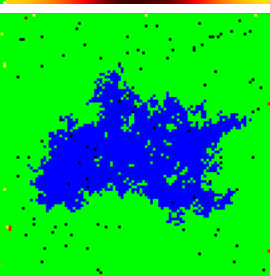
Timestep	No Drug / Constant Drug / Pulsed Drug		
0			
400			
1100			
5500			
10000			

Table 2: Selected model visualization PNGs. Blue cells are drug sensitive, Green cells are drug resistant, background heatmap colors show drug concentration. At timestep 0 and timestep 400 (right before drug application starts), all 3 models are identical. At tick 1100 the differences in treatment application show different effects: when no drug is applied, the rapidly dividing sensitive cells quickly fill the domain, when drug is applied constantly, the resistant cells overtake the tumor. Pulsed drug kills some sensitive cells, but leaves enough alive to prevent growth of the resistant cells. At tick 5500, the resistant cells have begun to emerge from the center of the pulsed drug model. At tick 10000, all domains are filled. Interestingly, the sensitive cells are able to survive in the center of the domain because drug is consumed by cells on the outside. This creates a drug-free zone in which the sensitive cells outcompete the resistant cells.

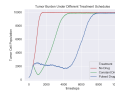


Figure 3: FileIO population output. This plot summarizes the changes in tumor burden over time for each model. This plot was constructed in R using data accumulated in the program output csv file. Displayed using GGPlot in R

This modeling example illustrates the power of HAL's approach to model building. Writing relatively little complex code, we setup a 3 model experiment with nontrivial dynamics along with methods to collect data and visualize the models. We now briefly review the model results.

As can be seen in Figure 3 and Table 2, the pulsed therapy is the most effective at preventing tumor growth, however the resistant cells ultimately succeed in breaking out of the tumor center and outcompeting the sensitive cells on the fringes of the tumor. It may be possible to maintain a homeostatic population of sensitive and resistant cells for longer by using a different pulsing schedule or possibly by using adaptive therapy. As the presented model is primarily an example, we do not explore how to improve treatment further. For a more detailed exploration of the potential of adaptive therapy for prolonging competitive release, see [1].

## References

- [1] Jill A Gallaher, Pedro M Enriquez-Navas, Kimberly A Luddy, Robert A Gatenby, and Alexander RA Anderson. Adaptive vs continuous cancer therapy: Exploiting space and trade-offs in drug scheduling.

## Acknowledgements

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