BiMat: Start Guide

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1 Description

1.1 Main Goal

The main goal of BiMat is to facilitate the analysis of nestedness and modularity of bipartite ecological networks.

1.2 System Requirements

- MATLAB® 2009b or superior. The software may work in previous versions, however we could not test in those versions.
- The user is expected to have basic MATLAB® knowledge.
- MATLAB® Parallel Computing Toolbox in case you want to work with the parallel BiMat version.

1.3 Functionality

BiMat is a MATLAB[®] library whose main function is the analysis of modularity and nestedness in bipartite ecological networks. Its main features are:

- Modularity and nestedness calculation.
- Diversity calculation using Shannon and Simpson's indexes.
- Different null models for the creation of random bipartite networks.
- Statistics of the network.
- Internal statistics of the modules (multi-scale analysis).
- Group statistical analysis (analysis of many networks).
- Plotting in matrix or graph layouts.

1.4 Workflow

The workflow of the BiMat package can be visualized in Figure 1.

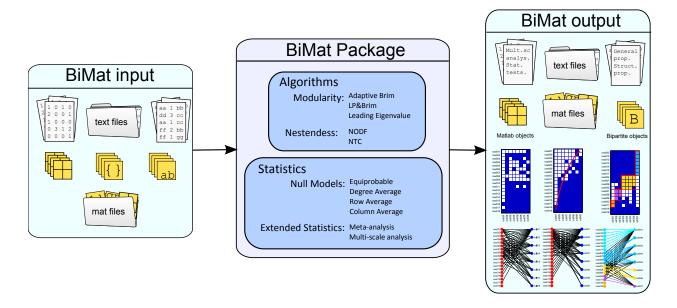


Figure 1: BiMat Workflow. The figure shows the main scheme of the BiMat package. BiMat can take matlab objects or text files as main input. The input is analyzed mainly around modularity and nestedness using a variety of null models. The user may also perform an additional multi-scale analysis on the data, or if he have more than one matrix to perform a meta-analysis in the entire data. Finally, the user can observe the results via matlab objects, text files, and plots.

2 Installation

2.1 Downloading BiMat

BiMat comes in two versions: (i) un-parallel and (ii) parallel version. While both have exactly the same functions, the last one has been coded for using more than one cpu core for performing the statistical tests, and therefore increase the speed of simulations. If you chose to use the parallel version be sure that the user have the Parallel Computing Toolbox installed in MATLAB[®] . The user can check if he has this toolbox by running in the MATLAB[®] command line:

>>ver

After entering this command, MATLAB® will display information about the version of MATLAB® the user is running, including a list of all toolboxes installed on the user's system.

No matter the version the user want to use, both versions can be downloaded from our group website: http://ecotheory.biology.gatech.edu/downloads. Equally possible, the user can download the last available version from the main developer GitHub website: https://github.com/cesar7f/BiMat.

2.2 Installing BiMat and adding it to the MATLAB® path

To install BiMat, copy the downloaded zip file to a directory of interest and unzip it. Next, you will need to add BiMat to the MATLAB® path either temporally or permanently:

• Temporal path: Add the BiMat directory (and sub-directories) to the MATLAB® path. You can do that by typing in the MATLAB® command line:

```
>>g=genpath('bimat_directory_location');
>>addpath(g);
```

You should replace bimat_directory_location with the full path to the directory in which you installed BiMat.

• **Permanent path:** Alternatively, the user can update permanently (also temporally) by accessing the MATLAB® path configuration. The path configuration can be accessed via menu File -> Set Path.

2.3 BiMat configuration: Options.m file

Most of the BiMat functions can work without the need of parameters by the user. However, if the user does not specify the required arguments, BiMat will assume that default values will be used. These default values are specified on the file main/Options.m that the user can modified according to his needs. A description of each parameter with its default value is indicated below:

- Statistical Significance: A two-tail test is the default way of testing for significance in BiMat . Notice that the user can perform a one-tail test by just duplicating the values below:
 - P_VALUE = 0.05: The p-value for testing statistical significance using a percentile test approach. Anything above the percentil 100*(1-p/2) will be significant, while anything below the percentile 100*(p/2) will be anti-significant.
 - Z_SCORE = 1.96: The z-score for testing statistical significance using a z-test approach. Anything above |z| will be considered significant, while anything below -|z| will be considered antisignificant. z = 1.96 has been chosen in order to correspond to p = 0.05.
- Null Models:

- DEFAULT_NULL_MODEL = @NullModels.EQUIPROBABLE: The default function for creating random networks (see corresponding section for a description of each null model).
- ALLOW_ISOLATED_NODES = true: When the network is sparse, a random network may be created with nodes with no links at all (matrix with empty rows or columns). BiMat by default allow this kind of random networks for performing the statistical test. However, the user may want to change this value to false and like this avoid the creation of this kind of random networks. However, the user must be aware that the time required for creating a random network without empty nodes will growth with the sparsity of the matrix.
- TRIALS_FOR_NON_EMPTY_NODES = 1000: This value is only used when the user changes the value of the previous parameter to false. In some extreme cases (a very sparse network), BiMat will not be able to find a random network without empty nodes. Hence, in order to avoid infinite loops, BiMat will stop looking for them after the number of trials specified in this parameter. If BiMat can not create a random network without empty nodes before this number of trials, BiMat will just create a random network without this constraint and will print the next message in the MATLAB® command line:

Warning: Not possible to create a matrix with non isolated nodes.

The random matrix was created without this constraint instead.

Consider to modify Options.ALLOW_ISOLATED_NODES and/or Options.INCLUDE_EMPTY_NODES

- INCLUDE_EMPTY_NODES = true: Sometimes the user may have data with empty nodes (a matrix with empty rows and/or columns). Depending on the value of this parameter BiMat will chose between keeping these nodes (true) or deleting them from the adjacency matrix (false). Further, the user must be aware that including or not empty nodes will have an effect during the statistical tests of his data.
- REPLICATES = 100: The amount of replicates that BiMat performs in order to test for statistical significance. The value of 100 was chosen with the idea of getting quick results. However, the user must be aware that this value is no appropriate for accurate testing. The right value will depend on the kind of network (or networks) that the user is analyzing. It will depend mostly in two quantities: the fill and the size of the adjacency matrix. Experience from the developers indicate that if matrices are small $\sim 10 \times 10$ the appropriate number is $\sim 10,000$, while for big matrices $\sim 200 \times 200$, the appropriate number is $\sim 1,000$. However, the right way for testing the appropriate value is by looking and how the variance decrease as the number of replicates increase. The variance stops decreasing considerably with the number of replicates, increasing this last number does not have any effect on the statistical results.
- Algorithms: All the next parameters refer to the modularity algorithms behavior. The user can change the values here, or he can change the parameters dynamically by modifying the corresponding properties in the BipartiteModularity instance.
 - OPTIMIZE_COMPONENTS = false: Modularity is a function that depends in the global information of the network. However, sometimes, the user may have a network which is not connected (it has isolated components). By using the default value false, BiMat will optimize the modularity value at the entire adjacency matrix, while by using the value true, BiMat will optimize the modularity at the component level. Optimizing at the component level may decrease the global modularity value, thought the modularity configuration may be more appropriate for the user.
 - MODULARITY_ALGORITHM = @AdaptiveBrim: BiMat has three algorithms for optimizing the modularity equation and hence find the module configuration of the network. See the corresponding section for a description of these algorithms.
 - TRIALS_MODULARITY = 20: The results of the modularity algorithms depends strongly in some initial random assignment of the communities. Therefore, BiMat restart the algorithm using this amount of times.

2.4 Getting help

At any moment you can access help from the command line using any of the next commands:

- help class_name: For a summary of the class file (i.e. help StatisticalTest).
- help class_name.method_name: For a summary of what the methods does and what kind of arguments it gets (i.e. help StatisticalTest.DoNulls).
- help class_name.property_name: For a summary of the property (i.e. help StatisticalTest.replicates).

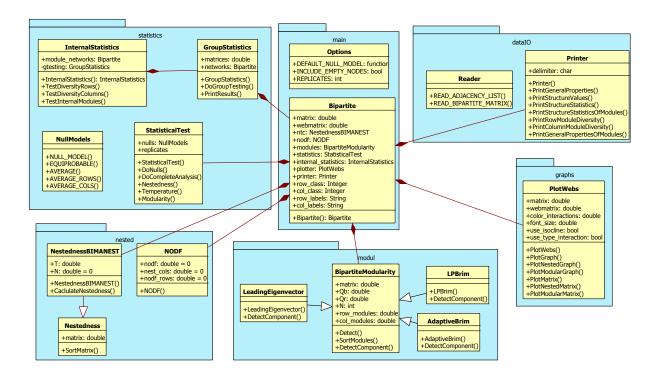


Figure 2: Class diagram of the BiMat package. This diagram shows the main classes of the BiMat package together with the relationships among them. Bipartite class is the main one inside the package and it is basically a bridge between all the functionality inside BiMat.

3 Main Modules

BiMat has been coded using an Object-Oriented-Programming approach. The main BiMat modules with their most important functions are show as class diagram in Figure 2.

In this section we will describe with some detail some of the important packages, what they do and how to work with them.

3.1 Bipartite class

Notice in Figure 2 that the class Bipartite is the central class of BiMat package. Its main purpose in the a BiMat life cycle analysis is to work as a bridge between all the BiMat modules. However, most of the modules can work independent of this class. For example, if the user is only interested in getting modularity and/or nested values of an adjacency matrix, he can just focus in on the algorithm classes. However, as the need of the user increases (e.g. need for algorithms, statistical tests, plotting) the best way is to work with a Bipartite instance. Hence, the user will need to create a instance of this class in order to perform all the statistics and plotting that he need to perform. There is two ways of creating a Bipartite instance that are described in the next subsubsections.

3.1.1 Creating a Bipartite instance from MATLAB® objects

I order to create an instance of the bimat class the user need as minimum a MATLAB[®] matrix that represents the bipartite adjacency matrix of the network he will work with. Optionally, the user may want to use two MATLAB[®] string cell arrays that will represent labels of row and column nodes, and two MATLAB[®] vectors

that will represent the classification id's for row and columns nodes. Assuming that the user has specified the these values in the variables matrix, row_labels, col_labels, row_classes, col_classes respectively, the next two ways will create the same instance (or object).

```
>>bip = Bipartite(matrix,'my network',row_labels,col_labels,row_classes,col_classes);
>>bip = Bipartite(matrix,'my network'); % The string is optional
>>bip.row_labels = row_labels; bip.col_labels = col_labels;
>>bip.row_class = row_classes; bip.col_class = col_classes;
```

The user must be aware that the way that matrix can contain any natural number, but that only the Boolean version of the matrix will be used during the algorithm calculations. However, number different than 1 and 0 may be used during plotting for differentiating categorical levels of interactions. In other words, BiMat can not deal with real adjacency matrices. Therefore, the user must either prepare an integer or boolean version of his data. See Example I for more details.

3.1.2 Creating a Bipartite instance from text files

Sometimes the available data for the user may be encoded in text files only. The user can still create Bipartite instance with the help of the static class Reader:

If the user does not have his data already encoded in a MATLAB® matrix, he can make use of the static class Reading which make use of two functions for reading the data directly from text files using two type of formats:

• bip = Reader.READ_BIPARTITE_MATRIX(filename): Each row in the file represent a different row of the bipartite matrix, such that spaces will separate column values. Below we show an example of how to create this type of data. All values different from 0 are counted as interactions. As already metionend, the algorithms will use the boolean version of the data and numbers different from 1 and 0 are only used for plotting:

```
1 0 0 2 0 0 0
1 2 0 0 0 2 1
1 1 0 0 1 2 1
1 2 3 0 0 1 1
2 1 1 1 0 0 0
```

Because no labels for row and column nodes can be specified by the user using this function, BiMat will use the default values: 'row_1', 'row_2', 'row_3',..., 'col_1', 'col_2', 'col_3',...

• bip =eader.READ_ADJACENCY_LIST(filename): The text file must have at least two columns separated by an space, such that the first and second column represent nodes from the two different entities that are interacting, respectively. In other words, a row for interaction must exist in the text file. This format can accept an extra column (in the middle of the previous ones) to indicate the strength of the interaction. Again, only boolean values are used in the algorithms, but integer values could be used for representing categorical interaction levels during plotting. Below we show an example of this kind of file (notice that in this version is possible to include labels but not classification id's for nodes)

```
row_label_1 1 col_label_1
row_label_1 1 col_label_2
row_label_1 2 col_label_3
row_label_3 1 col_label_2
row_label_3 3 col_label_1
row_label_2 3 col_label_2
```

As the user may already notice, currently BiMat can not read the classification's ids directly from text files. This functionality will be added in a future release. At the moment the user may want to read this data using any of the MATLAB® text file functions.

4 Examples

This section include three different examples to introduce the user to the main features of BiMat . All the code and data file can be found on the examples directory.

- creating_networks.m. It shows and explains the required input for BiMat .
- moebus_study.m. An analysis of the Moebus phage-bacteria bipartite network. It shows how to use the most important functions that are available to analyze a single matrix. This analysis include how to calculate most of the results published on [14].
- group_matrices.m. An analysis of a group of matrices that shows how to perform an analysis in many matrices at the same time. This example reproduce some of the results published on [13]. However, using this template all the results can be reproduced with a little extra effort.

4.1 BiMat - Creating networks

This example will introduce the user to the input of BiMat . It explains what input is required and how it is used by BiMat . This example is located on examples/creating_networks.m and make use of examples/input_adja.txt and examples/input_matrix.txt files.

4.1.1 Contents

- Add the source to the MATLAB® path
- Bipartite class and main input
- Optional input
- Creating input for Bipartite class
- Creating a Bipartite object from MATLAB® data
- Creating a Bipartite object from text files

4.1.2 Add the source to the MATLAB® path

```
5 %% Add the source to the matlab path
6 %Assuming that you run this script from examples directory
7 g = genpath('../'); addpath(g);
8 close all;
```

4.1.3 Bipartite class (main class)

The Bipartite is the fundamental class of the BiMat software. This class works as a communication bridge between all the available classes. Therefore, in order to work with BiMat we will always need to instantiate at least an object of this class.

4.1.4 Required input

The required input of the Bipartite class is a MATLAB® matrix, where the rows will represent the node set R and the columns the node set C, such that if the element matrix(i,j)>0 a link between node r_i and c_j exist. This matrix input can contain only non-negative integers $\{0,1,2,3...\}$. However, at present, values greater than 1 are only used for plotting purposes (e.g. color interactions according to weight) and not in the existing algorithms (which only work using the boolean version of the matrix).

4.1.5 Optional input

BiMat has two different types of optional input. The first type is for node labeling and the main use of it will be for labeling row and column nodes during plotting. The input must be encoded in a cell of strings for each set R and C nodes, such that each string in a cell corresponds to the label of a node. The size of such cells must corresponds to the number of nodes.

The second type of input consist of the type of node for either row and column nodes. For an example of type of nodes consider a bipartite network where R and C represent pollinators and plants respectively. In turn pollinators can be classified in birds and insects, which will be the classification for set R. The information of this classification is useful to explain modularity in terms of node classification. You can consult the Moebus study example for additional details. The classification input must be vectors of the same size than the number of nodes in rows and columns. The values must be positive integers $\{1, 2, 3, ...\}$ that represents the classification class of each node.

4.1.6 Creating input for Bipartite class

Here will show an example of the simplest way of creating a Bipartite object. We will create a bipartite networks using a MATLAB[®] matrix as input of the Bipartite object. This synthetic data matrix represents the interactions between a set of pollinators (rows) and a set of plants (columns). matrix(i,j)>0 means that pollinator i pollinates plant j with strength matrix(i,j).

```
%Creating the data
   matrix = [2 \ 0 \ 2 \ 2; ...
             1 2 2 1;...
48
             2 0 0 2;...
49
             0 1 2 2;...
50
             0 0 1 0];
52
  % For the next variables observe that the size of matrix 5x4 correlates with
53
  row_labels = {'insect 1', 'insect 2', 'insect 3', 'bird 1', 'bird 2'};
  col_labels = {'flower 1', 'flower 2', 'grass 1', 'gras 2'};
   %Notice that as long as each kind is represented by a differented positive
  %integer you will be fine.
   row_ids = [1 1 1 3 3];
58
   %Notice that 1 in col_ids not necessearly corresponds to 1's in row_ids.
  col_ids = [1 1 5 5];
```

4.1.7 Creating a Bipartite object from MATLAB® data

Using the data we just created we can now create our Bipartite object:

```
64 bp = Bipartite(matrix);
65 bp.row_labels = row_labels;
66 bp.col_labels = col_labels;
```

```
67 bp.row_class = row_ids;
68 bp.col_class = col_ids;
```

4.1.8 Creating a Bipartite object from text files

An additional way of creating data is by using the static functions from the Reading.m class. Currently two different formats are available. The first input format will contain only the information of the adjacency matrix (you will need to add row/column labels and classification id's if you need). A file example for creating the last data is on examples/input_matrix.txt, which contains:

```
2 0 2 2
1 2 2 1
2 0 0 2
0 1 2 2
0 0 1 0
```

The last format input can be called using:

```
bp = Reading.READ_BIPARTITE_MATRIX('input_matrix.txt');
s5 % We need to add labels and classification ids by ourselves
bp.row.labels = row.labels;
s7 bp.col.labels = col.labels;
s8 bp.row.class = row.ids;
s9 bp.col.class = col.ids;
```

The second input format consist on writing the adjacency list. This input format will read also the row and column node labels. However if you need ids for the classification you will need to add by yourself. An example for the last data format is located on examples/input_adja.txt and is shown below:

```
insect_1 2 flower_1
insect_1 2 grass_1
insect_1 2 grass_2
insect_2 1 flower_1
insect_2 2 flower_2
insect_2 2 grass_1
insect_2 1 grass_2
insect_3 2 flower_1
insect_3 2 grass_2
bird_1 1 flower_2
bird_1 2 grass_1
bird_1 2 grass_2
bird_2 1 grass_1
```

The middle column is optional. If it is not used, the reading function will assume that is composed of ones only. We can now just call:

```
bp = Reading.READ_ADJACENCY_LIST('input_adja.txt.');

113 % Wee need to add classification ids by ourselves
114 bp.row_class = row_ids;
115 bp.col_class = col_ids;
```

Now that you know how to create a network object, you can proceed to the next example that shows how to perform a complete analysis in a bipartite network.

4.2 BiMat Use case using Moebus cross-infection matrix data

This example will introduce the user to the most basic features of the BiMat Software. In order to do that we will calculate some of the results presented on the Flores et al 2012 paper (Multi-scale structure and geographic drivers of cross-infection within marine bacteria and phages) [14]. We will show how to plot, evaluate modularity and nestedness, and perform some statistics at the global and internal modular structure.

This example is located on examples/moebus_study.m and makes use of examples/moebus_use_case.mat data file.

4.2.1 Contents

- Add the source to the MATLAB® path
- Creating the Bipartite network object
- Calculating Modularity
- Calculating Nestedness
- Plotting in Matrix Layout
- Statistical analysis in the entire network
- Statistical analysis of the internal modules

4.2.2 Add the source to the MATLAB® path

```
10 %Assuming that you run this script from examples directory
11 g = genpath('../'); addpath(g);
12 close all; %Close any open figure
```

We need also to load the data from which we will be working on:

```
15 load moebus_use_case.mat;
```

load moebus_use_case.mat;

The loaded data contains the bipartite adjacency matrix of the Moebus and Nattkemper study [22], where 1's and 2's in the matrix represent either clear or turbid lysis spots. It also contains the labels for both bacteria and phages and their geographical location from which they were isolated across the Atlantic Ocean.

4.2.3 Creating the Bipartite network object

We can print the general properties of the network with:

```
30 bp.printer.PrintGeneralProperties();
```

```
General Properties
```

```
Number of species: 501

Number of row species: 286

Number of column species: 215

Number of Interactions: 1332

Size: 61490

Connectance or fill: 0.022
```

4.2.4 Calculating Modularity

The modularity algorithm is encoded in the property modules of the Bipartite object (bp.modules). Tree algorithms are available:

- 1. Adaptive BRIM (AdaptiveBrim.m)
- 2. LP&BRIM (LPBrim.m)
- 3. Leading Eigenvector (NewmanAlgorithm.m)

Each algorithm optimizes the same modularity equation [3] for bipartite networks using different approaches. Only the Newman algorithm may return the same result. The other two perform at some point random module pre-assignments, and by consequence they may return the same result in each call. The default algorithm is specified on Options.MODULARITY_ALGORITHM. However, we can assign another algorithm dynamically. Here, for example, we will use the Newman's algorithm (Leading eigenvector):

```
bp.modules = NewmanModularity(bp.matrix);

48 % The next flag is exclusive of Newman Algorithm and what it does is to

49 % performn a final tuning after each sub—division (see Newman 2006).

50 bp.modules.DoKernighanLinTunning = false; %Very slow, so we turn off.
```

We need to calculate the modularity explicitly by calling:

```
53 bp.modules.Detect();
```

If we are interested only in node module indexes we can use bp.modules.row_modules and bp.modules.col_modules. However for modularity values we need to call bp.modules.Qb or bp.modules.Qr as follows:

```
60 fprintf('The modularity value Qb is %f\n', bp.modules.Qb);
61 fprintf('The fraction inside modules Qr is %f\n', bp.modules.Qr);
```

```
The modularity value Qb is 0.770942
The fraction inside modules Qr is 0.917417
```

Because AdaptiveBrim is not deterministic you may get a different result. In order to improve the result you may increase the parameter Options.TRIALS_MODULARITY, which specify how many random restarts will perform the algorithm.

The value $0 \le Q_b \le 1$ is calculated using the standard bipartite modularity function (introduced by Barber) [3] while the value $0 \le Q_r \le 1$ represents the fraction of interactions that fall inside modules.

4.2.5 Calculating Nestedness

Two algorithms exist for calculating nestedness. Contrary to the case of modularity, in this case there is no need to change the algorithm because all the algorithms have an independent property in the Bipartite object. These algorithms are:

- NODF (Nestedness metric based on overlap and decreasing filling). With value in the interval [0,1] [1].
- NTC (Nestedness Temperature Calculator) With value in the interval [0 1] [2].

The first algorithm is runned during the creation of the Bipartite object, but because the NTC algorithm is slow, you need to run the algorithm explicitally:

```
85 bp.ntc.CalculateNestedness();
```

Finally to show the values of the two algorithms you need to call::

```
88 % The same value will be printed all the times
89 fprintf('The nestedness NODF value is %f\n', bp.nodf.nodf);
90 % Because the value depends in the sorting of rows and columns, it may
91 % variate from trial to trial.
92 fprintf('The nestedness NTC value is %f\n', bp.ntc.N);
```

```
The nestedness NODF value is 0.034053
The nestedness NTC value is 0.954287
```

We can print all the structure values by just calling:

```
95 bp.printer.PrintStructureValues();
```

```
Modularity
Used algorithm:
NewmanModularity
N (Number of modules):
Qb (Standard metric):
Qr (Ratio of int/ext inter):
0.8979
Nestedness
NODF metric value:
0.0341
```

4.2.6 Plotting in Matrix Layout

NTC metric value:

You can print the layout of the original, nestedness, and modular sorting. If you matrix is weighted in a categorical way using integers (0,1,2...) you can visualize a different color for each interaction, where 0 is no interaction. For using this functionality you need to assign a color for each interaction and specifically indicate that you want a color for each interaction before calling the plot function:

0.9543

```
figure(1);
104 % Matlab command to change the figure window;
105 set(gcf,'Position',[19 72 932 922]);
106 bp.plotter.font_size = 2.0; %Change the font size of the rows and labels
107 % Use different color for each kind of interaction
108 bp.plotter.use_type_interaction = true; %
```

```
bp.plotter.color_interactions(1,:) = [1 0 0]; %Red color for clear lysis
bp.plotter.color_interactions(2,:) = [0 0 1]; %Blue color for turbid spots
bp.plotter.back_color = 'white';
% After changing all the format we finally can call the plotting function.
bp.plotter.PlotMatrix();
```

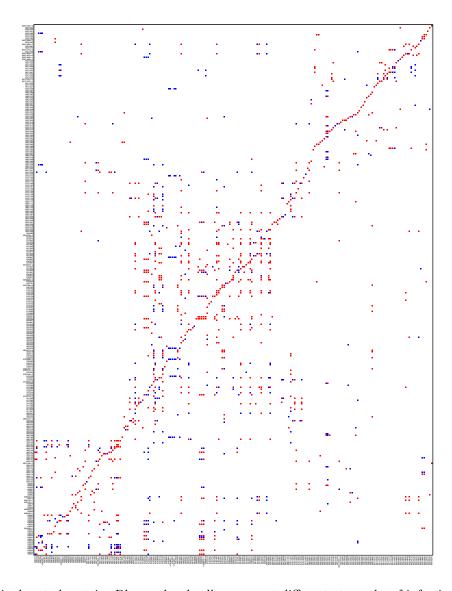


Figure 3: Original sorted matrix. Blue and red cells represent different strengths of infection between virus and bacteria. Rows and columns represent bacteria and phages, respectively.

For plotting the nestedness matrix you may decide to use or not an isocline. The nestedness pattern is just the matrix sorted in decreasing degree for row and column nodes.

```
figure(2);
figure
```

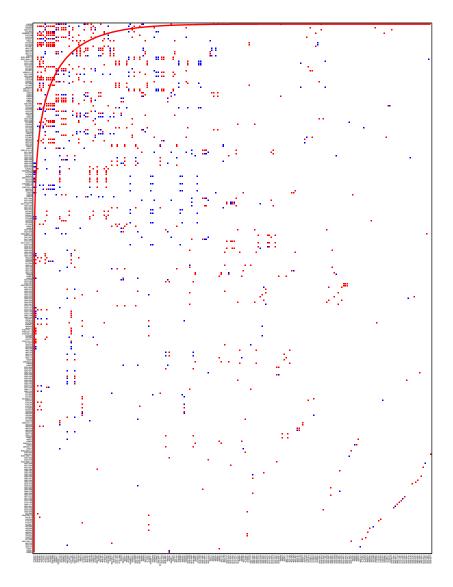


Figure 4: Nested sorted matrix. Blue and red cells represent different strengths of infection between virus and bacteria. In a perfectly nested pattern of the same fill than the current matrix, all the interaction cells will lay above the isocline (red line).

For plotting the modularity sort, the plot function will calculate the modularity (call bp.modules.Detect()) if you have not previously called it.

```
figure(3);
figure
```

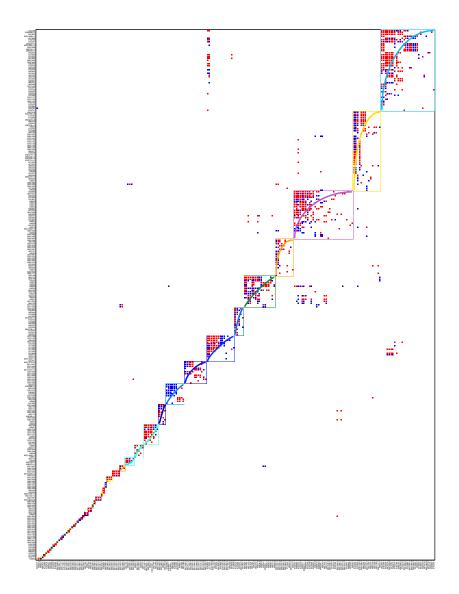


Figure 5: Modular sorted matrix. Blue and red cells represent different strengths of infection between virus and bacteria. Each block represent a different module.

4.2.7 Plotting in graph layout

Plotting in graph layout use the same three functions than matrix layout. You just need to replace the part Matrix in the function name by Graph. For example, for plotting the graph layout of modularity we will need to type:

```
figure (4);
140 % Matlab command to change the figure window;
141 set(gcf,'Position',[19+600 72 932 922]);
142 bp.plotter.PlotModularGraph();
```

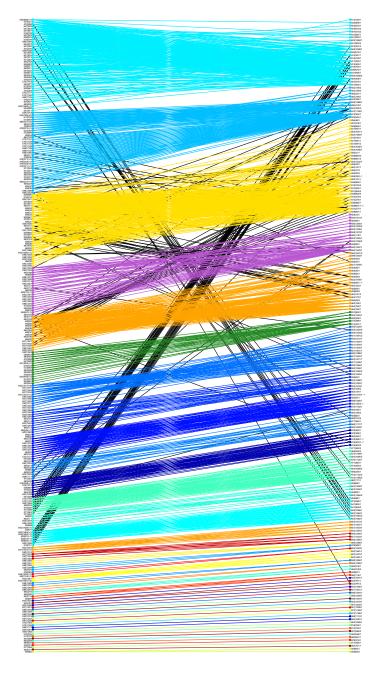


Figure 6: Modular graph layout. Nodes and interactions are colored according to the module they belong to. Black color is used for interaction across modules. Left and right side nodes represent bacteria and phages, respectively.

4.2.8 Statistical analysis in the entire network

We can perform an statistical analysis in the entire network for nestedness and modularity. In order to make an statistical analysis of the structure values we need to decide how many replicates we will need and what null model is more convenient for what we need. File NullModels.m contain all the available null models, while file StatisticalTest.m contains all the functions required for performing this analysis. The current

null models are:

- NullModels.EQUIPROBABLE: A random matrix in which all the interactions are uniformelly permuted. Another common name for this matrix is Bernoulli Matrix.
- NullModels.AVERAGE: A random matrix in which each element has an interaction with probability that depends on the sum of both the row and column to which the cell belongs to.
- NullModels.AVERAGE_ROWS: A random matrix in which each element has an interaction with probability that depends on the sum of the row to which the cell belongs to.
- NullModels.AVERAGE_COLS: A random matrix in which each element has an interaction with probability that depends on the sum of the column to which the cell belongs to.

To perform the statistical analysis of all the structure values we can just type bp.statistics.DoCompleteAnalysis(), which will perform an analysis using the default number of random matrices (Options.REPLICATES) and the default null model (Options.DEFAULT_NULL_MODEL). However, here we will chose directly those parameters:

```
171 % Do an analysis of modularity, nodf, and ntc values using 100 random
172 % matrices and the EQUIPROBABLE (Bernoulli) null model.
173 bp.statistics.DoCompleteAnalysis(100, @NullModels.EQUIPROBABLE);
```

```
Creating 100 null random matrices...

Performing NODF statistical analysis...

Performing Modularity statistical analysis...

Performing NTC statistical analysis...
```

The last function call printed information about the current status of the simulation. For printing the results we need to call:

```
pp.printer.PrintStructureStatistics(); %Print the statistical values
```

Modularity

Null model: NullModels.EQUIPROBABLE
Replicates: 100
NODF value: 0.0341
z-score: 18.4316
percent: 100.0000
NTC Nestedness

 Null model:
 NullModels.EQUIPROBABLE

 Replicates:
 100

 NTC value:
 0.9539

 z-score:
 15.9336

 percent:
 100.0000

The printed information is as follows:

- 1. Null Model: The null model used to created the random matrices.
- 2. Replicates: The number of random matrices that were created for performing the statistical analysis.
- 3. Value: The value in the tested real matrix (Qb, NTC, NODF).
- 4. z-score: The z-score of the real matrix using the values of the random matrices.
- 5. percent: The percent of random matrices than have an smaller value than the matrix that is being evaluated.

Additional information that can be acceded via code includes the mean, standard deviation, and t-test results. Be aware that the number of replicates is especially critical parameter for the results of the statistical analysis. To chose this number consider the size and fill of the matrix. As a rule of thumb, 100 works fine as quick analysis, and 10,000 for a more accurate result (up to a matrix size of 300 by 300).

4.2.9 Statistical Analysis of the internal modules

In addition to be able to perform structure analysis in the entire network, we may be able (depending in the size and module configuration of the tested matrix) to perform a structural analysis in the internal modules. We will show next (i) how to do an analysis of modularity and nestedness in the internal modules and (ii) how to test for a possible correlation between node labeling and module configuration. All the functions for performing this kind of analysis is encoded in file InternalStatistics.m. For calculating the statistical structure of the internal modules we just need to call:

```
% 100 random matrices using the EQUIPROBABLE null model.
204 bp.internal_statistics.TestInternalModules(100,@NullModels.EQUIPROBABLE);
```

The last function call will print information about what is the current matrix (module) that is being evaluated. Like this, the user knows at every moment the current status of the analysis:

```
Testing Matrix: 1 . . . Testing Matrix: 2 . . . . Testing Matrix: 3 . . . . Testing Matrix: 4 . . . . Testing Matrix: 5 . . . . Testing Matrix: 6 . . . . Testing Matrix: 7 . . . . . . and so on . . .
```

Finally, to print the results we just need to call:

```
pp.printer.PrintStructureStatisticsOfModules(); % Print the results
```

```
2,0.42241,0.33384,
                         7.6765,
                                         100,0.29082,
                                                         0.2388,
                                                                        3.5568,
                                                                                          100,0.77327,
                                                                                                        0.51871,
                                                                                                                       5.7448,
3,0.31017,0.29024
                         1.9282
                                          97,0.40425,
                                                         0.25719,
                                                                       13.2868
                                                                                          100,0.76455,
                                                                                                        0.46967
                                                                                                                        8.866,
 1.0.39294.0.39632
                                                          . 23815
                                                                        4.6112
                                                                                          100.0.75558.
5, 0.274,0.22035,
6,0.37778,0.30769,
                         2.0488
                                          97,0.38468,
                                                                       0.39141,
                                                                                           63,0.73545,
                                                                                                                       1.4505
  .0.20003.0.24335
                         -3.8697
                                           0. 0.6963
                                                         0.42384
                                                                        7.9444
                                                                                          100.0.87392
                                                                                                           47566
                                                                                                                       6.7485
                                                                        -1.6833,
-1.9416,
                                                                                            2,0.56773
    .47396,0.31748
                           6554
                                                          0.3541
                         2.9189,
10,0.31293, 0.2756
                                          91,0.36885
                                                          .43368
                                                                       -0.85541,
                                                                                           20,0.65236
                                                                                                           66395
                                                                                                                      -0.14178,
11,0.16272,0.18304
                                           6.0.76951
                                                         0.57219
                                                                        3.5834
                                                                                          100.0.86983
                                                                                                           61222
13,0.18343,0.19118,
                                                                        1.4436,
                                                                                                                      0.92978,
     0.32, 0.2488
                        0.67173
                                          32,0.66667
                                                           0.46
                                                                        1.1346
                                                                                           71,0.87346,
                                                                                                        0.87029
                                                                                                                      0.028445,
                            NaN,
                                                                            NaN
                                                                                                                          NaN,
                            NaN
                                                                            NaN
                                                                                                                          NaN.
                                                                                                                          NaN,
                                                                                                                           NaN.
                                                                                            0,0.99999, 0.99999,
```

```
24, 0, 0, NaM, 0, 0, NaM, 0, 0, 0, NaN, 0,0.99999, 0.99999, -0.99499, 0 0, 0, 0, NaN, 0, 0, 0, 0, NaN, 0, 0, 0, 0, NaN, 0, 0, 0, NaN, 0, 0, 0, 0, 0, NaN, 0, 0, 0, 0, NaN, 0, 0, 0, 0, NaN, 0, 0,
```

The module indexing is in the same order that the plotted modularity matrix, in which Network 1 corresponds to the one located at the top right of Figure 5. This last created table shows the same values previously described. However it is specially usefull for describing some of the possible results that the user may get at some point. What follows summarize some of the important points:

- NaN values appear in many of the z-scores. The reason of those values is because they are fully connected and mostly composed of only one node of each type (a matrix of size 1×1 . Therefore only one permutation of the matrix exist and by consequence all the random matrices are of the same size than the one being analyzed. This makes the standard deviation to be 0, and therefore the z-score to be 0/0 = NaN.
- Some of the matrices that have NaN values dot not have on the NTC values. However, they are also fully connected. The reason for which the z-score is not NaN (as theoretically predicted) is a computational error. In other words, for the z-score we are dividing a very small number over another very small number.
- The last module (49) has NaN values not only in the z-score values, but also in the values of modularity. The reason of these results, is because this module is compose of only one column (phage) node and the matrix is of size 0 × 1.

We can also study if a correlation exists between the row labeling and the module configuration. For performing this analysis we always will need a classification for rows and/or columns that group them in different sets. In this case we have as labeling the station number from which the bacteria and phages were extracted. Therefore what we will study is if there exist a correlation between the station location (geography) and the module configuration. We will use the same method that was used in Flores et al 2012 [14]. Given the labeling this method calculates the diversity index of the labeling inside each module and compare it with random permutations of the labeling across the matrix.

```
%Using the labeling of bp and 1000 random permutations
bp.internal_statistics.TestDiversityRows(1000);
% Using specific labeling and Shannon index
bp.internal_statistics.TestDiversityColumns( ...
1000,moebus.phage_stations,@Diversity.SHANNON_INDEX);
%Print the information of column diversity
bp.printer.PrintColumnModuleDiversity();
```

Diversity index: Diversity.SHANNON_INDEX
Random permutations: 1000

Module, index value, zscore, percent

```
1,
          2.662, -1.7609,
                                5.7
 2,
         2.3959,-0.2332,
                               31.1
         2.3421,-5.1204,
 3,
                                  0
         1.6434,-3.3856,
                                0.4
 4,
 5,
         1.2622, -9.1915,
                                   0
         1.6094,0.54671,
 6,
                               25.6
                                0.6
         2.0262,-2.8726,
 7,
         1.0751,-8.3588,
                                   0
 9,
         1.4979, -4.2423,
                                0.3
                                0.8
10,
         1.0397,
11,
         1.4942,-2.9839,
                                   1
         1.3322,-1.2265,
                                2.6
12,
        0.95027,-3.7459,
13,
                                0.1
         1.0986, 0.32725,
14,
                               10.3
15,
         1.0397,-1.9639,
                                0.5
16,
         1.0986,0.31089,
                                   9
               0,
                                   0
17,
                      NaN,
18,
               0,
                       NaN,
                                   0
                                   0
19,
               0,
                      NaN,
20,
        0.63651, -2.982,
                                   0
21,
               0,
                      NaN,
                                   0
22,
               0,
                      NaN,
                                   0
23,
               0,-4.7735,
                                   0
24,
        0.69315, 0.1633,
                                2.6
25,
               0,
                       NaN,
                                   0
26,
               0,
                       NaN,
                                   0
27,
               0,
                       NaN,
                                   0
28,
               0,
                                   0
                      NaN,
29,
               0,
                                   0
                       NaN,
30,
               0,
                       NaN,
                                   0
31,
               0,
                       NaN,
                                   0
32,
               0,
                      NaN,
                                   0
33,
               0,
                       NaN,
                                   0
                                   0
34,
               0,
                      NaN,
35,
               0,
                       NaN,
                                   0
36,
                                   0
               0,
                      NaN,
37,
               0,
                       NaN,
                                   0
38,
               0,
                                   0
                      NaN,
               0,
39,
                      NaN,
                                   0
               0,
                                   0
40,
                      NaN,
               0,
                                   0
41,
                      NaN,
42,
               0,
                      NaN,
                                   0
               0,
                                   0
43,
                      NaN,
                                   0
44,
               0,
                       NaN,
               0,
                                   0
45,
                       NaN,
               0,
46,
                       NaN,
                                   0
                                   0
47,
               0,
                      NaN,
               0, -5.0991,
                                   0
48,
49,
               0,
                      NaN,
                                   0
```

Using a two tailed p-value of 0.05 we can see that 3, 4, 5, 7, ... are not as diverse as random labeling and conclude that those modules have phages that were isolated from similar locations. The module indexing is

in the same order that the plotted modularity matrix, in which module 1 corresponds to the one located at the top of the plot. The NaN values happens because such modules have only a single phage and therefore the standard deviation used for calculating the z-score is 0.

4.3 BiMat - Group Statistics Example

This example will introduce the user to the features about how to perform an statistical analysis of a group of bipartite networks (matrices). For doing that we will use the data from Flores et Al 2011. This data consist of 38 bipartite adjacency matrices of different sizes. Each matrix is named according to the first author paper from which it was extracted. We will perform an analysis of modularity and nestedness in the entire set.

This example is located on examples/group_matrices.m and make use of examples/group_testing_data.mat data file.

4.3.1 Contents

- Add the source to the MATLAB® path
- Creating a Group Testing object
- Perform an statististical analysis in the set of matrices
- Using a GroupStatistics object to create your own plots

4.3.2 Add the source to the MATLAB® path

```
11 %Assuming that you run this script from examples directory
12 g = genpath('../'); addpath(g);
13 close all; %close all open figures
```

We need also to load the data from which we will be working on

```
16 load group_testing_data.mat;
```

The loaded data is a set of 38 matrices together with a name that refer to the first author and year from the paper from which the matrix was extracted. These matrices were published by Flores et Al 2011 [13].

4.3.3 Creating a Group Statistics object

If the number of random matrices and the null model are not assigned, 100 and AVERAGE are used as default. Here we will use 100 random matrices with the EQUIPROBABLE null model

```
22 gp = GroupStatistics(grouptesting.matrices); % Create the main object
```

4.3.4 Perform an statistical analysis in the set of matrices

Suppose that we are interested in calculating the modularity and nestedness using the NTC algorithm as Flores et Al 2011 did. In addition, following the approach of Flores et Al 2011 [13], we want to use the equiprobable model as null model in our random networks. The way to perform this analysis is by running the next lines:

```
30 gp.replicates = 100; %How many random networks we want for each matrix 31 gp.null_model = @NullModels.EQUIPROBABLE; %Our Null model
```

```
gp.modul_class = @AdaptiveBrim; %Algorithm for modularity.
gp.do_temp = 1; % Perform NTC analysis (default)
gp.do_modul = 1; % Perform Modularity analysis (default)
gp.do_nest = 1; % Perform Modularity analysis (default)
gp.names = grouptesting.name;
gp.DoGroupTesting(); % Perform the analysis.
```

```
Testing Matrix: 1 . . . Testing Matrix: 2 . . . . Testing Matrix: 3 . . . . Testing Matrix: 4 . . . . Testing Matrix: 5 . . . . Testing Matrix: 6 . . . . Testing Matrix: 7 . . . . . . and so on . . .
```

Notice that DoGroupTesting method prints information about the current networks that is being analyzed, such that the user will know at every moment the current status of the analysis. After the analysis is finished a simple statistical measure to say that a matrix is nested and/or modular is to chose a two tail p-value = 0.05 as Flores et al 2011 did. Therefore, the next lines of code will show how many matrices are found nested and/or modular

```
46 fprintf('Number of NTC nested matrices: %i\n',sum(gp.tempvals.percent ≥ 97.5));
47 fprintf('Number of NODF nested matrices: %i\n',sum(gp.nestvals.percent ≥ 97.5));
48 fprintf('Number of modular matrices: %i\n',sum(gp.qb_vals.percent ≥ 97.5));
```

```
Number of nested matrices: 29
Number of modular matrices: 6
```

Because we only did 100 random matrices you may get different results. For a more accurate result you may try 1.000 or even 10,000. Finally we can show detailed results for the entire set of matrices:

```
53 gp.PrintResults();
```

```
NODF, NODF mean, NODF z-score, NODF percent,
                                             93, 0.28197,
                                                                                                   2,0.65025, 0.66945,
 1, 0.30992, 0.27149,
                            1.2134,
                                                             0.45503,
                                                                             -2.2113,
                                                                                                                             -0.21996,
                                               0, 0.19418,
     0.2144, 0.40165,
                           -5.1897.
                                                              0.27106.
                                                                             -1.4879
                                                                                                   6.0.80601, 0.68326
                                                                                                                                1.451
                                                                             0.62112
                                                                                                                0 64117
 5, 0.25652, 0.30552
                            -2.7321.
                                                   0.6114.
                                                              0.37957
                                                                               5.288.
                                                                                                 100.0.94524.
                                                                                                                 0.5527
                                                                                                                               5.2917
     0 2699
              0 50685
                            -3 6399
                                                                              -2 9556
                                                                                                   0.0.82486. 0.78899
                                                                                                                              0 41664
                                                                                                 100,0.79783, 0.43154
                                                              0.40415,
 9, 0,21395,
               0.34563.
                           -10.4795.
                                               0. 0.28559.
                                                              0.24741
                                                                               2.4581
                                                                                                  99.0.85564.
                                                                                                                0.54169
                                                                                                                                7.8469
                                               4, 0.5081, 0, 0.17111,
                                                                              1.0616,
-3.0679,
    0.29191
                27974
                            0 57535
                                                                                                  85 0 64376
                                                                                                                                0 6057
     0.4821, 0.40186,
                                             100, 0.2197,
1, 0.19444,
                                                                              -1.1726,
                                                                                                   9,0.86948,
                                                                                                                                2.4762
                0.4842,
                            -2.3729,
                                                              0.15741
                                                                               0.7342,
                                                                                                  75,0.92493, 0.88036
    0.31667.
               0.34872
                                                                             0.80359
                                                                                                  85,0.77249,
                                                                                                                                 4.645
15, 0.20023,
16, 0.18696,
               0.19292,
                                             10, 0.61153,
                                                                .40412,
                                                                             17.0309,
                                                                                                 100,0.83459,
                                                                                                                              17.5724
                            -1.7491
                                                              0.70271
                                                                              -3.2974
                                                                                                   2.0.99875.
                                                                                                                0.84967
                                                                                                                               3.6965
                                                                                                                                3.7284
069048
                                                                                                  88.0.90103
19, 0,
20, 0.31027,
                                               0, 0.11655,
               0.47264,
                            -7.6013,
                                                               0.1328,
                                                                              -1.7354,
                                                                                                    7,0.85639,
                                                                                                                0.64653
                                                                                                                               4.9957
                            -4.1428,
7.0829,
-5.1781,
21. 0.19136.
                                               0. 0.34694.
                                                                             0.45722
                                                                                                  64,0.97793,
                                                                                                                0.73159
                                                                                                                               2.5262
                                                                             25.9477,
-4.4116,
                                                                                                                               20.3579,
6.0492,
     0.4102,
               0.34576,
                            3.7243,
                                             100, 0.19474,
                                                               0.2912
                                                                                                   0,0.70409,
                                                                                                                0.54842
                                                                                                                               2.4715
                                               1, 0.58971.
25, 0.14966
               0.19488
                            -2.3858
                                                                           0.0099181
                                                                                                  49.0.85489
                                                                                                                                 85917
                                                                                                  0,0.87649,
75,0.83643,
27, 0.20209,
                                             14, 0.51298,
              0.21213,
                                                                             0.62949,
                                                                                                                0.43449
                                                                                                                              14.4161
28. 0.35584.
                            -1.6314
                                               4, 0.30619,
                                                              0.20129
                                                                               6.7904.
                                                                                                 100.0.80589.
                                                                                                                0.59973
                                                                                                                                3.7898
                                                                                                 58,0.64478, 0.
100,0.93098, 0.
87,0.97517, 0
    0.37622,
                                            100, 0.41751,
46, 0.53377,
                                                                                                                                5.6463,
3.4536,
31, 0.22893, 0.47055,
                            -6.6212,
                                               0, 0.18127,
                                                               0.1555,
                                                                                                                0.7038
32. 0.18341.
                                            100, 0.71354.
                                                                                                 100,0.78597, 0.61381
```

```
34, 0.4876, 0.51298, -0.34778, 35, 0.18421, 0.13342, 1.1952, 89,0.86017, 0.89187, -0.53745, 22
35,0.084203,0.084955, -0.29141, 42, 0.7063, 0.71613, -0.29018, 36,0.94762, 0.6857, 4.2764, 100
36, 0.61983, 0.63331, -0.15015, 40, 0.04023, 0.029799, 0.73559, 79,0.98846, 0.94324, 1.3188, 100
37, 0.21079, 0.27282, -6.8961, 0, 0.70141, 0.26149, 49.6956, 100,0.88768, 0.82511, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 2021, 20
```

4.3.5 Plotting results

The user can visualize the results of the last output in a graphical way. For example for visualizing the results of modularity and NTC nestedness value, the user can type:

```
gp.plotter.p_value = 0.05; %p-value for error bars
gp.plotter.font_size = 10; %Size for x-labels.
gp.plotter.PlotModularValues();
gp.plotter.PlotNTCValues();
```

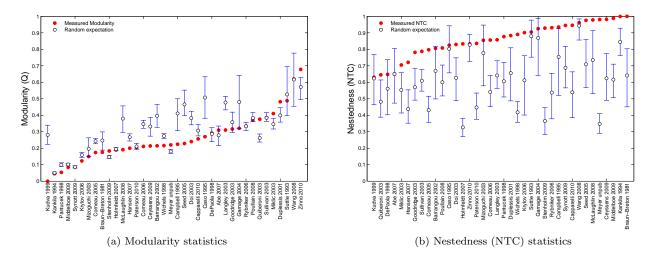


Figure 7: Visual representation of the statistical tests in the set of matrices. Red circles represent the value of the analyzed networks. White circles represent the mean of the null model, while the error bars represent the networks that falls inside a two-tailed version of the random null model values. The margin of the error bars are (p-value,1-p-value), where p-value can is an optional argument of the plot functions.

```
gp.plotter.p_value = 0.05; %p—value for color labeling
for % Plot of nested graphs
gp.plotter.bead_color_rows = 'blue'; %Color of row nodes
gp.plotter.bead_color_columns = 'red'; %Color of column nodes
gp.plotter.link_width = 0.5; %Edge width
gp.plotter.nest_test = 2; %Use NODF for color labels (1 is for NTC).
gp.plotter.PlotNestedGraphs(5,8);
gp.plotter.PlotNestedGraphs(5,8);
gp.plotter.PlotModularMatrices
```

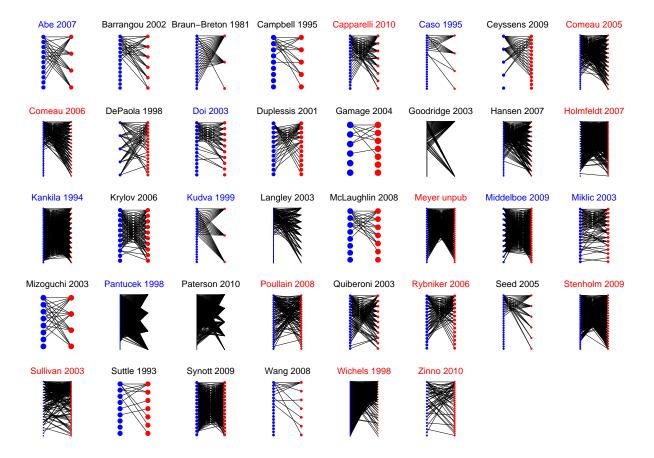


Figure 8: NODF nestedness values of a set of 38 matrices of phage-bacteria networks. A two-tail p-value of 0.05 was used for labeling the names. Blue and and red lebels represent anti and statistical significance, respectively. Notice that this Figure shows an smaller number of nested matrices than the NTC plot of the previous figure.

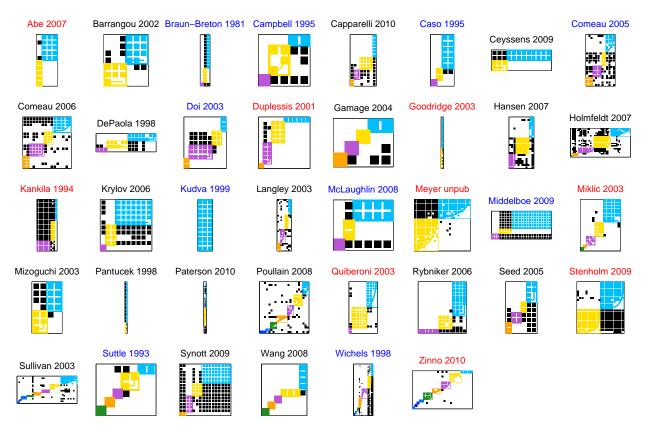


Figure 9: The meta-set collected on Flores et al [13] plotted using the modularity algorithm of the BiMat library. Red and blue labels represent significant modularity ($p \ge 0.975$) and anti-modularity ($p \le 0.275$), respectively. For bibliographic information about these matrices see [13].

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