# ChromHMM Version 1.04 User Manual

# Jason Ernst and Manolis Kellis

# Overview

ChromHMM is a Java program for the learning and analysis chromatin states using a multivariate Hidden Markov Model that explicitly models the observed combination of marks.

ChromHMM can be run on any computer supporting Java 1.5 or later. ChromHMM is executed from the command line with a command such as:

java -mx4000M -jar ChromHMM.jar Command [commandoptions] commandparameters

Where the 4000 specifies the amount of memory given to Java and could be adjusted based on the size of the data, the model size, and the *Command* being executed. In some cases the memory flag could be omitted.

ChromHMM has ten top level commands which then determine the required and optional set of parameters. The top level commands are briefly described here and a detailed description of each command, the required and optional parameters can be found in the remaining sections.

LearnModel – takes a set of binarized data files, learns chromatin state models, and by default produces a segmentation, generates browser output with default settings, and calls OverlapEnrichment and NeighborhoodEnrichments with default settings for the specified genome assembly. A webpage is a created with links to all the files and images created.

BinarizeBed – converts a set of bed files of aligned reads into binarized data files for model learning and optionally prints the intermediate signal files

BinarizeSignal – converts a set of signal files into binarized files.

MakeSegmentation – takes a learned model and binarized data and outputs a segmentation.

MakeBrowserFiles - can convert segmentation files into a browser viewable format.

OverlapEnrichment – shows the enrichment of each state of a segmentation for a set of external data

NeighborhoodEnrichment – shows the enrichment of each state relative to a set of anchor positions

Compare Models - can compare models with different numbers of states in terms of correlation in emission parameters

Reorder - allows reordering the states of the model, the columns of the emission matrix, or adding state labels

StatePruning – can be used to prune states from a model in order to initialize models when using the non-default two pass approach

The usage for any of these commands can be obtained at the command line by typing at the command line java —jar ChromHMM.jar *Command* 

# LearnModel

### Description

This command takes a directory with a set of binarized data files and learns a chromatin state model. Binarized data files have '\_binary' in the file name. The format for the binarized data files are tuat the first line contains the name of the cell separated by a tab with the name of the chromosome. The second line contains in tab delimited form the name of each mark. The remaining lines correspond to consecutive bins on the chromosome. The remaining lines in tab delimited form corresponding to each mark, with a '1' for a present call or '0' for an absent call and a '2' if the data is considered missing at that interval for the mark. As an example:

```
Cell chr1
Mark1 Mark2 Mark3
0 0 0
0 1
0 1 1
```

The emission parameters in table format are printed to text, png, and svg image files named beginning with 'emissions\_' then having the number of states and optionally an identifier followed by a '.txt', '.png', and '.svg' respectively. Similarly the transition parameters are printed to text and image files prefixed with 'transitions'. All the model parameters in an internal format are printed to a file beginning with the name 'model\_'. After each iteration across the genome these files are updated. At the termination of the learning process, LearnModel effectively calls MakeSegmentation to produce segmentation files, MakeBrowserFiles, OverlapEnrichments, and NeighborhoodEnrichments with all the files corresponding to the specified assembly. Also upon termination webpage is created called webpage\_NUMSTATES.html summarizing all the files and images created, and if a browser can be found, then is automatically opened with this file. The current search progress in terms of estimated log likelihood and change after each iteration is printed to the terminal. If one terminates the search early, a segmentation can still be produced by applying MakeSegmentation to the current model file.

### **Usage**

```
LearnModel [-b binsize][-color r,g,b][-d convergedelta]
[-e loadsmoothemission][-f inputfilelist][-h informationsmooth]
[-holdcolumnorder][-i outfileID][-init information|random|load]
[-l chromosomelengthfile][-m modelinitialfile][-nobed][-nobrowser][-noenrich]
[-printposterior][-printstatebyline][-r maxiterations][-s seed]
[-stateordering emission|transition][-t loadsmoothtransition]
[-x maxseconds][-z zerotransitionpower]
inputdir outputdir numstates assembly

Note items in [] are optional
```

### **Required Parameters**

inputdir – This is the directory containing the binarized input files. Only file names containing '\_binary' are used.

outputdir – This is the directory where the output files are written.

numstates – This parameter specifies the number of states to include in the model.

assembly – specifies the assembly. overlap and neighborhood enrichments will be called with default parameters using this assembly.

- -b binsize The number of base pairs in a bin determining the resolution of the model learning and segmentation. By default this parameter value is set to 200 base pairs.
- -color r,g,b This specifies the color of the heat map. r,g,b are integer values between 0 and 255 separated by commas. By default this parameter value is 0,0,255 corresponding to blue.
- -d convergedelta The threshold on the change on the estimated log likelihood that if it falls below this value, then parameter training will terminate. If this value is less than 0 then it is not used as part of the stopping criteria. The default value for this parameter is 0.001.
- -e loadsmoothemission This parameter is only applicable if the load option is selected for the init parameter. This parameter controls the smoothing away from 0 when loading a model. The emission value used in the model initialization is a weighted average of the value in the file and a uniform probability over the two possible emissions. The value in the file gets weight (1-loadsmoothemission) while uniform gets weight loadsmoothemission. The default value of this parameter is 0.02.
- -f inputfilelist A list of files to include in the segmentation. If this option is not provided then by default all files in inputdir with '\_binary' in the name will be included in the model learning.
- -h informationsmooth A smoothing constant away from 0 for all parameters in the information based initialization. This option is ignored if random or load are selected for the initialization method. The default value of this parameter is 0.02.
- -holdcolumnorder -Including this flag suppresses the reordering of the mark columns in the emission parameter table display.
- -i outfileID If this option is included the string outfileID is included in the file names of all the output files.
- -init information|random|load This specifies the method for parameter initialization method. information is the default method described in (Ernst and Kellis, Nature Methods 2012). Random-randomly initializes the parameters from a uniform distribution. load loads the parameters specified in modelinitialfile and smooths them based on the value of the loadsmoothemission and loadsmoothtransition parameters. The default is information.
- -1 chromosomelengthfile This file specifies the length of the chromosomes. It is a two column tab delimited file with the first column specifying the chromosome name and the second column the length. If this file is provided then no end coordinate will exceed what is specified in this file. By default BinarizeBed excludes the last partial bin along the chromosome, but if that is included in the binarized data input files then this file should be included to give a valid end coordinate for the last interval. Also by default if a file exists in CHROMSIZES for the provided assembly that will be used.

- -m modelinitialfile This specifies the model file containing the initial parameters which can then be used with the load option
- -nobed If this flag is present, then this suppresses the printing of segmentation information in the four column format. The default is to generate a four column segmentation file
- -nobrowser If this flag is present, then browser files are not printed. If -nobed is requested then browserfile writing is also suppressed.
- -noenrich If this flag is present, then enrichment files are not printed. If -nobed is requested then enrichment file writing is also suppressed.
- -printposterior If this flag is present the posterior probabilities over state assignments are also printed in a file. These files end with '\_posterior.txt'. One file is generated per cell type and chromosome. The first line of these files specify the chromosome and cell type, followed by a header line for each column, and then the posterior probabilities one per line. By default these files are not printed.
- -printstatebyline If this flag is present the state assignment are printed to a file one per line. These files end with '\_maxstate.txt'. One file is generated per cell type and chromosome. The first line specifies the cell type and chromosome and the second line says MaxState and the state ordering methods. The remaining lines have the state assignments. By default these files are not printed.
- -r maxiterations This option specifies the maximum number of iterations over all the input data in the training. By default this is set to 200.
- -s seed This allows the specification of the random seed. Randomization is used to determine the visit order of chromosomes in the incremental expectation-maximization algorithm used to train the parameters and also used to generate the initial values of the parameters if random is specified for the init method.
- -stateordering emission | transition This determines whether the states are ordered based on the emission or transition parameters. See (Ernst and Kellis, Nature Methods) for details. Default is emission.
- -t loadsmoothtransition This parameter is only applicable if the load option is selected for the init parameter. This parameter controls the smoothing away from 0 when loading a model. The transition value used in the model initialization is a weighted average of the value in the file and a uniform probability over the transitions. The value in the file gets weight (1-loadsmoothtransition) while uniform gets weight loadsmoothtransition. The default value is 0.5.
- -x maxseconds This parameter specifies the maximum number of seconds that can be spent optimizing the model parameters. If it is less than 0, then there is no limit and termination is based on maximum number of iterations or a log likelihood change criteria. The default value of this parameter is -1.
- -z zerotransitionpower This parameter determines the threshold at which to set extremely low transition probabilities to 0 durining training. Setting extremely low transition probabilities makes model learning more efficient with essentially no impact on the final results. If a transition probability falls below 10^-zerotransitionpower during training it is set to 0. Making this parameter to low and thus the cutoff too high can potentially cause some numerical instability. By default this parameter is set to 8.

# BinarizeBed

# Description

This command converts coordinates of aligned reads into binarized data form from which a chromatin state model can be learned. The binarization is based on a poisson background model. If no control data is specified the parameter to the poisson distribution is the global average number of reads per bin. If control data is specified the global average number of reads is multiplied by the local enrichment for control reads as determined by the specified parameters. Optionally intermediate signal files can also be outputted and these signal files can later be directly converted into binary form using the BinarizeSignal command.

### Usage

```
BinarizeBed [-b binsize][-c controldir][-center]
[-colfields chromosome,start,end[,strand]][-e offsetend][-f foldthresh]
[-n shift][-o outputcontroldir][-p poissonthresh][-peaks]
[-s offsetstart][-strictthresh][-t outputsignaldir][-u pseudocountcontrol]
[-w flankwidthcontrol] chromosomelengthfile inputbeddir cellmarkfiletable outputbinarydir
```

Note items in [] are optional

## **Required Parameters**

chromosomelengthfile —A two column tab delimited file with the first column being the chromosome and the second being the chromosome length. For genome assemblies hg18, hg19, mm9, dm3, and ce6 these can be found in the directory CHROMSIZES. For other assemblies these can be obtained with the fetchChromSizes script available from the UCSC browser <a href="http://hgdownload.cse.ucsc.edu/admin/exe/">http://hgdownload.cse.ucsc.edu/admin/exe/</a> specifying the desired assembly and redirecting the output to a text file.

inputbeddir - The directory containing the input bed files.

cellmarkfiletable — A tab delimited file each row contains the cell type or other identifier for a groups of marks, then the associated mark, then the name of a bed file, and optionally a corresponding control bed file

```
cell1 mark1 cell1_mark1.bed cell1_control.bed cell1 mark2 cell1_mark2.bed cell1_control.bed cell2 mark1 cell2_mark1.bed cell2_control.bed cell2_mark2.bed cell2_control.bed
```

If a mark is missing in one cell type, but not others it will receive a 2 for all entries in the binarization file and -1 in the signal file

outputbinarydir - The output directory to which the binarized data files should be written. These files will be named CELL\_CHROM\_binary.txt

- -b binsize The number of base pairs in a bin determining the resolution of the model learning and segmentation. By default this parameter value is set to 200 base pairs.
- -c controldir A directory containing the control input files. If this is not specified then by default a uniform background will be used in determining the binarization thresholds.
- -center If this flag is present then the center of the interval is used to determine the bin to assign a read. This can make sense to use if the coordinates are based on already extended reads. If this option is selected, then the strand information of a read and the shift parameter are ignored. By default reads are assigned to a bin based on the position of its 5' end as determined from the strand of the read after shifting an amount determined by the –n shift option.
- -colfields chromosome, start, end[, strand] This is a comma delimited list of integers specifying the 0-based index of the columns of the chromosome, the start position, the end position, and the strand ('+' or '-') of the reads. If the option -center is present, then the strand can be omitted. By default the chromosome is found in the first column, the start in the second, and the end position in the third, and the strand in the sixth column unless there are fewer columns in which case the last column of the file is assumed to contain the strand.
- -e offsetend Specifies the amount that should be subtracted from the end coordinate of a read so that both coordinates are inclusive and 0 based. The default value is 1 corresponding to standard bed convention of the end interval being 0-based but not inclusive.
- -f foldthresh This indicates a threshold for the fold enrichment over expected that must be met or exceeded by the observed count in a bin for a present call. The expectation is determined in the same way as the mean parameter for the poission distribution in terms of being based on a uniform background unless control data is specified. This parameter can be useful when dealing with very deeply and/or unevenly sequenced data. By default this parameter value is 0 meaning effectively it is not used.
- -n shift The number of bases a read should be shifted to determine a bin assignment. Bin assignment is based on the 5' end of a read shifted this amount with respect to the strand orientation. By default this value is 100.
- -o outputcontroldir This specifies the directory to which control data should be printed. The files will be named CELL\_CHROM\_controlsignal.txt. Control data will only be outputted if there are control bed files present and an output control directory is specified.
- -p poissonthreshold This option specifies the tail probability of the poisson distribution that the binarization threshold should correspond to. The default value of this parameter is 0.0001.
- -peaks This option specifies to treat the bed files as peak calls directly and give a '1' call to any bin overlapping a peak call.
- -s offsetstart The amount that should be subtracted from the interval start coordinate so the interval is inclusive and 0 based. Default is 0 corresponding to the standard bed convention.
- -strictthresh If this flag is present then the poisson threshold must be strictly greater than the tail probability, otherwise by default the largest integer count for which the tail includes the poisson threshold probability is used.

- -t outputsignaldir If specified signal files will be generated and outputted to the given directory. The files will be named CELL\_CHROM\_signal.txt. These files could later be binarized directly at different thresholds with the BinarizeSignal command. By default no output signal is written.
- -u pseudocountcontrol An integer pseudocount that is uniformly added to every bin in the control data in order to smooth the control data from 0. The default value is 1.
- -w flankwidthcontrol This determines the extent of the spatial smoothing in computing the local enrichment for control reads. The local enrichment for control signal in the  $x^{th}$  bin on the chromosome after adding pseudocountcontrol is computed based on the average control counts for all bins within x-w and x+w. If no controldir is specified, then this option is ignored. The default value is 5.

# BinarizeSignal

# Description

This command converts data already processed into signal files into binarized data files. Signal data files have '\_signal' in the file name. The format for the signal file is the first line contains the name of the cell separated by a tab with the name of the chromosome. The second line contains in tab delimited form the name of each mark. The remaining lines correspond to consecutive bins on the chromosome and contain the integer signal value for each mark. As an example:

Cell	chr1	
Mark1	Mark2	Mark3
0	4	0
1	3	0
2	1	9

Control signal data are in files with '\_controlsignal' in the file name and have either one column which is used for all the marks or control data specified for every mark matched to a '\_signal' data file. Note the binarization from signal is designed only for signal data which represent counts of reads assigned to bins such as the optional output from the BinarizeBed command. If the signal was computed in other ways, then the binarization based on the poisson distribution may not give meaningful results. '-1' entries in a signal files are considered missing.

# **Usage**

```
BinarizeSignal [-c controldir][-f foldthresh][-p poissonthresh]
[-strictthresh][-u pseudocountcontrol][-w flankdwidth] signaldir outputdir

Note items in [] are optional
```

### **Required Parameters**

signaldir - Directory containing the signal input files. Only files with '\_signal' in the name will be read
outputdir - Directory to which the binarized data files should be written

- -c controldir If a controldir is specified then this directory has control signal files that will be used. Files containing '\_controlsignal' will be considered control files and there must be matched control signal file for each signal file.
- -f foldthresh, -p poissonthreshold, -strictthresh, -u pseudocountcontrol, -w flankwidthcontrol are same as described above for BinarizeBed

# MakeSegmentation

# Description

This command takes a saved model file and binarized data and outputs files with the segmentation state assignments and/or the posterior state probabilities. These files can also be outputted by LearnModel, but this enables producing the segmentations without the need to relearn the model

## **Usage**

```
MakeSegmentation [-b binsize][-f inputfilelist][-i outfileID]
[-l chromosomelengthfile][-nobed][-printposterior][-printstatesbyline]
modelfile inputdir outputdir
```

Note items in [] are optional

# **Required Parameters**

modelfile - the file with the model parameters to produce a segmentation

inputdir - the directory with the binarized data for which a segmentation should be produced

outputdir – the output directory for the segmentation results

```
-b binsize, -f inputfilelist , -i outfileID , -l chromosomelengthfile, -nobed, -printposterior, -printstatesbyline are the same as described above in LearnModel
```

# MakeBrowserFiles

# Description

This command converts segmentation coordinates in a .bed file into two files that can be viewed as custom tracks in the UCSC genome browser (http://genome.ucsc.edu). One of the files can give a dense view of the segmentation in a single track with states being differentiated solely by color. The other file gives an expanded view showing one state per track.

### Usage

MakeBrowserFiles [-c colormappingfile][-m labelmappingfile][-n numstates] segmentfile segmentationname outputfileprefix

Note items in [] are optional

### **Required Parameters**

segmentfile - The name of a four column tab delimited segmentation file. The four columns are chromosome, start, end, and label

segmentationname — A name for the segmentation that will be given to the segmentation and displayed in the browser

outputfileprefix – The output file prefix including possibly the directory name. Appended to this prefix is the suffix '\_browserexpanded.bed' and '\_browserdense.bed' for the expanded and dense browser files.

- -c colormappingfile The specified file under this option gives a mapping from state ID to desired color. The file is a two column tab delimited file. The first column contains the state number without the state ordering prefix. The second column contains r,g,b integer values between 0 and 255 delimited by commas specifying the colors for each state. If this file is not specified different colors are automatically generated for each state such that states closer in the order have more similar colors.
- -m labelmappingfile This option can specify a file which maps state IDs to descriptive names. The descriptive names can be appended to the state IDs when viewing the states in the browser. The format of this file is a two column tab delimited file. The first column contains each state ID without the state ordering letter prefix. The second column contains a descriptive name or mnemonic. If this file is not specified, then just the state IDs are displayed in the browser.
- -n numstates This option specifies the number of states on which the segmentation is based. By default the maximum state number is used to indicate how many states there are. This parameter can potentially influence the default state coloring if there locations in the genome which were not assigned to any state.

# OverlapEnrichment

# Description

This command can be used to compute the enrichment of each state of the segmentation for a set of external coordinates by default in bed format. Signal values can optionally be associated with each coordinate to weight the enrichments. The enrichment is outputted as a table in both text and image format. Included in the download are the directories COORDS/hg18, COORDS/hg19, COORDS/mm9, COORDS/ce6, COORDS/dm3. These directories include bed files for the RefSeq transcription start site, transcript end site, gene, exon, and regions within 2kb of the transcription start site obtained from the UCSC genome browser. Also CpG islands are included for hg18, hg19 and mm9, and for hg18 and hg19 a set of coordinates on NuclearLamin domains from (Guelen et, Nature 2008). Coordinates were obtained from the UCSC genome browser.

### Usage

OverlapEnrichment [-a cell][-b binsize][-binres][-color r,g,b][-center]
[-colfields chromosome,start,end[,signal]][-e offsetend][-f coordlistfile][-m
labelmappingfile][-multicount][-posterior][-s offsetstart][-signal][-t
title][-uniformscale] inputsegment inputcoorddir outfileprefix

Note items in [] are optional

### **Required Parameters**

inputsegment — either a segmentation bed file or if the —posterior option is selected the directory containing the posterior files

inputcoorddir – a directory containing the external coordinates for enrichment analysis. If inputcoorddir is a file instead of a directory then enrichments for just the file are computed.

outfileprefix – the prefix of the text and image files to which the fold enrichments should be written. The enrichment text file has a '.txt' extension added and a heat map image has a '.png' and '.svg' extension added.

- -a cell-If the -posterior flag is specified then this option can be used to specify the cell type for which to compute the enrichment. By default the posterior enrichment is computed in aggregate over all cell types.
- -b binsize The number of base pairs in a bin determining the resolution of the model learning and segmentation. By default this parameter value is set to 200 base pairs.
- -binres If this flag is present, then this flag indicates enrichments should be computed at the bin resolution just requiring single base overlap of a coordinate for a bin to fully count the bin opposed to the default of base resolution.
- -center Use the center base for computing enrichments instead of the entire interval

- -colfields chromosome, start, end[, signal] This is a comma delimited list of integers which specify the 0-based index of the column fields contains the chromosome, start, end, and if signal is being used the the signal data. By default the values of these parameters are 0,1,2,3.
- $-color\ r$ , g, b This specifies the color of heat map. r,g,b are integer values between 0 and 255 separated by commas. By default this parameter value is 0,0,255 corresponding to blue.
- -e offsetend Specifies the amount that should be subtracted from the end coordinate so the coordinate is inclusive and 0 based. The default value is 1 corresponding to standard bed convention of the end interval being 0-based and exclusive.
- -f coordlistfile This option specifies a file which lists one per line the coordinate files from the inputcoorddir to compute enrichment for and the order in which they should be displayed in the enrichment table. By default enrichments are computed for all files in inputcoorddir.
- -m labelmappingfile This option can specify a file which maps state IDs to descriptive names. The descriptive names can be appended to the state IDs when viewing the states in the browser. The format of this file is a two column tab delimited file. The first column contains each state ID without the state ordering letter prefix. The second column contains a descriptive name or mnemonic. If this file is not specified, then just the state IDs are displayed.
- -multicount This flag indicates to count overlaps multiple times when the -signal flag is not present. If the -signal flag is present overlaps are always counted multiple times. By default without the -signal overlaps are only counted once. Overlaps are defined to either be at the base resolution or the bin resolution based on the -binres flag. If the input coordinate data for enrichments is known to be non-overlapping then including this flag can speed up the enrichment calculation without effecting the final results.
- -posterior Indicates that the full posterior should be used for the enrichment opposed to the default of the maximum probability state assignments. If this flag is present then inputsegment should be a directory with the posterior files.
- -s offsetstart This parameter specifies the value that should be subtracted from the interval start coordinate so the interval is inclusive and 0 based. Default is 0 corresponding to the standard .bed convention
- -signal If this flag is given, then signal information will be used in the enrichments if available otherwise all overlaps are given equal weight. If signal is used coordinates without signal given will be assumed to have 0 signal in the enrichment calculation
- -t title This option specifies a title for the heat map image. A default title of "Fold Enrichments" is used.
- -uniformscale If this flag is given, then in coloring the heat map all columns will be on the same color scale. The default is to have a column specific color scale which subtracts the minimum value in the column and then divides by the maximum column value.

# NeighborhoodEnrichment

### Description

This command given a set of anchor positions determines the enrichment for each state at fixed positions relative to the anchor positions. Signal values can optionally be associated with each coordinate to weight the enrichments. Strand information can also be optionally used to compute the positional enrichments in a strand aware manner. The enrichments are outputted both as a text file and in image format. The directory ANCHORFILES has subdirectories hg18, hg19, mm9, dm3, and ce6 included containing anchor position files for transcription start sites (TSS) and transcript end sites (TES) for the corresponding assemblies.

### Usage

```
usage NeighborhoodEnrichment [-a cell][-b binsize][-color r,g,b]
[-colfields chromosome,position[,optionalcol1|,optionalcol1,optionalcol2]]
[-l numleftintervals][-m labelmappingfile][-nostrand][-o anchoroffset]
[-posterior][-r numrightintervals][-s spacing][-signal][-t title]
inputsegment anchorpositions outfileprefix
```

Note items in [] are optional

### **Required Parameters**

inputsegment — Either a segmentation bed file or if the —posterior option is selected the directory containing the posterior file

anchorpositions – Specifies a file containing the coordinates of the anchor positions around which state enrichments will be determined. Positions are specified by a chromosome and coordinate.

outfileprefix – The prefix of the text and image files to which the fold enrichments should be written. The enrichment text file has a '.txt' extension added and a heat map images have a '.png' or '.svg' extensions added.

#### **Optional Parameters**

-colfields chromosome, position[,optionalcol1|,optionalcol1,optionalcol2] - This is a comma delimited list of integers which specify the 0-based index of the column fields contains the chromosome, position. If the -nostrand option is not present optionalcol1 is the strand and if -signal is also specified optionalcol2 is the signal. If -nostrand is present and -signal is selected optionalcol1 has the signal column. By default the values of these parameters are 0,1,2,3.

- -l numleftintervals The number of enrichment columns to the left of the anchor position to display. By default this value is 10.
- -m labelmappingfile This option can specify a file which maps state IDs to descriptive names. The descriptive names can be appended to the state IDs when viewing the states in the browser. The format of this file is a two column tab delimited file. The first column contains each state ID without the state ordering letter prefix. The second column contains a descriptive name or mnemonic. If this file is not specified, then just the state IDs are displayed.

- -o anchoroffset The value that should be subtracted from the anchor coordinate so it is 0 based. By the default this value is 0.
- -r numrightintervals The number of enrichment columns to the right of the anchor position to display. By default this value is 10.
- -s spacing The spacing in base pairs at which column enrichments should be displayed. Default is binsize.
- -nostrand If this flag is present then no strand information is used in the enrichment calculations
- -a cell, -b binsize, -color r,g,b, -posterior, -signal, -t title are the same as described for OverlapEnrichment

# CompareModels

### Description

This command allows the comparison of the emission parameters of a selected model to a set of models in terms of correlation. The output is a table in both heat map and text file form showing the maximum correlation of each state in the selected model with its best matching state in each other model.

### **Usage**

```
CompareModels [-color r,g,b] referencemodel comparedir outputprefix

Note items in [] are optional
```

## **Required Parameters**

referencemodel – the file, including the path, to the reference set of emission parameters. This file must start with 'emissions\_' and end with '.txt'

comparedir — the directory containing all the emission parameters to be compared to. Only files that start with 'emissions\_' and end with '.txt' are compared

outputprefix – The prefix for the output files containing parameter correlation files in both .txt, .png, and .svg form. The columns are ordered by the number of states in the models and each row corresponds to a state in the reference model. The values correspond to the best correlation in emission parameters of any state in the comparison model for that state in the reference model.

# **Optional Parameters**

 $-color\ r$ , g, b – This determines specifies the color of heatmap. r,g,b are integer values between 0 and 255 separated by commas. By default this parameter value is 0,0,255 corresponding to blue.

# Reorder

### Description

This command allows reordering the states of a model without relearning the model, and outputs a model file and emission and transition tables with the states reordered. The states can be reordered based on the emission or transition parameters or based on a user provided state ordering. Also the columns of the emission parameters can be ordered based on their original order in the data file, based on the correlation of the columns, or a user specified order. Note that this command does not reorder any existing segmentation or analysis file. In the segmentation files the state number is prefixed by an 'E' if the ordering is based on emissions, 'T' if it was based on the transition parameters, and 'U' if it was user provided. The command can also be used to add descriptive state labels to the parameter heatmaps with the current ordering.

### Usage

```
usage: Reorder [-color r,g,b][-f columnorderingfile][-holdcolumnorder]
[-i outfileID][-m labelmappingfile][-o stateorderingfile][-stateordering
emission|transition] inputmodel outputdir
```

Note items in [] are optional

# **Required Parameters**

inputmodel - The file for the model that should be reordered

outputdir - The directory to which the reordered model, emission, and transition files should be written

## **Optional Parameters**

- -f columnorderingfile A text file which gives the ordering for the names of the columns one per line
- -m labelmappingfile This option can specify a file which maps state IDs to descriptive names. The descriptive names can be appended to the state IDs when viewing the states in the browser. The format of this file is a two column tab delimited file. The first column contains each state ID without the state ordering letter prefix. The second column contains a descriptive name or mnemonic. If this file is not specified, then just the state IDs are displayed.
- -o stateorderingfile The contents of this file gives a user supplied ordering of states. Giving such a file will override the stateordering option if that is also specified. The format of a stateordering file is a two column tab delimited file. The first column is the old state number and the second column is the new state number in order. The state prefix should not be given.

```
-stateordering emission|transition,-color r,g,b, -holdcolumnorder,
```

-i outfileID, have the same meaning as in LearnModel

# **StatePruning**

# Description

This command takes as input a set of learned models generally learned across different initializations and then generates a nested set of models by pruning states from the highest scored model. The highest score model is the state with the highest estimated log-likelihood. States are greedily pruned from the highest scoring model. The criteria to a prune a state is that it causes the least decrease on the total distance for all the states in other models to their nearest state in the pruned model. If there is only one model, then distance to other states in that model are considered. This can be used as one heuristic to initialize model parameters to get a roughly comparable set of models across different number of states while biasing the learning procedure to avoid redundant or non-representative states. By default ChromHMM uses a simpler single pass method for producing nested initializations.

### **Usage**

```
StatePruning [-correlation] inputdir outputdir

Note items in [] are optional
```

### **Required Parameters**

inputdir - The name of the directory containing model files to eliminate. Only files with the prefix 'model\_' in the directory are used.

outputdir -The name of the output directory where the nested set of model files should go. Each model file is prefixed with the name 'elim\_'.

### **Optional Parameters**

-correlation - An optional flag indicating to use 1-correlation coefficient to define distance instead of the default of the euclidean distance

# Acknowledgements

The software includes the open source jheatchart library, the open source batik library, and RefSeq coordinates and CpG island coordinates obtained from the UCSC genome browser, and nuclear lamina data from (Guelen, Nature 2008) obtained through the UCSC genome browser.