

A quick guide to JSNMF algorithm with mouse brain (RNA+H3K4me3, Paired-Tag) data

Load data

```
addpath(' ../data')
load('h3k4me3_5k_10k.mat')
num_clu = length(unique(label));
```

Marker genes and housekeeping genes are given by user

```
marker_genes = ["Dnah12"; "Cfap299"; 'Flt1'; 'Slco1a4'; 'Inpp5d'; 'Hexb'; 'Slc1a3';
'Atp1a2'; 'Slc1a2'; 'Gpc5'; 'Prr51'; 'Plp1'; 'Rnf220'; 'Lhfp13'; 'Cacng4'; 'Erbb4';
'Kcnc2'; 'Reln'; 'Grin3a'; 'Adarb2'; 'Fam19a2'; 'Ahcy12'; 'Gm32647'; 'Shisa6';
'Epha6'; 'Galnt14'; 'Hs3st4'; 'Tshz2'; 'Fam19a1'; 'Il1rap12'; 'Cdh12'; 'Rgs6';
'Lingo2'; 'Prr16'; 'Zfp804b'; 'Pex51'; 'Gm26883'; 'Cdh18'; 'Gm28928'; 'Galnt16';
'Spag16'; 'Cfap43'; 'Wdr49'; 'Ebf1'; 'Mecom'; 'Ptprb'; 'Tgfbr1'; 'Zfhx3'; 'Apbb1ip';
'Gpc5'; 'Plpp3'; 'Slc1a2'; 'Igsf8'; 'St18'; 'Mog'; 'Mag'; 'St18'; 'Mbp'; 'Vcan';
'Tnr'; '6030443J06Rik'; 'Nxph1'; '6330411D24Rik'; 'Kcnmb2'; 'Grip1'; 'Gm45341';
'Gm45455'; 'Grip1'; 'Trpm3'; 'Rfx3'; 'Dgkh'; 'Hs3st4'; 'Ryr3'; 'Hs6st3'; 'Gm2164';
'Grik4'; 'Cpne7'; 'Gm2164'; 'Ryr3'; 'Hs6st3'; 'Sdk1'; 'Foxp2'; 'Garnl3hrm2'; 'Grm8';
'Vwc21'; 'Olfm3'; 'Grik1 Gm2164'; 'Gm28928'; 'Sgcz'; 'Prr16'; 'Chrm3'; 'Pdzn3';
'Kcnq5'; 'Unc5d'; 'Car10'; 'Pcdh15'; 'Nrg1'];
marker_genes = unique(marker_genes);
load('housekeeping_genes.mat')
housekeeping_genes = hp_genes;
```

Preparing for RAGI

```
load('h3k4me3_rna_RAGI.mat')
genes = genes_h3k4me3; data = rna_h3k4me3;
[genes_h3k4me3, rna_h3k4me3] = rna_ragi(barcodes, share_bd, genes, RNA,1);

[~, ~, ind1] = intersect(marker_genes,genes,'stable');
[~, ~, ind2] = intersect(housekeeping_genes,genes,'stable');
```

Run JSNMF

Parameter selection

```
addpath(' ../codes')
addpath(' ../external')
```

```
[alpha, gamma, Inits] = parameter_selection(X1, X2, label);
```

```
disp(['value of alpha:',num2str(alpha));  
disp(['value of gamma:',num2str(gamma));
```

```
value of alpha:0.36157  
value of gamma:3734.8227
```

Start time

```
tic  
[W1, W2, H1, H2, S, iter, objs] = jsnmf(X1, X2, alpha, gamma, Inits);  
disp('JSNMF runtime:');  
toc
```

```
iteration starts!  
number of iteration:10   obj:2167309.0214  
number of iteration:20   obj:2160203.2657  
number of iteration:30   obj:2157360.4942  
number of iteration:40   obj:2155750.9988  
number of iteration:50   obj:2154672.8747  
number of iteration:60   obj:2153795.968  
number of iteration:70   obj:2152994.5733  
number of iteration:80   obj:2152529.4533  
number of iteration:90   obj:2152224.5205  
converged!
```

```
JSNMF runtime:  
Elapsed time is 89.678800 seconds.  
ac: 0.8536      393/2684      nmi:0.8358  
ave_marker_gene: 0.4256      ave_hk_gini:0.1896      diff_gini:0.2361
```

Evaluating performance

```
% KNN on S  
A = Wtrim(S,50);  
% implement louvain clustering  
[clust,~,~] = getNCluster(A,num_clu,0,3,20);  
if length(unique(clust))== num_clu  
    [ac, nmi_value, ~] = CalcMetrics(label, clust);  
    [ave_mk_gini, ave_hk_gini, difgini] = RAGI(data,ind1,ind2,clust);  
else  
% using spectral clustering instead if number of clusering doesn't equals to the  
pointed number  
    [~, clust, ~] = SpectralClustering(A, num_clu);  
    [ac, nmi_value, ~] = CalcMetrics(label, clust);  
    [ave_mk_gini, ave_hk_gini, difgini] = RAGI(data,ind1,ind2,clust);  
end
```

Downstream analysis including visualization

visualization on S obtained from JSNMF

```
% load UMAP package
addpath(' ../umapFileExchange/umap')
addpath(' ../umapFileExchange/util')

term = '.';
label_name = label_h3k4me3(:,2);
label_name = strrep(label_name, '_', '\_');
colors = generateColors(max(length(unique(label_name))),length(unique(term)));

D_jsnfm = 1-S; D_jsnfm = D_jsnfm-diag(diag(D_jsnfm));
[reduction_jsnfm, ~, ~, ~] =
run_umap(D_jsnfm, 'metric', 'precomputed', 'min_dist', 0.68, 'n_neighbors', 12);
gscatter(reduction_jsnfm(:,1), reduction_jsnfm(:,2), label_name, colors, [], 4);
set(gca, 'xtick', [], 'ytick', []);
title('UMAP for JSNMF')

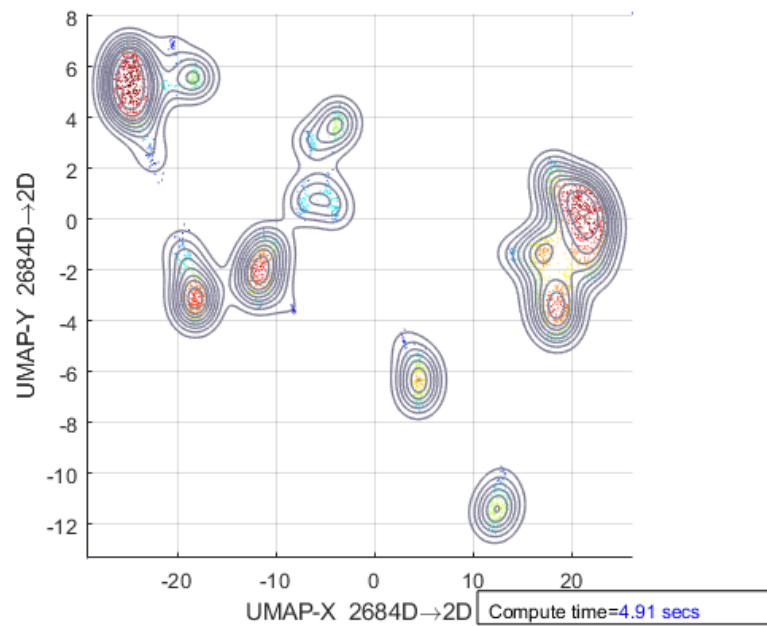
legend('Location', 'westoutside', 'Box', 'off', 'FontSize', 9.5);
legendmarkeradjust(16)
legend('boxoff')
```

Parallelizing UMAP with MATLAB's 8 assigned logical cores for nn_descent_tasks, sgd_tasks

Running basic (ub) reduction, v2.1.3
(ub=basic, us=supervised, ubt=template, ust=supervised template)

Warning: Start point not provided, choosing random start point.

```
UMAP(method=MEX, n_neighbors=12, n_components=2, metric='precomputed',
n_epochs=[], learning_rate=1, init=spectral, min_dist=0.68, spread=1,
set_op_mix_ratio=1, local_connectivity=1, repulsion_strength=1,
negative_sample_rate=5, transform_queue_size=4, a=0.341586509870361,
b=1.53964927967393, randomize=true, target_n_neighbors=-1,
target_metric='categorical', target_metric_kwds=[], target_weight=0.5,
verbose=true, initial_alpha=1, sparse_data=false, small_data=true,
distance_func='precomputed', dist_args=[])
```



UMAP reduction finished (cost 4.93 secs)

Finished basic (ub) reduction

