clone_fate_scanpy_separateConditions_SplitDonors

March 23, 2021

```
[1]: import pandas as pd
     import scanpy as sc
     import os
     from glob import glob
     from os.path import join, exists
     from src.config import ROOT_DIR
     from src.utils.parse_config import read_config_file
     os.chdir(ROOT_DIR)
     import anndata as ad
     import numpy as np
     import matplotlib.pyplot as plt
     from src.utils.scanpy_utils import *
    Project Directory: /data2/mito_lineage
    here
[2]: prefix = "jan21_2021"
[3]: config = read_config_file(join(ROOT_DIR, "Analysis",
     →"lineage_and_peakclusters", prefix, "config.yaml"))
     config
[3]: {'outdir':
     '/data2/mito_lineage/Analysis/lineage_and_peakclusters/results/jan21_2021/',
      'lineage_in': '/data2/mito_lineage/Analysis/multiplex/data/jan21_2021/chrM/pseu
     do/minC200_minAF0.01/numC25000_ispropFalse/flt3/',
      'aggregate_in': '/data2/isshamie/mito_lineage/data/processed/mtscATAC/jan21_202
     1/MTblacklist/reanalysis/outs/analysis/',
      'cell_names': ['/data2/mito_lineage/Analysis/multiplex/data/jan21_2021/chrM/P2_
     cellSNP minC200 minAF0.01/cellSNP.samples.tsv',
       '/data2/mito_lineage/Analysis/multiplex/data/jan21_2021/chrM/J2_cellSNP_minC20
     0 minAF0.01/cellSNP.samples.tsv'],
      'samples': 'J2,P2',
      'n_donors': 4,
      'only_donors': False,
      'nclones_values': '20,100',
      'prefix': 'jan21_2021',
      'genome': 'MTblacklist',
```

```
'name': 'cd34',
     'sample_moniker': ['Control', 'Flt31']}
[4]: dfs = {}
    for nclones_val in ["20"]:#confiq["nclones_values"].split(","):
        print(nclones_val)
        dfs[nclones_val] = pd.read_csv(config["outdir"]+f"/
     \neg cells_merged_lin_and_peak_nclones{nclones_val}.overlap_percent_normClone.
     full df = dfs["20"]
    full df.index = full df.index.astype("str")
    full_df.columns = full_df.columns.astype("str")
    df = full_df.drop(["Cluster", "Sample"], axis=0)
    ## Not including the different conditions into the same lineage. So (e.g. donor)
     \rightarrowA - lineage 2 is now D3-L2-Flt3l and D3-L2-Ctrl)
    ## Note the difference is the shape of the matrix:
    ## Before it was (#conditions*#cell clusters)-by-(lineages*donors),
    # Here it's (#Cell-clusters)-by-(lineage*donors*conditions)
    sample df = \{\}
    for ind, val in full_df.transpose().groupby("Sample"):
        val = val.set index("Cluster")
        val = val.drop("Sample", axis=1)
        cols = val.apply(lambda x: x.name+"_"+ind)
        val.columns = cols
        sample_df[ind] = val
    sample_df[ind]
    sample_df = pd.concat(sample_df.values(),axis=1)
    sample df.index = sample df.index.astype("str")
    sample_df.columns = sample_df.columns.astype("str")
    sample df
   20
[4]:
               0_0_J2
                                                      10_0_J2
                                                              10_1_J2 \
                         0_{1}J_{2}
                                   0_2_J2
                                             0_3_J2
    Cluster
            0.0588235 0.0714286 0.0408163
                                           0.117647
                                                     0.166667
                                                              0.0384615
    1
    2
            0.0392157 \quad 0.0714286 \quad 0.0204082 \quad 0.0588235 \quad 0.0333333 \quad 0.0384615
    3
            0.0980392 0.0714286 0.326531 0.0588235
                                                          0.1
                                                               0.192308
    4
             0.313725 0.0714286 0.0204082 0.117647
                                                          0.2 0.0769231
    5
            0.0384615
            6
                                                              0.0384615
    7
            0.0769231
             0.176471 0.0714286 0.0204082 0.0588235 0.0333333
                                                               0.115385
```

```
9
         0.0196078 0.0714286 0.0204082
                                            0.0588235
                                                        0.0333333
                                                                   0.0384615
10
         0.0196078
                     0.0714286
                                 0.244898
                                            0.0588235
                                                        0.0666667
                                                                   0.0769231
                                 0.102041
                                            0.0588235
11
         0.0392157
                     0.0714286
                                                        0.0666667
                                                                   0.0769231
12
         0.0196078
                     0.0714286
                                0.0204082
                                            0.0588235
                                                        0.0333333
                                                                   0.0384615
         0.0196078
                     0.0714286
                                0.0204082
                                            0.0588235
                                                        0.0333333
                                                                   0.0384615
13
14
         0.0196078
                    0.0714286
                                0.0612245
                                            0.0588235
                                                              0.1
                                                                    0.115385
        10_2_J2
                    10_3_J2
                               11_0_J2
                                           11_1_J2
                                                         7_1_P2
                                                                    7_2_P2 \
Cluster
1
            0.1
                   0.151515
                             0.0535714
                                         0.0909091
                                                        0.09375
                                                                  0.037037
2
           0.06
                 0.0606061
                             0.0178571
                                          0.030303
                                                                  0.296296
                                                          0.125
3
           0.12
                   0.030303
                             0.0892857
                                         0.0606061
                                                        0.09375
                                                                  0.111111
4
           0.16
                  0.0606061
                             0.0714286
                                          0.272727
                                                         0.0625
                                                                 0.0740741
5
           0.08
                  0.0909091
                             0.0535714
                                         0.0606061
                                                         0.0625
                                                                  0.037037
6
           0.08
                   0.030303
                              0.214286
                                         0.0606061
                                                          0.125
                                                                 0.0740741
7
           0.04
                  0.181818
                             0.0535714
                                          0.121212
                                                          0.125
                                                                  0.037037
            0.1
8
                  0.151515
                             0.0892857
                                         0.0909091
                                                         0.0625
                                                                  0.037037
9
           0.08
                  0.030303
                              0.107143
                                          0.030303
                                                         0.0625
                                                                 0.0740741
           0.02
10
                   0.030303
                             0.0892857
                                          0.030303
                                                        0.03125
                                                                  0.037037
11
           0.06
                   0.030303
                             0.0535714
                                          0.030303
                                                        0.03125
                                                                  0.037037
           0.04
                  0.0606061
                                                         0.0625
12
                             0.0714286
                                         0.0606061
                                                                  0.037037
           0.02
                  0.0606061
                             0.0178571
                                          0.030303
                                                        0.03125
                                                                  0.037037
13
14
           0.04
                  0.030303
                             0.0178571
                                          0.030303 ...
                                                        0.03125
                                                                 0.0740741
         7_3_P2
                     8_0_P2
                                8_1_P2 8_2_P2
                                                   9_0_P2
                                                               9_1_P2
                                                                           9_2_P2 \
Cluster
                             0.0540541
                                          0.05 0.055556
1
          0.125
                  0.111111
                                                           0.0882353
                                                                        0.0666667
2
         0.0625
                  0.111111
                               0.27027
                                           0.2
                                                0.0555556
                                                             0.338235
                                                                              0.3
3
         0.0625
                  0.111111
                             0.0540541
                                           0.1
                                                0.055556
                                                             0.132353
                                                                              0.2
4
         0.0625
                  0.0740741
                              0.027027
                                                0.0555556
                                                            0.0441176
                                                                             0.05
                                          0.05
5
         0.0625
                  0.0740741
                              0.027027
                                           0.1
                                                 0.111111
                                                            0.0441176
                                                                       0.0666667
         0.0625
                                          0.05
                                                 0.111111
6
                  0.037037
                             0.0810811
                                                            0.0882353
                                                                        0.066667
7
                              0.162162
         0.0625
                  0.111111
                                          0.05
                                                 0.111111
                                                            0.0735294
                                                                        0.0333333
                                                0.0555556
8
         0.0625
                  0.037037
                             0.0810811
                                          0.05
                                                            0.0147059
                                                                        0.0166667
9
          0.125
                   0.037037
                              0.027027
                                           0.1
                                                 0.111111
                                                            0.0294118
                                                                              0.1
10
         0.0625
                  0.0740741
                             0.0810811
                                          0.05
                                                0.055556
                                                            0.0735294
                                                                       0.0333333
         0.0625
                  0.111111
                             0.0540541
                                          0.05
                                               0.0555556
                                                            0.0294118
                                                                       0.0166667
11
                                                            0.0147059
12
         0.0625
                   0.037037
                              0.027027
                                          0.05
                                                0.055556
                                                                       0.0166667
13
         0.0625
                   0.037037
                              0.027027
                                          0.05
                                                0.055556
                                                            0.0147059
                                                                        0.0166667
         0.0625
                   0.037037
                              0.027027
                                          0.05
                                                0.0555556
                                                            0.0147059
                                                                       0.0166667
14
            9_3_P2
Cluster
1
         0.0588235
2
         0.0588235
3
         0.0588235
4
         0.0588235
```

```
5
          0.176471
6
         0.0588235
7
         0.0588235
8
         0.0588235
9
          0.117647
         0.0588235
10
         0.0588235
11
12
         0.0588235
13
         0.0588235
14
         0.0588235
```

[14 rows x 158 columns]

[5]:		Barcode	Donor	Condition
	0_0_J2	0	0	J2
	0_1_J2	0	1	J2
	0_2_J2	0	2	J2
	0_3_J2	0	3	J2
	10_0_J2	10	0	J2
				•••
	8_2_P2	8	2	P2
	9_0_P2	9	0	P2
	9_1_P2	9	1	P2
	9_2_P2	9	2	P2
	9_3_P2	9	3	P2

[158 rows x 3 columns]

1 Run with the sample-lineages separated as different barcodes

Not including the different conditions into the same lineage. So (e.g. donor A - lineage 2 is now D3-L2-Flt3l and D3-L2-Ctrl)

Note the difference is the shape of the matrix:

Before it was (#conditionsX#cell clusters)-by-(lineages*donors), Here it's (#CellXclusters)-by-(lineageXdonorsXconditions)

```
[6]: # def run_single_donor(df, curr_obs):
                      print("Donor", d)
          #
                      \#df = df.iloc[:, np.flatnonzero(sample_df.columns.isin(curr_obs.index))]
                      #run single_donor(curr_sample_df, curr_obs)
          #
                      adata = create_scanpy(df=df.transpose(), sample_df=curr_obs)
          #
                      sc.pl.highest_expr_genes(adata, n_top=20, )
          #
                      sc.tl.pca(adata, svd_solver='arpack')
                      sc.pl.pca(adata, color='1')
                      sc.tl.pca(adata, svd_solver='arpack')
                      sc.pl.pca(adata, color=list(adata.var_names.values)+['leiden', 'Donor', _
           \rightarrow 'Condition'7)
          #
                      sc.pp.neighbors(adata, n_neighbors=10, n_pcs=40)
          #
                      sc.tl.umap(adata)
                      sc.pl.umap(adata, color=adata.var_names)
          #
                      sc.tl.leiden(adata)
                      ax = sc.pl.umap(adata, color=list(adata.var_names.values)+['leiden', ___ | 
           → 'Donor', 'Condition'], show=False)
                      sc.tl.rank_genes_groups(adata, 'leiden', method='t-test')
                      sc.pl.rank_genes_groups(adata, n_genes=25, sharey=False)
                      c_ax = sc.pl.matrixplot(adata, adata.var_names, groupby='Condition', ___
           \rightarrow cmap='viridis', dendrogram=False,show=False)
                      c_ax = sc.pl.matrixplot(adata, adata.var_names, groupby='leiden',_
           → cmap='viridis', dendrogram=False, show=False)
                      ax1 = sc.pl.heatmap(adata, adata.var names, groupby='Condition', |
           → cmap='viridis', dendrogram=True, show=False)
                      ax2_ = sc.pl.heatmap(adata, adata.var_names, groupby='Condition',_
           → cmap='viridis', dendrogram=True, show=False)
                      ax3_ = sc.pl.heatmap(adata, adata.var_names, groupby='Donor',_
           → cmap='viridis', dendrogram=True, show=False)
                                  ax4 = sc.pl.heatmap(adata, adata.var_names, groupby='leiden'_
           \hookrightarrow, cmap='RdBu_r',
                                                                        vmin=0, vmax=0.1, dendrogram=True, show=False)
                                  ax5 = sc.pl.heatmap(adata, adata.var_names, groupby='Condition'
           \hookrightarrow, cmap='RdBu_r',
                                                                         vmin=0, vmax=0.1, dendrogram=True, show=False)
                      \#ax = sc.pl.heatmap(adata, adata.var_names, groupby=['Donor', 'Condition', United States and Condition', United States and C
           → 'leiden'], cmap='viridis', dendrogram=True)
                      ### Violinplot grouping by graph clusters, showing how much of that,
           → feature is in that cluster (first 5 cell types shown)
                      sc.pl.violin(adata, adata.var_names, groupby='leiden',size=4,)
          #
                      sc.pl.violin(adata, adata.var_names, groupby='Donor', size=4,)
```

```
# return

# for d, curr_obs in obs_data.groupby("Donor"):
# print("Donor", d)

# curr_sample_df = sample_df.iloc[:, np.flatnonzero(sample_df.columns.

isin(val.index))]

# run_single_donor(curr_sample_df, curr_obs)

# curr_sample_df
```

```
[]: for d, curr_obs in obs_data.groupby("Donor"):
        print("Donor", d)
        df = sample_df.iloc[:, np.flatnonzero(sample_df.columns.isin(curr_obs.
     →index))]
        #run_single_donor(curr_sample_df, curr_obs)
        adata = create_scanpy(df=df.transpose(), sample_df=curr_obs)
        sc.pl.highest_expr_genes(adata, n_top=20, )
        print("PCA")
        sc.tl.pca(adata, svd_solver='arpack')
        sc.pl.pca(adata, color=list(adata.var names.values)+['Donor', 'Condition'])
        sc.pp.neighbors(adata, n_neighbors=10, n_pcs=40)
        print("UMAP & Constructing Fate clusters")
        sc.tl.umap(adata)
        sc.tl.leiden(adata)
        ax = sc.pl.umap(adata, color=list(adata.var_names.values)+['leiden',_
     → 'Donor', 'Condition'], show=False)
        sc.tl.rank_genes_groups(adata, 'leiden', method='t-test')
        sc.pl.rank_genes_groups(adata, n_genes=25, sharey=False)
        c_ax = sc.pl.matrixplot(adata, adata.var_names, groupby='Condition',_
     c_ax = sc.pl.matrixplot(adata, adata.var_names, groupby='leiden',u
     ax1_ = sc.pl.heatmap(adata, adata.var_names, groupby='Condition',_
     ax2_ = sc.pl.heatmap(adata, adata.var_names, groupby='leiden',_
     ax4 = sc.pl.heatmap(adata, adata.var_names, groupby='leiden'
     \hookrightarrow, cmap='RdBu_r',
                               vmin=0, vmax=0.1, dendrogram=True, show=False)
        #
             ax5 = sc.pl.heatmap(adata, adata.var_names, groupby='Condition'
     \rightarrow, cmap='RdBu r',
                               vmin=0, vmax=0.1, dendrogram=True, show=False)
        \#ax = sc.pl.heatmap(adata, adata.var_names, groupby=['Donor', 'Condition', \sqcup
     → 'leiden'], cmap='viridis', dendrogram=True)
```

```
### Violinplot grouping by graph clusters, showing how much of that feature

is in that cluster (first 5 cell types shown)

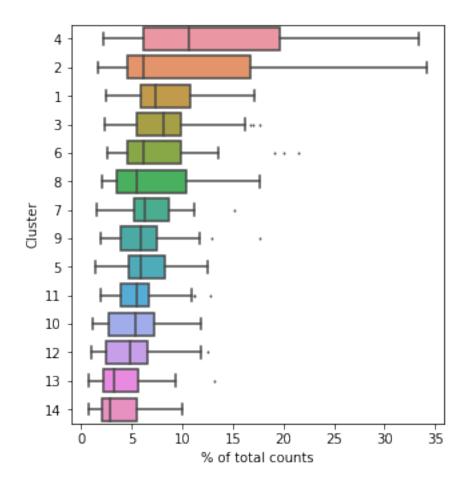
sc.pl.violin(adata, adata.var_names, groupby='leiden',size=4,)

sc.pl.violin(adata, adata.var_names, groupby='Donor',size=4,)

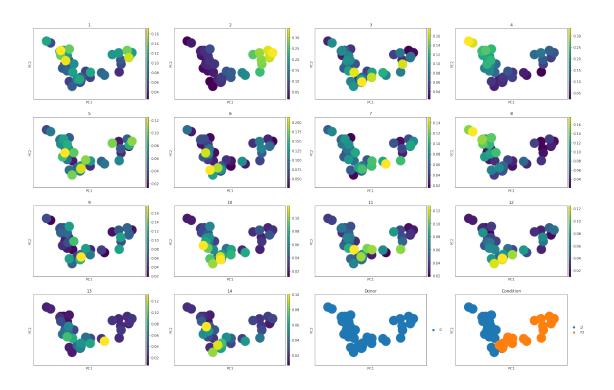
print("\n")

print("\n")
```

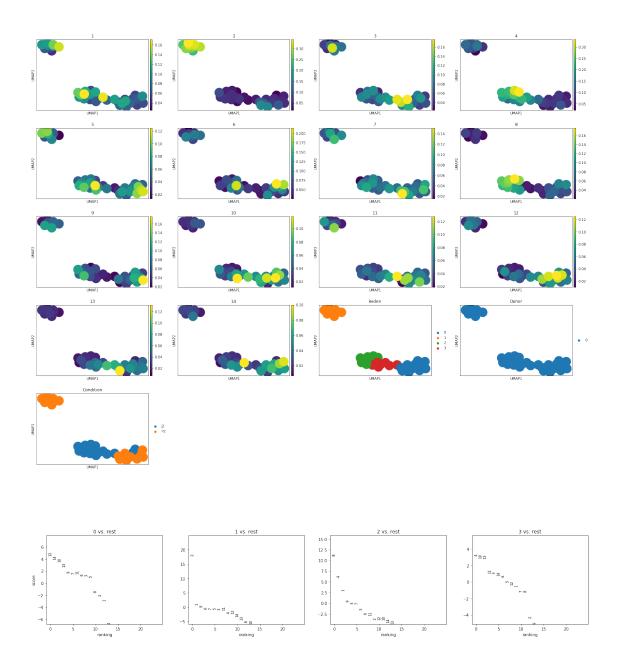
Donor 0



```
... storing 'Barcode' as categorical
... storing 'Donor' as categorical
... storing 'Condition' as categorical
PCA
```

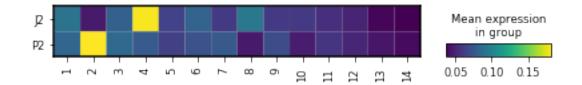


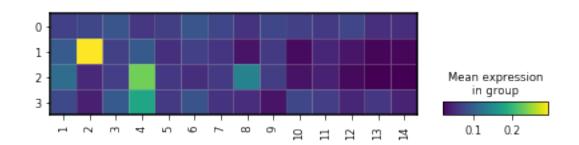
UMAP & Constructing Fate clusters

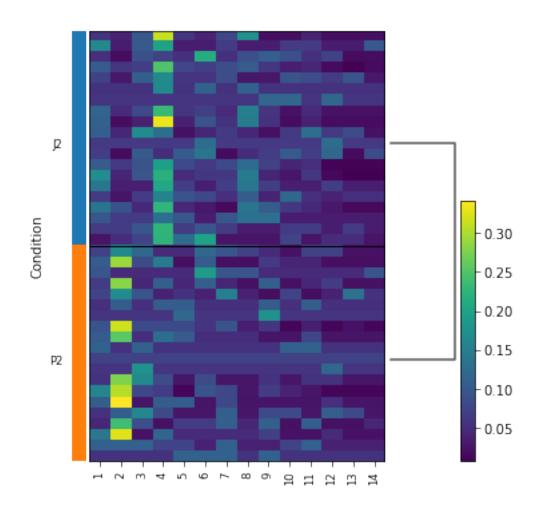


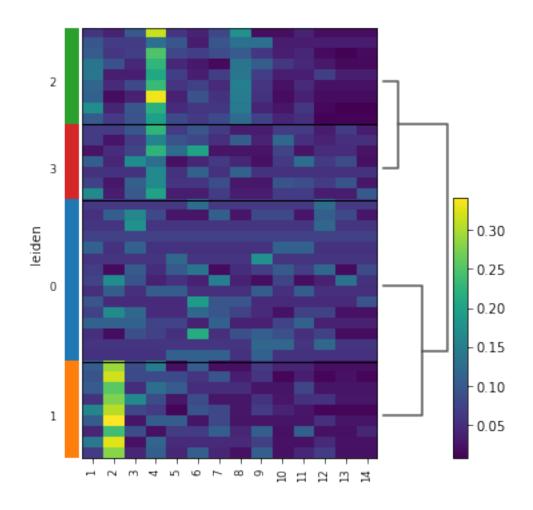
WARNING: dendrogram data not found (using key=dendrogram_Condition). Running `sc.tl.dendrogram` with default parameters. For fine tuning it is recommended to run `sc.tl.dendrogram` independently.

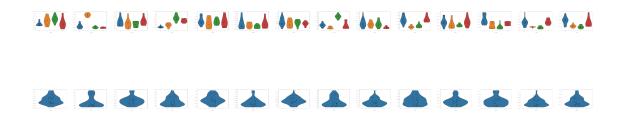
WARNING: dendrogram data not found (using key=dendrogram_leiden). Running `sc.tl.dendrogram` with default parameters. For fine tuning it is recommended to run `sc.tl.dendrogram` independently.



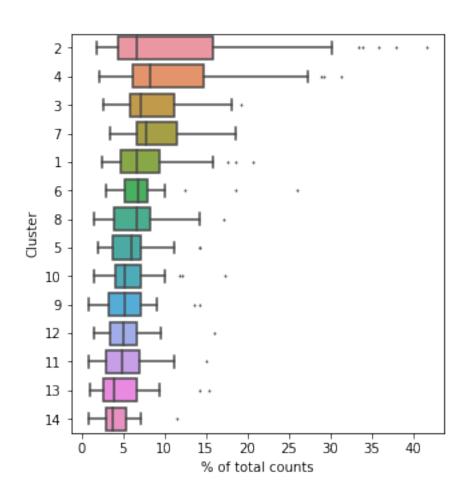








Donor 1

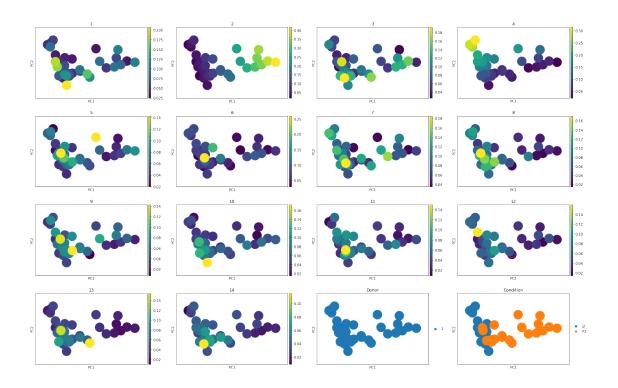


```
... storing 'Barcode' as categorical
```

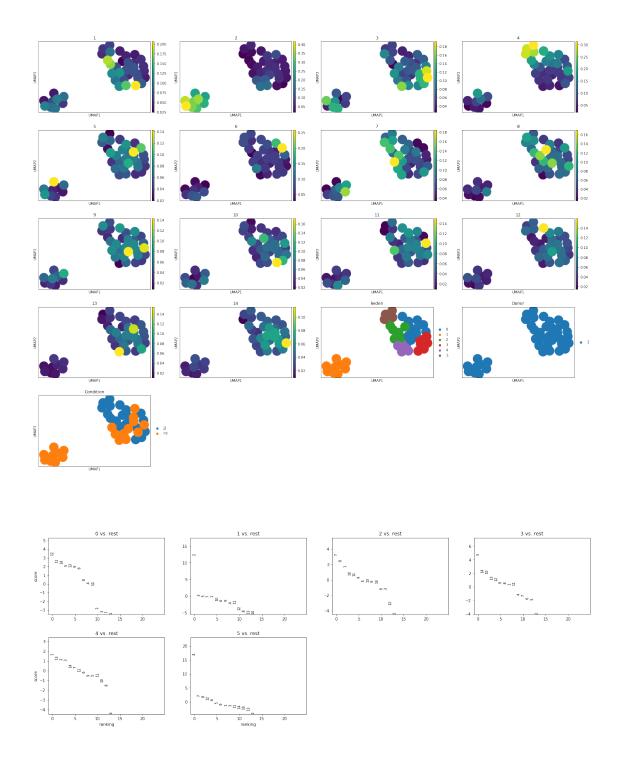
PCA

^{...} storing 'Donor' as categorical

^{...} storing 'Condition' as categorical



UMAP & Constructing Fate clusters



WARNING: dendrogram data not found (using key=dendrogram_Condition). Running `sc.tl.dendrogram` with default parameters. For fine tuning it is recommended to run `sc.tl.dendrogram` independently.

WARNING: dendrogram data not found (using key=dendrogram_leiden). Running `sc.tl.dendrogram` with default parameters. For fine tuning it is recommended to run `sc.tl.dendrogram` independently.

