Understanding Outputs

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Overview

We will analyze example output found in SLIDE/test/out/0.01_1_out

This is the general workflow:

- 1. Evaluate latent factor performance using ControlPerformancePlot.png
- 2. Look at top features in each latent factor using plotSigGenes_marginals.png and plotSigGenes.png
- 3. Look at interactions between latent factors with plotInteractions.png
- 4. Look at gene lists in gene_list_Z.txt files
- 5. View SLIDE results and parameters in SLIDE LFs.rds
- 6. View correlation networks of significant latent factors after running plotCorrelationNetworks

Evaluate latent factor performance using ControlPerformancePlot.png

This plot shows how the significant latent factors perform relative to their knockoff counterparts:

- The red line is the true performance of the significant latent factors (either correlation for continuous response/Y or AUC for binary response/Y).
- The blue density shows the performance of the knockoffs for the stand-alone/marginal latent factors
- The green density shows the performance of the true marginal latent factors paired with knockoffs for the interaction latent factors

A good model should have:

- **performance red line should be highest** (better performance) and to the right of both the blue and green densities this indicates that there signal in the significant latent factors.
- green density should be slightly higher than blue, since we are using the true marginal latent factors paired with knockoff interaction latent factors we are trying to evaluate whether the interaction latent factors are meaningful
- blue density has the lowest performance, as knockoffs for our marginal latent factors should not have signal

knitr::include_graphics("examples/out/0.01_1_out/ControlPerformancePlot.png")

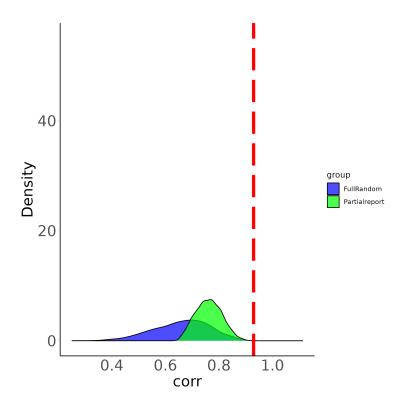


Figure 1: Significant Latent Factor Performance

Above is an example of a model with great performance. In some cases, you may find that the green density overlaps perfectly with the blue - in this case, the interaction latent factors are not meaningful. If you find that all three (blue, green and red) perfectly overlap, then this indicates a lack of signal in the significant latent factors and there are a few options:

- 1. Increase spec parameter (increase FDR control for choosing latent factors)
- 2. Re-cluster latent factors using different delta and lambda parameters it is possible the latent factors are a poor fit to the data (too many latent factors will overfit = increase delta; too few latent factors will be over-clustered = decrease delta)

The data for this plot is stored in ControlPerformance.rds; the ControlPerformancePlot.rds has the plot object for the ggplot above.

Look at top features in each latent factor using plotSigGenes_marginals.png and plotSigGenes.png

The plotSigGenes_marginals.png shows the top features in the stand-alone/marginal significant latent factors. By default, we show the top 10 features by loading (features that contribute most to the latent factor), as well as the top 10 univariate features (features that have high correlation or AUC with response/Y). Each latent factor shows 20 features - the bottom 10 are the highest loading and the top 10 are highest univariate.

knitr::include_graphics("examples/out/0.01_1_out/plotSigGenes_marginals.png")

SLIDE Marginal Variables

| | 26 | 45 Significant L | 70 atent Factor | 77 |
|--|-------------|---------------------|--------------------|-------------|
| | c.4.TCIM | c.1.APOE | c.3.WISP2 | c.14.PRX |
| | c.10.FOSB | c.3.GSN | c.1.APOE | c.2.SELE |
| | c.3.POSTN | c.3.IGFBP2 | c.2.CLU | c.7.LYZ |
| Ge | c.3.WIF1 | c.3.STC2 | c.3.STC2 | c.3.WISP2 |
| Genes Associated with Significant Latent Factors | c.3.STC2 | c.13.CCDC80 | c.6.CCL19 | c.1.APOE |
| Ass | c.3.TNC | c.3.TNC | c.13.CCDC80 | c.14.CFD |
| ocia | c.3.SAA1 | c.10.FOS | c.5.IGFBP5 | c.6.SFRP4 |
| ted | c.3.STMN2 | c.3.STMN2 | c.3.TNC | c.16.KRT6B |
| with | c.12.AREG | c.6.CXCL9 | c.10.FOS | c.7.CCL8 |
| Sig | c.7.IFI27 | c.15.DCD | c.11.KRT15 | c.13.CRIP1 |
| nific | c.17.CCL4L2 | c.7.IFI27 | c.5.FABP4 | c.3.SFRP4 |
| ant | c.3.GPX3 | c.3.WIF1 | c.15.CD59 | c.4.RGS5 |
| Late | c.8.CTGF | c.5.TM4SF1 | c.14.CFD | c.16.KRT15 |
| int F | c.8.MT1X | c.10.FOSB | c.15.CD164 | c.16.CXCL14 |
| acto | c.6.MT1X | c.16.IFI27 | c.5.ACTA2 | c.1.CXCL14 |
| Ors | c.5.MT1X | c.11.IFI27 | c.5.POSTN | |
| | c.4.MT1X | c.1.IFI27 | c.15.PPP1R1B | |
| | c.3.MT1X | c.0.IFI27 | c.15.KRT7 | |
| | | | c.15.AQP5 | |

Figure 2: Marginal Significant Latent Factors

We also include plots showing both the marginal and interaction latent factors in this format; here, the bolded/italicized latent factors are the stand-alone/marginal latent factors and the plain text are the interactions.

knitr::include_graphics("examples/out/0.01_1_out/plotSigGenes.png")

| | | | c.13.DES | | | | c.16.LGALS1 | |
|--|-------------|-------------|-------------|------------|--------------|-------------|-------------|---------------|
| | | | c.13.ACTG2 | | c.15.AQP5 | | c.16.S100A2 | |
| | c.3.MT1X | c.0.IFI27 | c.13.PCP4 | | c.15.KRT7 | | c.16.KRT17 | |
| actors | c.4.MT1X | c.1.IFI27 | c.13.RERGL | c.13.DUSP1 | c.15.PPP1R1B | | c.16.CALML3 | c.16.FOS |
| | c.5.MT1X | c.11.IFI27 | c.13.CNN1 | c.13.CYR61 | c.5.POSTN | | c.16.APOE | c.16.HES1 |
| | c.6.MT1X | c.16.IFI27 | c.13.BCAM | c.17.GZMB | c.5.ACTA2 | c.1.CXCL14 | c.16.S100A6 | c.2.RBP7 |
| ent | c.8.MT1X | c.10.FOSB | c.13.TPM1 | c.13.APOE | c.15.CD164 | c.16.CXCL14 | c.16.IGFBP7 | c.8.IGFBP5 |
| Associated with Significant Latent Factors | c.8.CTGF | c.5.TM4SF1 | c.13.SMTN | c.13.ACTC1 | c.14.CFD | c.16.KRT15 | c.16.KRT75 | c.2.SELE |
| | c.3.GPX3 | c.3.WIF1 | c.13.MYLK | c.13.EGR1 | c.15.CD59 | c.4.RGS5 | c.16.KRT6B | c.4.IFITM1 |
| | c.17.CCL4L2 | c.7.IFI27 | c.13.CSRP1 | c.4.DCN | c.5.FABP4 | c.3.SFRP4 | c.16.MYC | c.16.SERPINB2 |
| | c.7.IFI27 | c.15.DCD | c.13.CCDC80 | c.14.PRX | c.11.KRT15 | c.13.CRIP1 | c.10.FOS | c.16.FGFBP1 |
| | c.12.AREG | c.6.CXCL9 | c.3.STC2 | c.13.ACTA1 | c.10.FOS | c.7.CCL8 | c.10.TXNIP | c.5.CFH |
| | c.3.STMN2 | c.3.STMN2 | c.1.APOE | c.8.TAGLN | c.3.TNC | c.16.KRT6B | c.10.KLF6 | c.11.AREG |
|)Clai | c.3.SAA1 | c.10.FOS | c.10.FOSB | c.3.STC2 | c.5.IGFBP5 | c.6.SFRP4 | c.6.CCL19 | c.3.GSN |
| 455 | c.3.TNC | c.3.TNC | c.3.POSTN | c.1.APOE | c.13.CCDC80 | c.14.CFD | c.3.STC2 | c.1.APOE |
| celles | c.3.STC2 | c.13.CCDC80 | c.13.RGS5 | c.13.TIMP3 | c.6.CCL19 | c.1.APOE | c.2.CLU | c.3.WISP2 |
| <u> </u> | c.3.WIF1 | c.3.STC2 | c.4.TCIM | c.13.HES1 | c.3.STC2 | c.3.WISP2 | c.1.APOE | c.3.POSTN |
| | c.3.POSTN | c.3.IGFBP2 | c.7.IFI27 | c.17.MTCO1 | c.2.CLU | c.7.LYZ | c.3.WISP2 | c.7.LYZ |
| | c.10.FOSB | c.3.GSN | c.13.MT1X | c.13.FLNA | c.1.APOE | c.2.SELE | c.16.SAT1 | c.17.MTCO1 |
| | c.4.TCIM | c.1.APOE | c.13.TIMP3 | c.13.CCL2 | c.3.WISP2 | c.14.PRX | c.3.POSTN | c.3.ADH1B |

Significant Latent Factors - Marginals (bold/italic) and Interactions

Figure 3: Marginal (bolded) and Interaction (plain) Significant Latent Factors

60 70 Significant Latent Factor 82

Look at interactions between latent factors with plotInteractions.png

This plot shows a graphical representation of how the interacting latent factors are paired with the marginals. Here, Z97 interacts with both Z147 and Z109; Z46, Z58, Z129, and Z137 have no interactions

knitr::include_graphics("examples/out/0.01_1_out/plotInteractions.png")



Figure 4: Latent Factor Interactions

Look at gene lists in gene_list_Z.txt files

These outputs provide a table of the top 20 features (same as above in the text plots), and provide the associated latent factor loadings A_loading and correlation/AUC corrs, as well as the association of each feature with the response color. For the color, Red is associated with a higher response/Y (or Y = 1 for binary) and Blue is associated with a lower response/Y (or Y = 0 for binary).

```
gene_list = read.csv("examples/out/0.01_1_out/gene_list_Z70.txt",
    row.names = 1)
knitr::kable(gene_list)
```

| | A_loading | corrs | color |
|--------------|-----------|------------|-------|
| c.15.AQP5 | 1.0000000 | -0.5396689 | Blue |
| c.15.KRT7 | 1.0000000 | 0.0387969 | Red |
| c.15.PPP1R1B | 1.0000000 | -0.5174597 | Blue |
| c.5.POSTN | 0.2546804 | 0.1633278 | Red |
| c.5.ACTA2 | 0.2097075 | -0.0135018 | Blue |
| c.15.CD164 | 0.2054907 | -0.3351158 | Blue |
| c.14.CFD | 0.1918422 | -0.2628597 | Blue |
| c.15.CD59 | 0.1816671 | -0.1307760 | Blue |
| c.5.FABP4 | 0.1496629 | -0.4573178 | Blue |
| c.11.KRT15 | 0.1481397 | 0.1398086 | Red |
| c.10.FOS | 0.0146592 | 0.6363250 | Red |
| c.3.TNC | 0.0578869 | 0.6215166 | Red |
| c.5.IGFBP5 | 0.0436693 | -0.5971263 | Blue |
| c.13.CCDC80 | 0.1019874 | 0.5884155 | Red |
| c.6.CCL19 | 0.0238157 | 0.5849312 | Red |
| c.3.STC2 | 0.0469672 | 0.5823180 | Red |
| c.2.CLU | 0.0029181 | -0.5696873 | Blue |
| c.1.APOE | 0.0111260 | -0.5548789 | Blue |
| c.3.WISP2 | 0.0065220 | -0.5531367 | Blue |
| | | | |

View SLIDE results and parameters in SLIDE_LFs.rds

We can see the parameters that were used to run SLIDE in SLIDE_LFs.rds

```
SLIDE_LFs = readRDS("examples/out/0.01_1_out/SLIDE_LFs.rds")
names(SLIDE_LFs)
```

```
#> [1] "SLIDE_res" "SLIDE_param" "marginal_vals" "interaction"
```

We can see the parameters:

```
knitr::kable(SLIDE_LFs$SLIDE_param)
```

| | x |
|--------|-----|
| method | 4.0 |
| spec | 0.3 |

| | 3 |
|---------------|-------|
| fdr | 0.1 |
| $_{ m niter}$ | 500.0 |
| f_size | 24.0 |

View correlation networks of significant latent factors after running plotCorrelationNetworks

After running optimizeSLIDE, we can run plotCorrelationNetworks to generate correlation network plots of the top features in each latent factor.

- Node color: Red nodes are associated with higher response/Y (for continuous Y) or Y = 1 (for binary Y); blue nodes are associated with lower response/Y (for continuous Y) or Y = 0 (for binary Y). This is the same color code as in the plotSigGenes.png plots above.
- Edge color: Purple edges are positive correlations and green edges are negative correlations between features



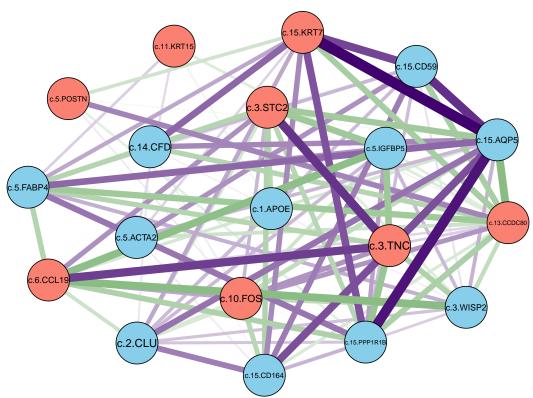


Figure 5: Correlation Network Example