Cancer programs - Kinker et al

```
library(Seurat)
library(plyr)
library(pheatmap)
library(RColorBrewer)
rhps <- read.delim('../GSKinker_RHPs.txt', check.names = F)</pre>
head(rhps)
     Cell Cycle - G1/S Cell Cycle - G2/M Skin Pigmentation
                                                                 EMT I
                                                                          EMT II
## 1
              HIST1H4C
                                     AURKA
                                                      PHACTR1
                                                                  DKK1
                                                                            MYL9
## 2
                 CLSPN
                                     CENPF
                                                          DCT
                                                                PMEPA1 SERPINE1
## 3
                  ATAD2
                                      PLK1
                                                         MITF C12orf75
                                                                           THBS1
## 4
                                                                 CYR61
                  E2F1
                                    TOP2A
                                                       CHCHD6
                                                                             FN1
## 5
                  HELLS
                                                          MBP
                                                                   IL8
                                     UBE2C
                                                                            IL32
                                                        GDF15
## 6
                   RRM2
                                      ASPM
                                                                SORBS2
                                                                           LAMC2
     IFN Response EMT III p53-Dependent Senescence Epithelial Senescence
## 1
            ISG20
                    LAMA3
                                              CDKN1A
                                                                        SLPI
## 2
            IFIT3
                    LAMB3
                                               NEAT1
                                                                       LCN2
## 3
            ISG15
                    LAMC2
                                               POLD4
                                                                       ELF3
## 4
             OASL CDKN1A
                                                MXD4
                                                                     S100A9
## 5
            IFIT1
                   FAM83A
                                           MMP24-AS1
                                                                       S100P
## 6
            IFIT2
                                               PNRC1
                    ITGA2
                                                                        SAA1
     Stress Response Protein Maturation Proteasomal Degradation
## 1
               DDIT3
                                    HSPA5
                                                             PSMA3
## 2
              SNHG12
                                     RPN2
                                                             PSMC4
## 3
              SLC3A2
                                  SLC3A2
                                                             PRDX1
            PPP1R15A
                                    PDIA3
                                                             PSMC2
## 5
             GADD45B
                                LGALS3BP
                                                            EIF4A3
             GADD45A
                                    MCM7
                                                              MDH1
subdataset <- readRDS('../tmp/crc_smc.malignantcells.Rds')</pre>
head(subdataset@meta.data); nrow(subdataset@meta.data) # 17334
##
                                        nCount_RNA nFeature_RNA
## AAACCTGCATACGCCG-1-PM-PS-0001-T-A1
                                             35998
                                                            4823 PM-PS-0001-T-A1
## AAACCTGGTCGCATAT-1-PM-PS-0001-T-A1
                                             31383
                                                            5252 PM-PS-0001-T-A1
## AAACCTGTCCCTTGCA-1-PM-PS-0001-T-A1
                                              7302
                                                            1713 PM-PS-0001-T-A1
## AAACGGGAGGGAAACA-1-PM-PS-0001-T-A1
                                              3759
                                                            1233 PM-PS-0001-T-A1
## AAACGGGGTATAGGTA-1-PM-PS-0001-T-A1
                                             23097
                                                            3874 PM-PS-0001-T-A1
## AAAGATGAGGCCGAAT-1-PM-PS-0001-T-A1
                                                            3282 PM-PS-0001-T-A1
##
                                        Patient Sample Cell_subtype RNA_snn_res.0.3
## AAACCTGCATACGCCG-1-PM-PS-0001-T-A1
                                          SMC01 SMC01-T
                                                                 CMS2
## AAACCTGGTCGCATAT-1-PM-PS-0001-T-A1
                                          SMC01 SMC01-T
                                                                 CMS2
                                                                                     \cap
```

SMC01 SMC01-T

SMC01 SMC01-T

CMS2

CMS2

2

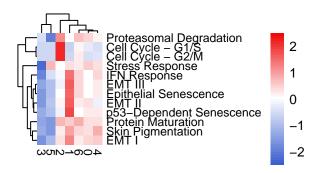
1

AAACCTGTCCCTTGCA-1-PM-PS-0001-T-A1

AAACGGGAGGAAACA-1-PM-PS-0001-T-A1

```
## AAACGGGGTATAGGTA-1-PM-PS-0001-T-A1
                                        SMC01 SMC01-T
                                                               CMS2
## AAAGATGAGGCCGAAT-1-PM-PS-0001-T-A1
                                        SMCO1 SMCO1-T
                                                               CMS2
## [1] 17334
label \leftarrow subdataset@meta.data[, c(3,4,7)]
colnames(label)[3] <- "res_0.3"</pre>
label <- subset(label, !res_0.3 %in% c('7'))</pre>
label <- droplevels(label)</pre>
head(label, n=3); nrow(label) # 17276
##
                                              Library Patient res_0.3
## AAACCTGCATACGCCG-1-PM-PS-0001-T-A1 PM-PS-0001-T-A1
                                                        SMC01
## AAACCTGGTCGCATAT-1-PM-PS-0001-T-A1 PM-PS-0001-T-A1
                                                         SMC01
                                                                     2
## AAACCTGTCCCTTGCA-1-PM-PS-0001-T-A1 PM-PS-0001-T-A1
                                                        SMC01
## [1] 17276
rhp_scores <- data.frame(matrix(ncol = length(levels(label$res_0.3)), nrow = ncol(rhps)))</pre>
colnames(rhp_scores) <- levels(label$res_0.3)</pre>
rownames(rhp_scores) <- colnames(rhps)</pre>
rhp_scores
                            0 1 2 3 4 5 6
## Cell Cycle - G1/S
                            NA NA NA NA NA NA
## Cell Cycle - G2/M
                          NA NA NA NA NA NA
                          NA NA NA NA NA NA
## Skin Pigmentation
## EMT I
                          NA NA NA NA NA NA
                       NA NA NA NA NA NA NA
NA NA NA NA NA NA NA
## EMT II
## IFN Response
## EMT III
                           NA NA NA NA NA NA
## p53-Dependent Senescence NA NA NA NA NA NA NA
## Epithelial Senescence \, NA NA NA NA NA NA NA
## Proteasomal Degradation NA NA NA NA NA NA NA
for (clst in colnames(rhp_scores)) {
  tmpObj <- subset(subdataset, idents = clst)</pre>
  tmpExpr <- as.matrix(GetAssayData(tmpObj, slot = 'data'))</pre>
  tmpExprMean <- data.frame(row.names = rownames(tmpExpr),</pre>
                            Symbol = unlist(lapply(rownames(tmpExpr), function (x) unlist(strsplit(as.c.
                            cluster = rowMeans(tmpExpr))
  head(tmpExprMean)
  for (rhp in colnames(rhps)) {
    rhp_genes <- as.character(rhps[, rhp][rhps[, rhp] != ""])</pre>
    tmpExprMean_rhp_genes <- subset(tmpExprMean, Symbol %in% rhp_genes)</pre>
    head(tmpExprMean_rhp_genes)
    rhp_scores[rhp, clst] <- mean(tmpExprMean_rhp_genes$cluster)</pre>
```

```
}
}
head(rhp_scores, n=3)
##
                                         1
## Cell Cycle - G1/S 0.08063864 0.05219887 0.2878732 0.05997496 0.06871475
## Cell Cycle - G2/M 0.07644820 0.06443931 0.3475173 0.06596869 0.08088368
## Skin Pigmentation 0.22521813 0.26241780 0.2219094 0.12645031 0.25496137
##
## Cell Cycle - G1/S 0.05527019 0.1178972
## Cell Cycle - G2/M 0.06457555 0.1415242
## Skin Pigmentation 0.10796311 0.2306110
min(rhp_scores)
## [1] 0.05219887
max(rhp_scores)
## [1] 1.036172
pheatmap(rhp_scores, scale = 'row', #filename = '../res_various/res_0.3/gskinker_rhps.pdf',
         treeheight_row = 10, treeheight_col = 10,
         cellwidth = 7, cellheight = 7, fontsize_row = 9, fontsize_col = 9, border_color = NA,
         clustering_distance_rows = 'euclidean', clustering_distance_cols = 'euclidean', clustering_met
         breaks = c(-25:25)/10,
         color = colorRampPalette(c("#3a5fcd", 'white', "#ee0000"))(n = 51))
```



#dev.off()