NETWORK-BASED DATA ANALYSIS

ELEONORA GIULIANI



DATASET GSE43837

19 human breast cancer brain metastases

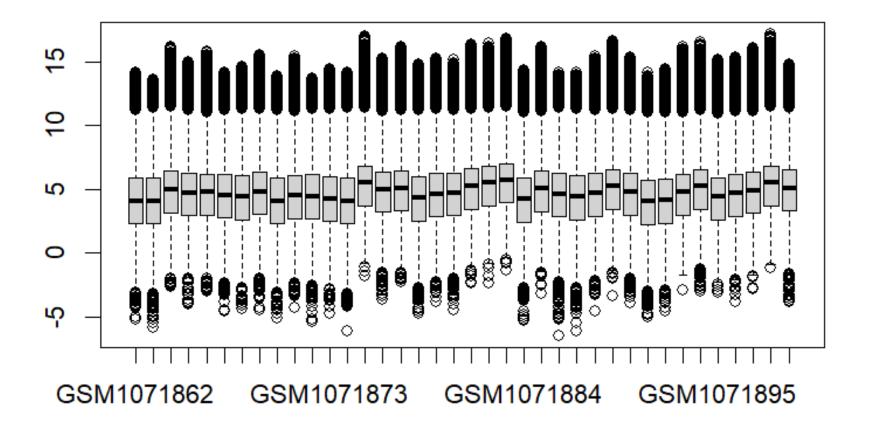
```
GSM1071862 brain_met_J3
GSM1071863 brain_met_BT25
GSM1071864 brain_met_J12
GSM1071865 brain_met_BT16
GSM1071866 brain met J8
GSM1071867 brain met BT34
GSM1071868 brain_met_J1
GSM1071869 brain_met_BT11
GSM1071870 brain_met_J2
GSM1071871 brain_met_BT30
GSM1071872 brain met J4
GSM1071873 brain_met_BT57
GSM1071874 brain_met_J5
GSM1071875 brain_met_BT51
GSM1071876 brain_met_BT31
GSM1071877 brain_met_J7
GSM1071878 brain_met_J10
GSM1071879 brain_met_J11
GSM1071880 brain met BT40
```

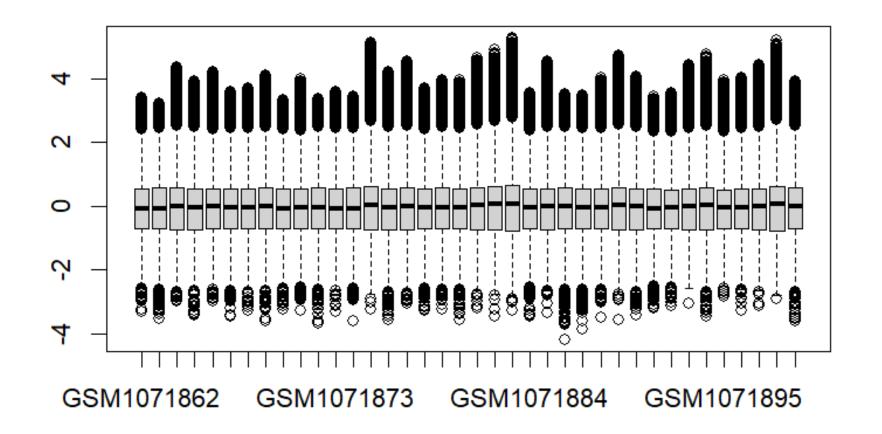
19 nonmetastatic primary human breast tumors

```
GSM1071881 breast primary MGH16
GSM1071882 breast_primary_MGH42
GSM1071883 breast primary MGH46
GSM1071884 breast_primary_MGH133
GSM1071885 breast_primary_MGH153
GSM1071886 breast primary MGH351
GSM1071887 breast_primary_MGH1104
GSM1071888 breast_primary_MGH574
GSM1071889 breast_primary_MGH434
GSM1071890 breast primary MGH450
GSM1071891 breast_primary_MGH421
GSM1071892 breast_primary_MGH482
GSM1071893 breast primary MGH963
GSM1071894 breast_primary_MGH455
GSM1071895 breast_primary_MGH1084
GSM1071896 breast primary MGH1038
GSM1071897 breast_primary_MGH1057
GSM1071898 breast_primary_MGH674
GSM1071899 breast_primary_MGH1102
```

Goal: Comparison of the gene expression between the two groups

PRE-PROCESSING



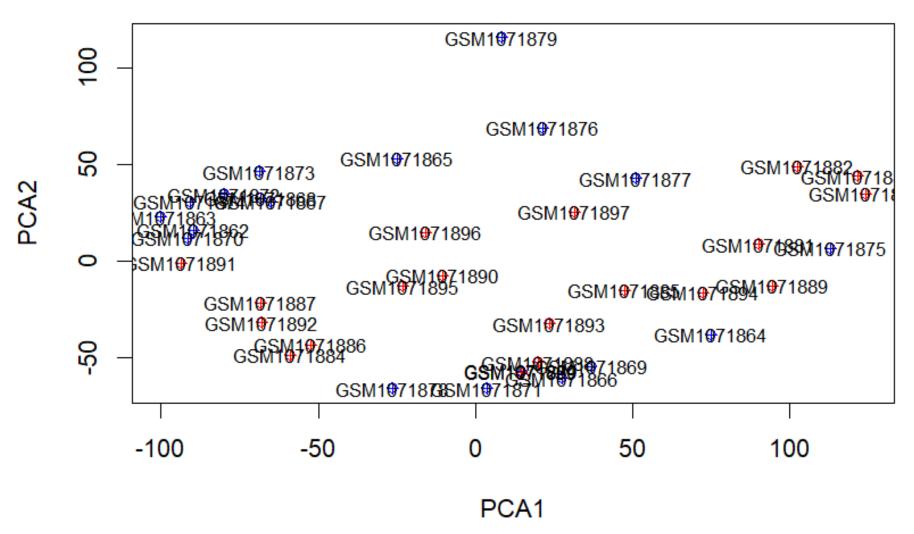


non-normalized dataset

normalized dataset

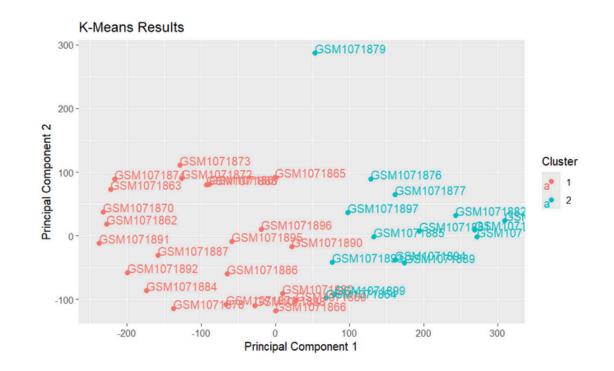
UNSUPERVISED LEARNING METHODS

PCA for components 1&2

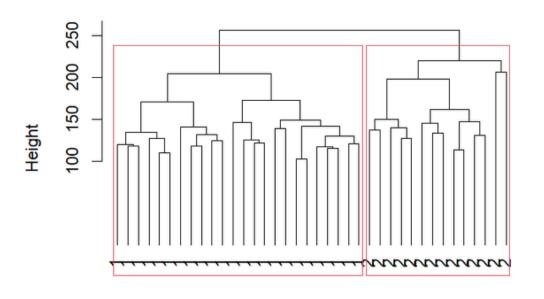


Red: nonmetastatic primary breast cancer (GSM1071881-GSM1071899)

Blue: breast cancer brain metastases (GSM1071862-GSM1071880)



Cluster Dendrogram



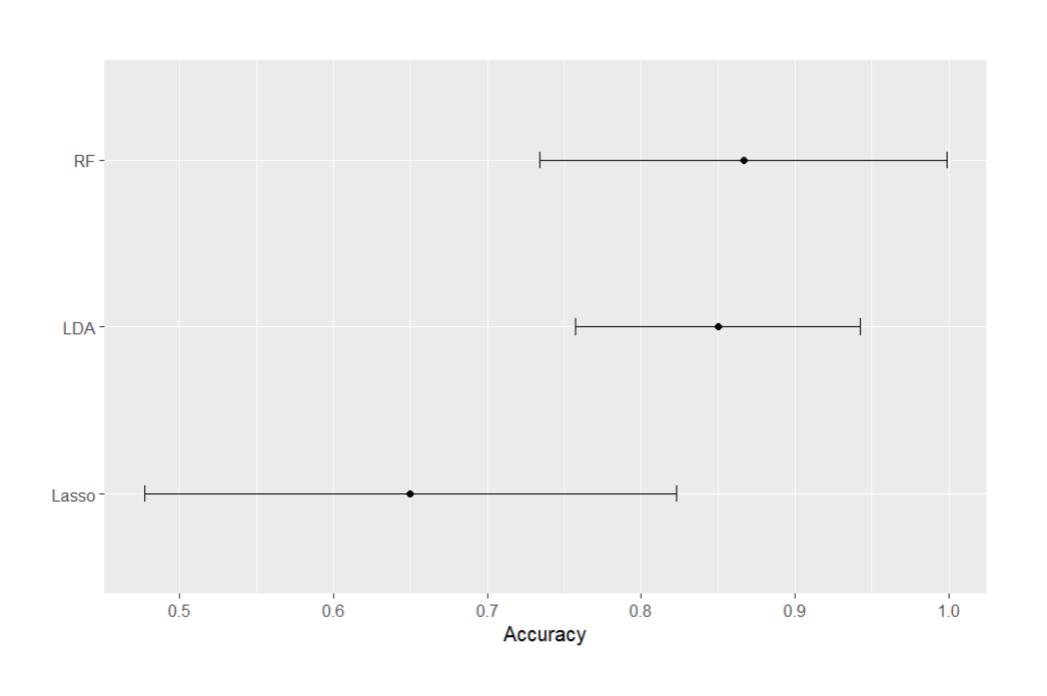
dist_matrix hclust (*, "complete") g1:24

g1: 23

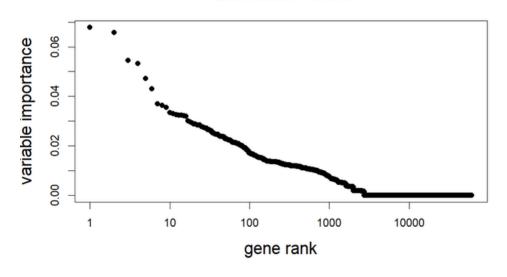
g2: 15

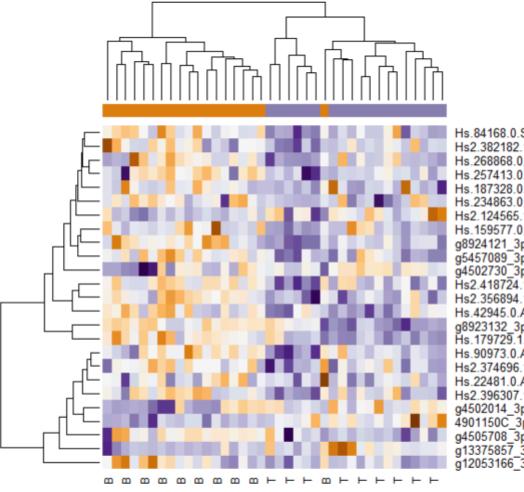
g2:14

SUPERVISED LEARNING METHODS



ALL subset results

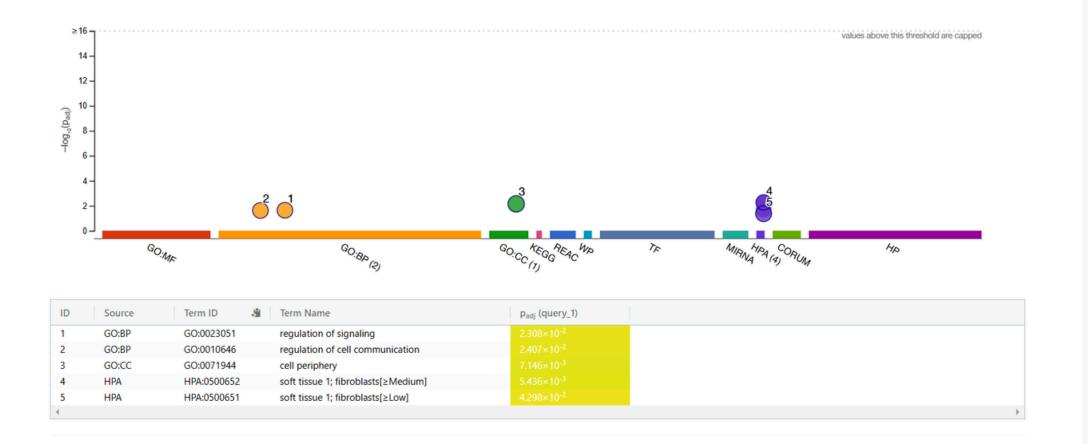




Hs.84168.0.S1_3p_at
Hs2.382182.1.S1_3p_at
Hs.268868.0.S1_3p_at
Hs.257413.0.A1_3p_at
Hs.257413.0.A1_3p_at
Hs.187328.0.A1_3p_at
Hs.234863.0.S1_3p_at
Hs2.124565.1.S1_3p_x_at
Hs2.124565.1.S1_3p_at
g8924121_3p_at
g5457089_3p_a_at
g5457089_3p_a_at
Hs2.418724.1.S1_3p_s_at
Hs2.356894.1.S1_3p_at
Hs2.356894.1.S1_3p_at
Hs.179729.1.S1_3p_a_at
Hs.179729.1.S1_3p_a_at
Hs.179729.1.S1_3p_a_at
Hs.2374696.1.S1_3p_at
Hs2.374696.1.S1_3p_at
Hs2.374696.1.S1_3p_at
Hs2.396307.1.S1_3p_at
g4502014_3p_at
4901150C_3p_at
g4505708_3p_at
g13375857_3p_a_at
g13375857_3p_a_at

FUNCTIONAL ENRICHMENT ANALYSIS

gProfiler

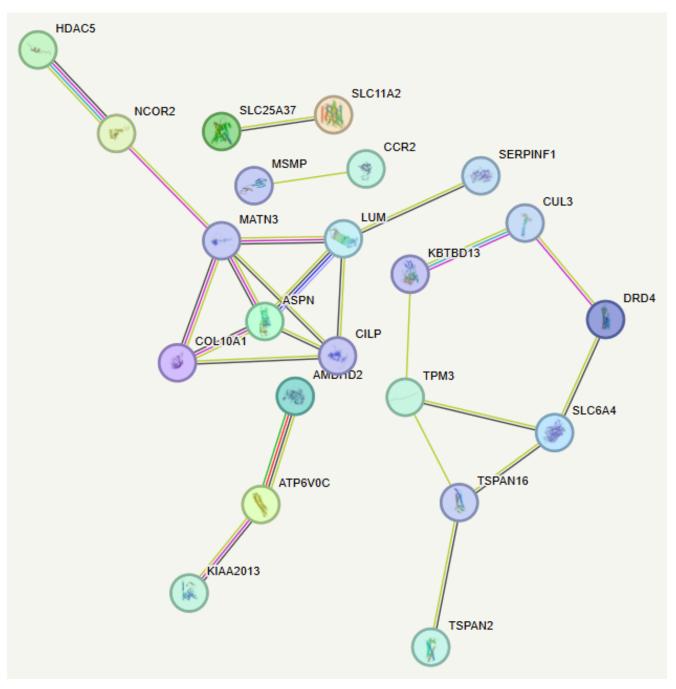


DAVID

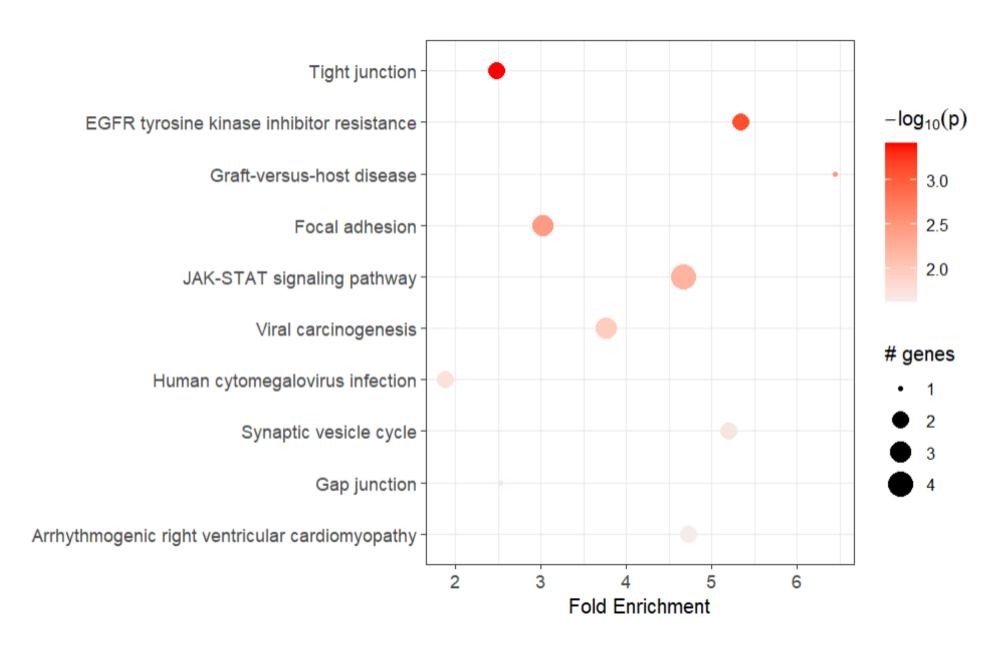
4 chart records						🖁 Download File		
Sublis	st <u>Category</u> \$	<u>Term</u>	RT	Genes	Count	<u>%</u>	P-Value	<u>Benjami</u>
	GOTERM_CC_DIRECT	membrane	RT		54	33,5	1,3E-4	3,5E-2
	GOTERM_CC_DIRECT	receptor complex	RT	=	7	4,3	3,9E-3	3,9E-1
	GOTERM_CC_DIRECT	endomembrane system	RT		6	3,7	4,5E-3	3,9E-1
	GOTERM_CC_DIRECT	nucleoplasm	RT		39	24,2	1,2E-2	7,5E-1
	GOTERM_BP_DIRECT	positive regulation of kinase activity	RT		4	2,5	5,8E-3	1,0E0
	GOTERM_BP_DIRECT	extracellular matrix organization	RT	=	6	3,7	6,0E-3	1,0E0
	WIKIPATHWAYS	Ferroptosis	RT		4	2,5	1,6E-2	1,0E0
	GOTERM_BP_DIRECT	microtubule cytoskeleton organization	RT		5	3,1	1,8E-2	1,0E0
	GOTERM_MF_DIRECT	protein binding	<u>RT</u>		98	60,9	2,2E-2	1,0E0
	WIKIPATHWAYS	Regulatory circuits of STAT3 signaling	RT	=	4	2,5	2,5E-2	1,0E0
	GOTERM_CC_DIRECT	synaptic membrane	RT		3	1,9	3,1E-2	1,0E0
	GOTERM_MF_DIRECT	protein tyrosine kinase activity	RT	<u>=</u>	4	2,5	3,3E-2	1,0E0
	GOTERM_CC_DIRECT	cell junction	<u>RT</u>		5	3,1	3,6E-2	1,0E0
	REACTOME_PATHWAY	MAPK family signaling cascades	RT	=	7	4,3	3,6E-2	1,0E0
	GOTERM_BP_DIRECT	cell surface receptor protein tyrosine kinase signaling pathway	RT		4	2,5	3,8E-2	1,0E0
	GOTERM_BP_DIRECT	cadmium ion transmembrane transport	RT	•	2	1,2	3,9E-2	1,0E0
	GOTERM_MF_DIRECT	ferrous iron transmembrane transporter activity	<u>RT</u>		2	1,2	4,0E-2	1,0E0
	GOTERM_CC_DIRECT	endosome membrane	RT	=	6	3,7	4,2E-2	1,0E0
	GOTERM_MF_DIRECT	extracellular matrix structural constituent	<u>RT</u>		4	2,5	4,5E-2	1,0E0
	GOTERM_BP_DIRECT	cellular defense response	RT	Ē	3	1,9	5,0E-2	1,0E0
	WIKIPATHWAYS	Microtubule cytoskeleton regulation	<u>RT</u>		3	1,9	5,2E-2	1,0E0
	GOTERM_CC_DIRECT	<u>plasma membrane</u>	RT		47	29,2	5,2E-2	1,0E0
	GOTERM_BP_DIRECT	cobalt ion transport	RT		2	1,2	5,2E-2	1,0E0
	GOTERM_MF_DIRECT	iron ion transmembrane transporter activity	RT	1	2	1,2	5,3E-2	1,0E0
	GOTERM_CC_DIRECT	collagen-containing extracellular matrix	RT		7	4,3	5,4E-2	1,0E0
	GOTERM_BP_DIRECT	positive regulation of protein localization to plasma membrane	RT	•	3	1,9	5,5E-2	1,0E0
	REACTOME_PATHWAY	Transcriptional Regulation by TP53	RT		7	4,3	5,6E-2	1,0E0
	GOTERM_BP_DIRECT		RT	=	2	1,2	5,8E-2	1,0E0
	GOTERM_BP_DIRECT		RT		2	1,2	5,8E-2	1,0E0
	GOTERM_BP_DIRECT		RT		9	5,6	6,0E-2	1,0E0
	EC_NUMBER	2.7.10.1	RT	_	3	1,9	6,4E-2	1,0E0
		unmethylated CpG binding	RT	_	2	1,2	6,5E-2	1,0E0
	WIKIPATHWAYS	Wnt signaling	RT		4	2,5	6,5E-2	1,0E0
	GOTERM_MF_DIRECT		RT	_	3	1,9	6,9E-2	
	REACTOME_PATHWAY		RT		27	16,8	6,9E-2	1,0E0
	GOTERM_CC_DIRECT	<u>endosome</u>	RT		6	3,7	7,0E-2	1,0E0
	GOTERM_MF_DIRECT		RT		2	1,2	7,2E-2	1,0E0
		Interleukin-21 signaling	RT	=	2	1,2	7,3E-2	
		peroxisomal membrane	RT		3		7,4E-2	
0	KEGG_PATHWAY	Regulation of actin cytoskeleton	RT	=	5	3,1	7,7E-2	
	KEGG_PATHWAY	Cytoskeleton in muscle cells	RT		5		7,9E-2	
	REACTOME_PATHWAY		RT		20		8,5E-2	
		double-stranded RNA binding	RT		3	1,9	8,9E-2	
	GOTERM_BP_DIRECT	positive regulation of calcium ion transport into cytosol	RT		2	1,2	9,5E-2	1,0E0

FUNCTIONAL ENRICHMENT ANALYSIS

STRING



pathfindR



CONCLUSION AND FUTURE PERSPECTIVE

PCA is able to divide the two groups

RF has the best performance

In the enrichment analysis, not many enriched terms were **specifically connected** to breast cancer or brain metastasis.

Repeat the analysis considering a larger number of samples

Repeat the analysis considering **BRCA1** gene

Repeat the analysis focusing on the genes highlighted by **rScudo**



RSCUDO

