

# miRNA pathway views for Alzheimer's Disease

# Description of miRNopath algorithm

- The algorithm attempts to identify pathways where the genes predicted to be affected by miRNA changes are present much more so than one would expect by random chance alone.
- The slides below detail the observations seen for some pathways of interest
- The counts represent the number of miRNAs and genes affected in a pathway, with emphasis on the consistent directionality of the changes.
  - In large part, most pathways predicted to be significantly affected by the miRNA changes are affected by miRNAs which are all consistently up-regulated, or all consistently down-regulated.
  - In some cases, there are some miRNA changes in an opposing direction from the majority, which either reflects that pathways are often defined with some positive- and negative-effectors, or that the miRNA changes are not in complete agreement. In these cases, the section “Both” is used to show any differences from “Up” or “Down.”
- Note the library size, which is defined as the total set of miRNAs measured and detected, which therefore varies for each tissue and Braaks stage.
  - The library size is used to define the random background, a uniquely challenging estimate based upon the overall set of genes predicted to be affected by the “measured and detected” miRNAs.
  - From this background, the “measured, detected, and statistically-significantly-differentially expressed miRNAs” are compared, using only their set of genes predicted to be affected.

# Understanding the miRNAPath views

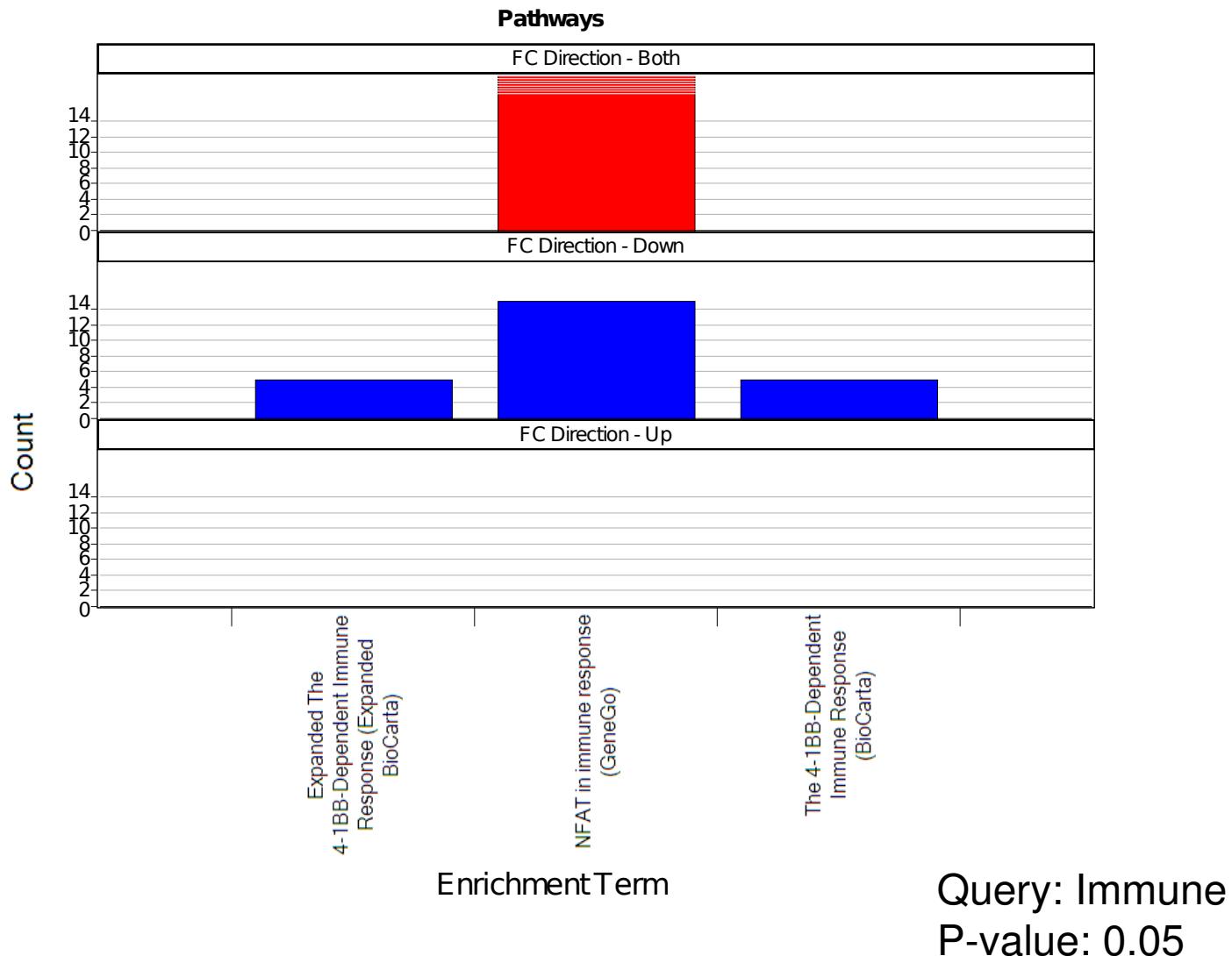
- The pathway views show the direction of the miRNAs regardless of the view (first is pathways, second is miRNAs, and third is targets)
- In the pathway and gene views, assume the genes and pathways are being modulated opposite to the direction of the miRNAs

# Color coding for Spotfire files

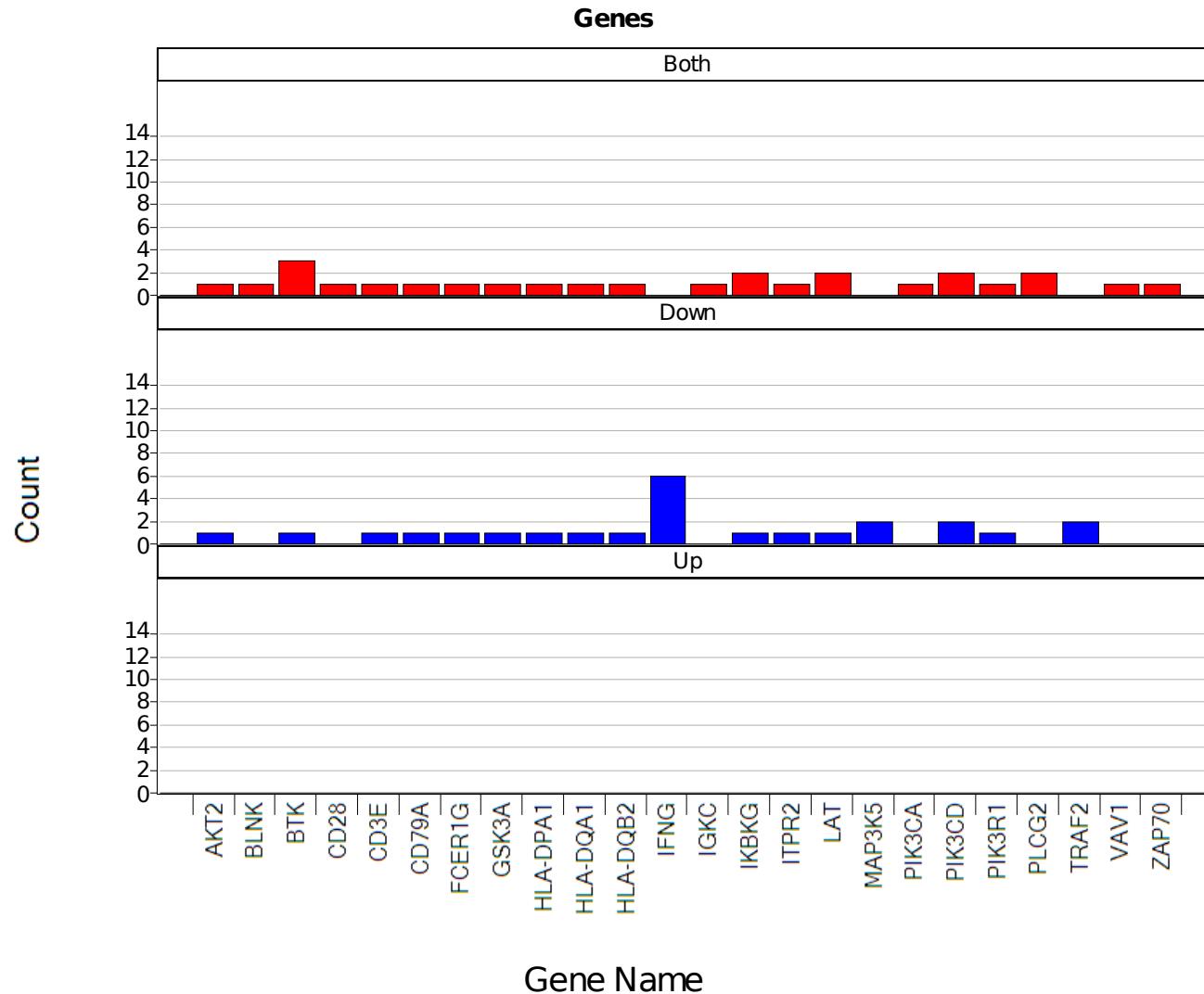
	B3,4	B5,6
Hippocampus		
Medial Frontal Gyrus		
Cerebellum		

	Up	Down
CSF		

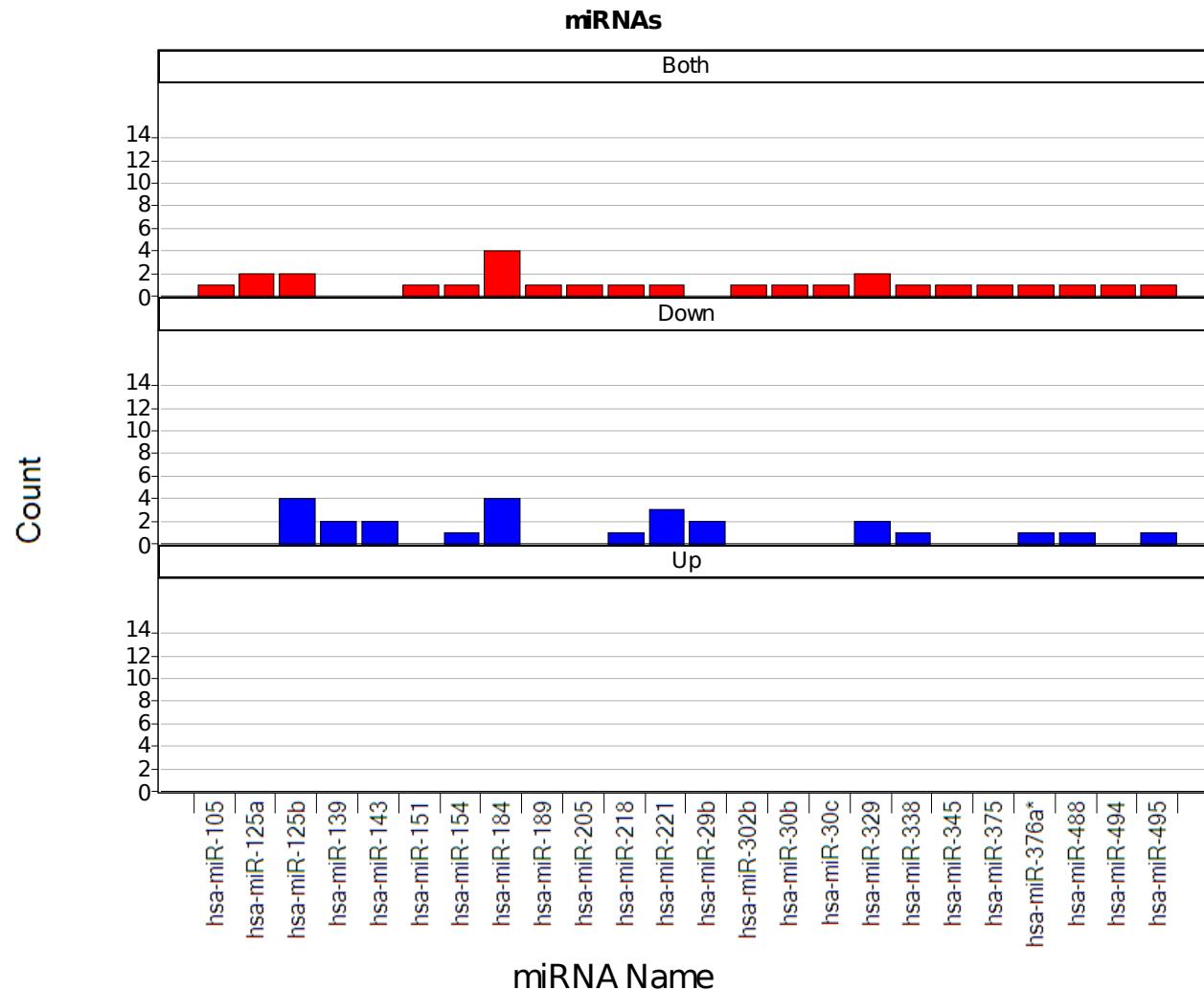
# AD CSF: miRNAs in T cell signaling pathways



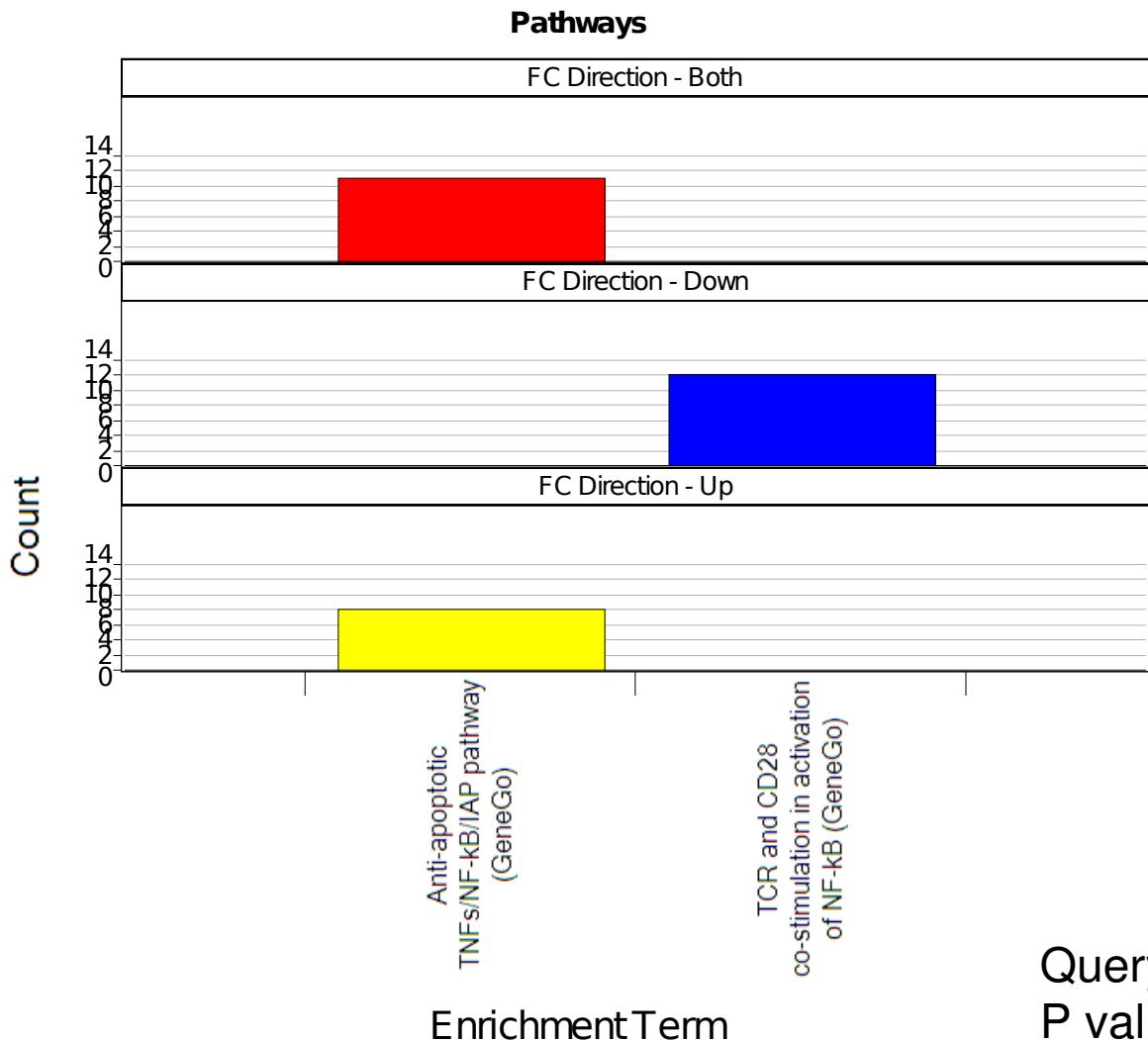
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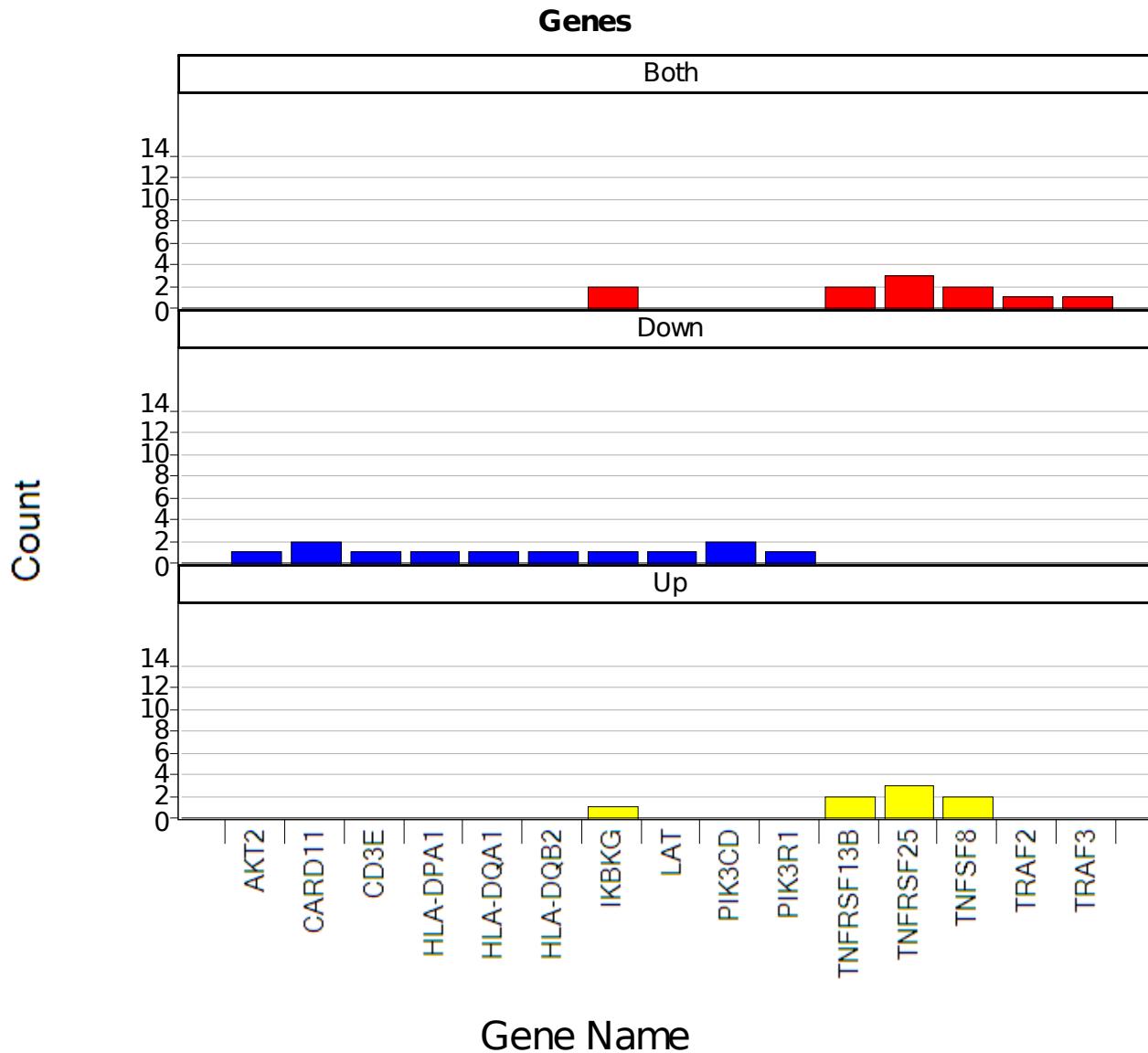


# AD CSF: miRNAs in NF-kB pathway

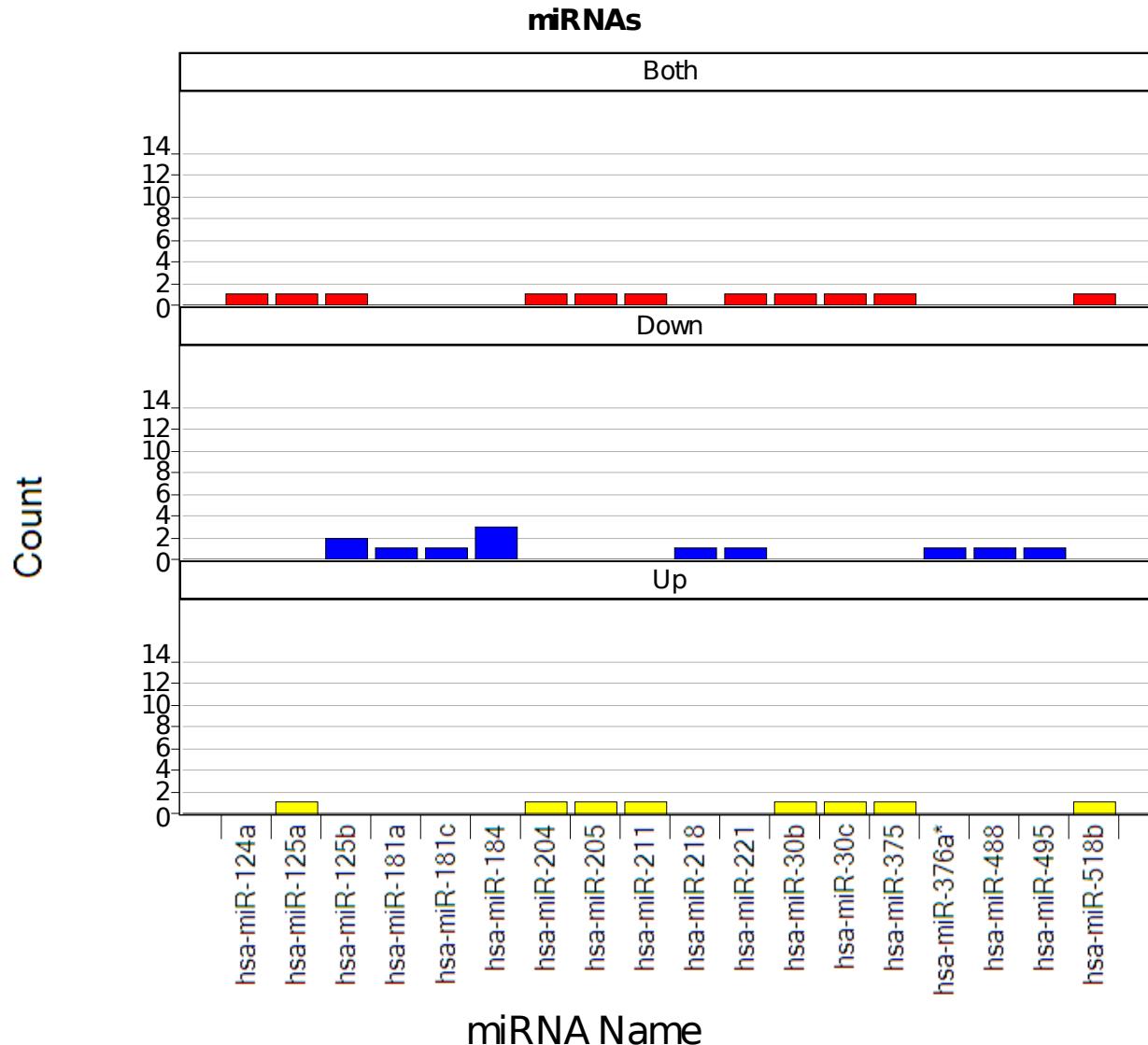


Query: NF-kB  
P value: 0.05

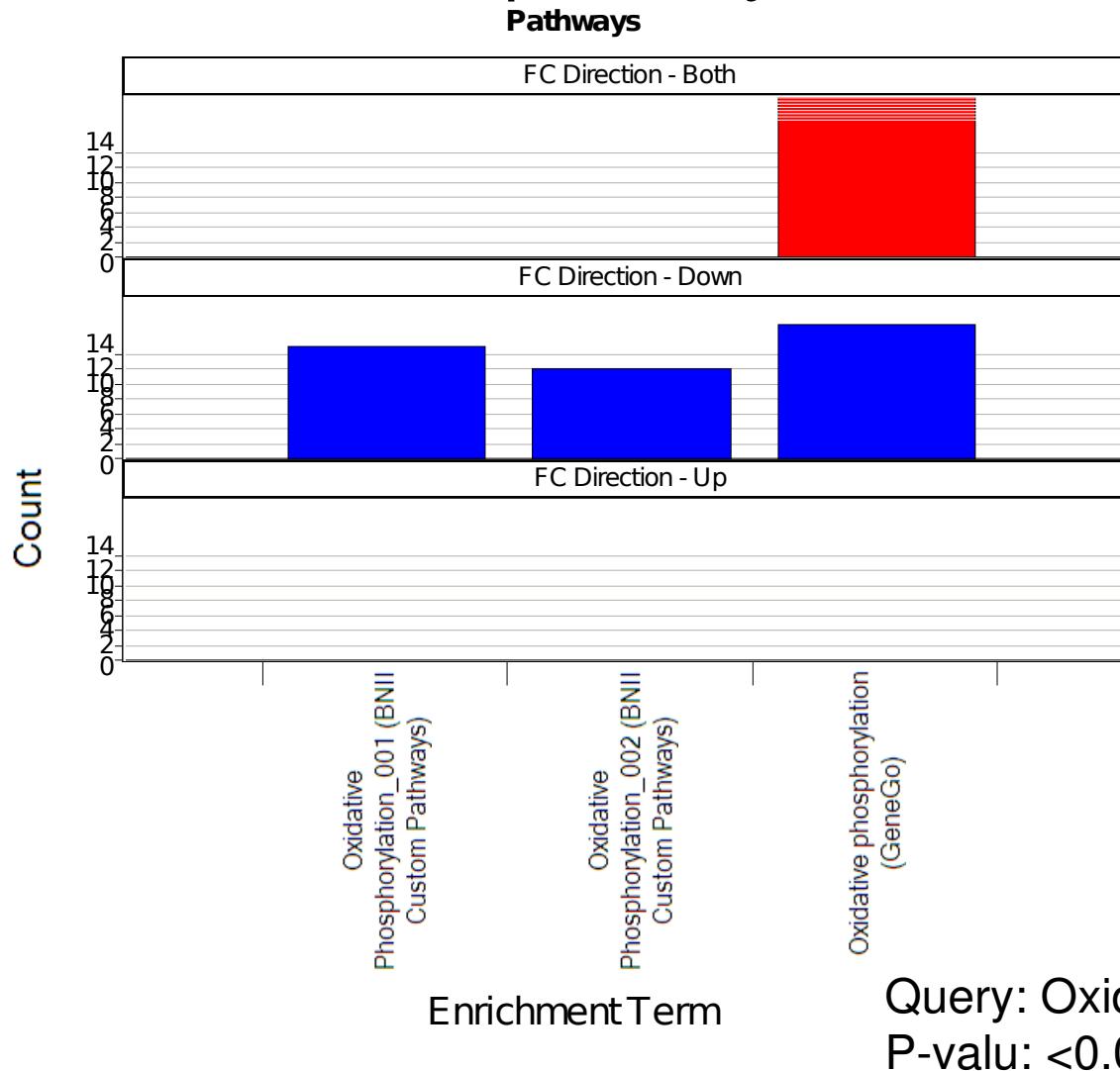
# AD CSF: miRNAs in NF- $\kappa$ B pathway



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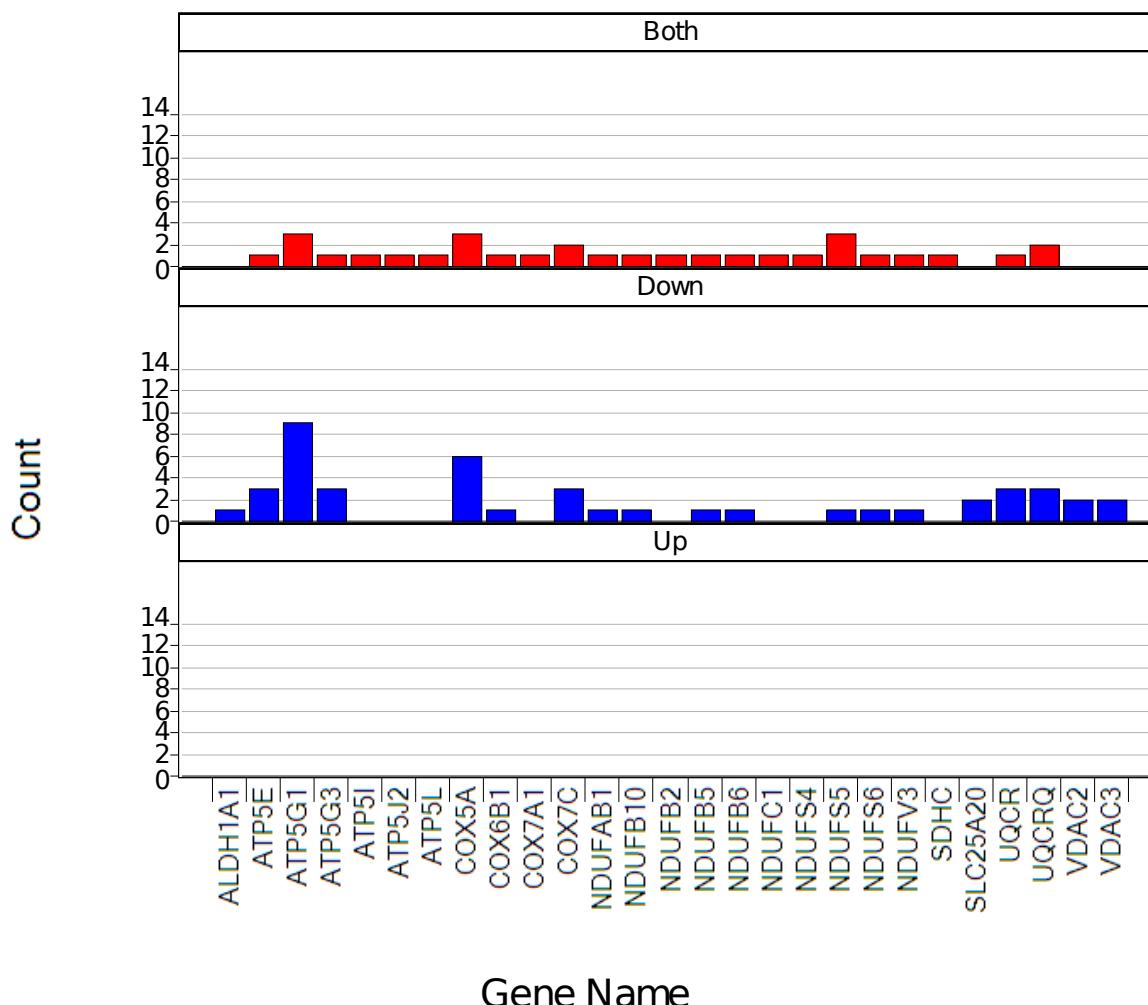


# AD CSF: miRNAs in Oxidative phosphorylation pathways

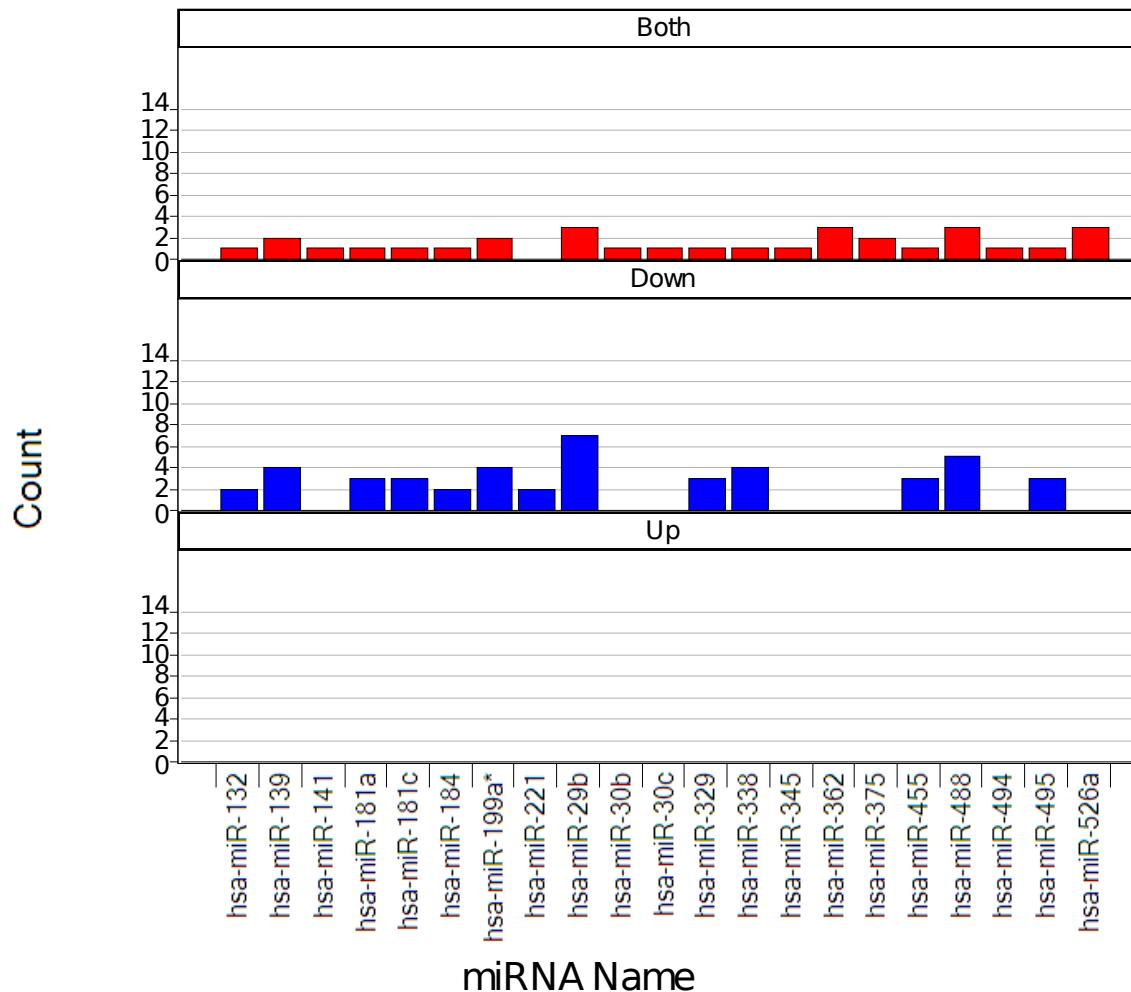


Query: Oxidative Phosphorylation  
P-val: <0.05

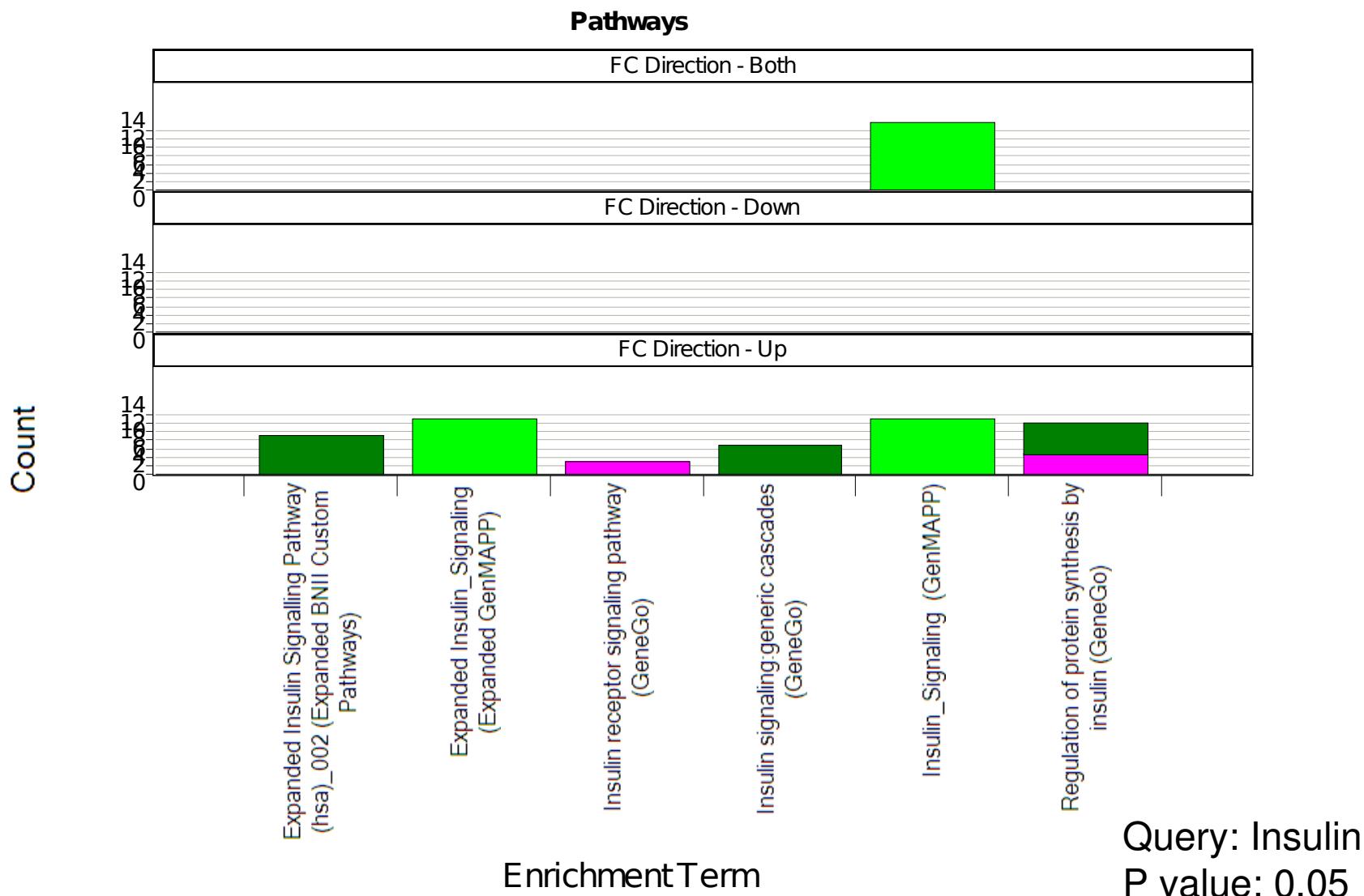
# AD CSF: miRNAs in Oxidative phosphorylation pathways



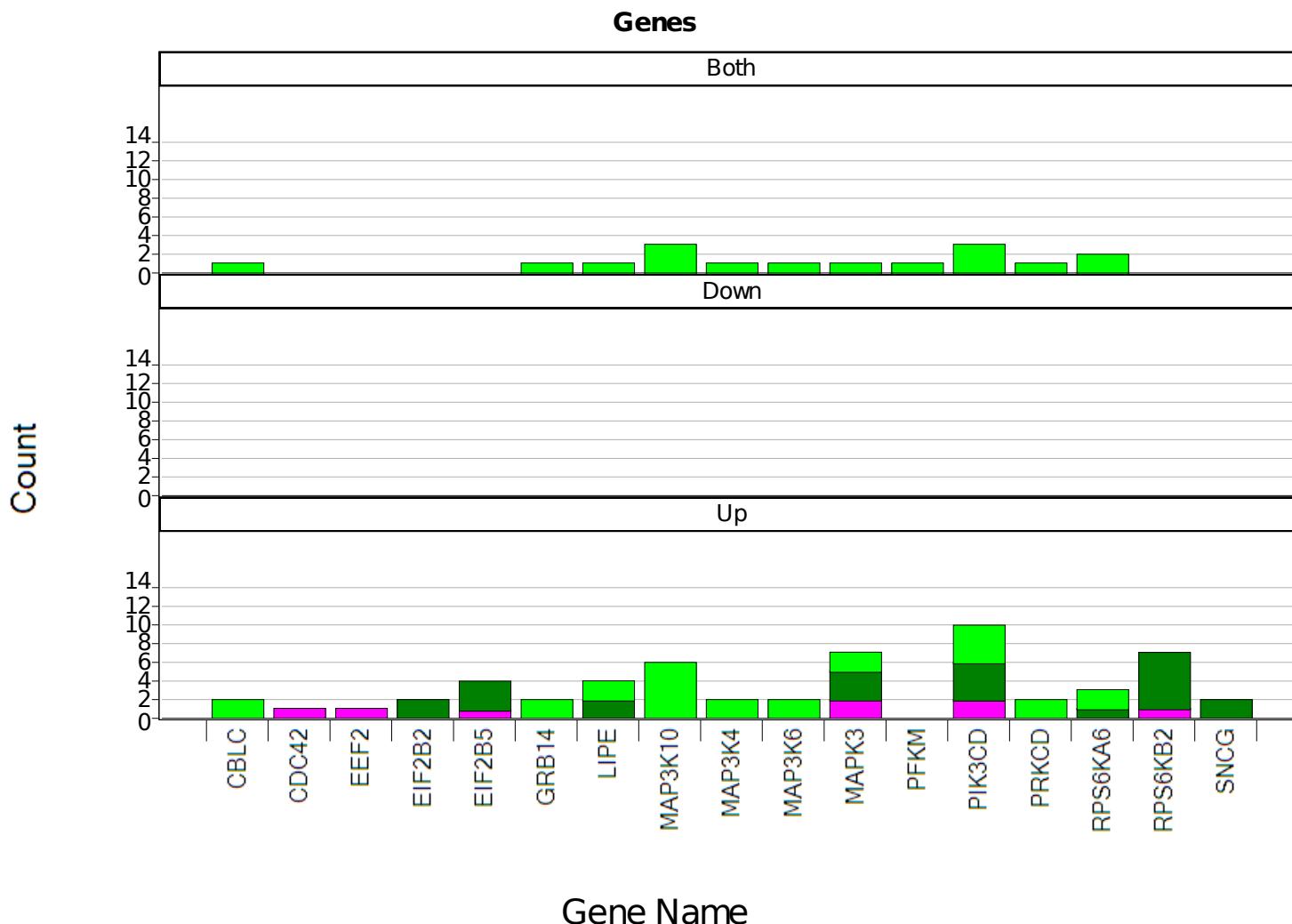
# AD CSF: miRNAs in Oxidative phosphorylation pathways



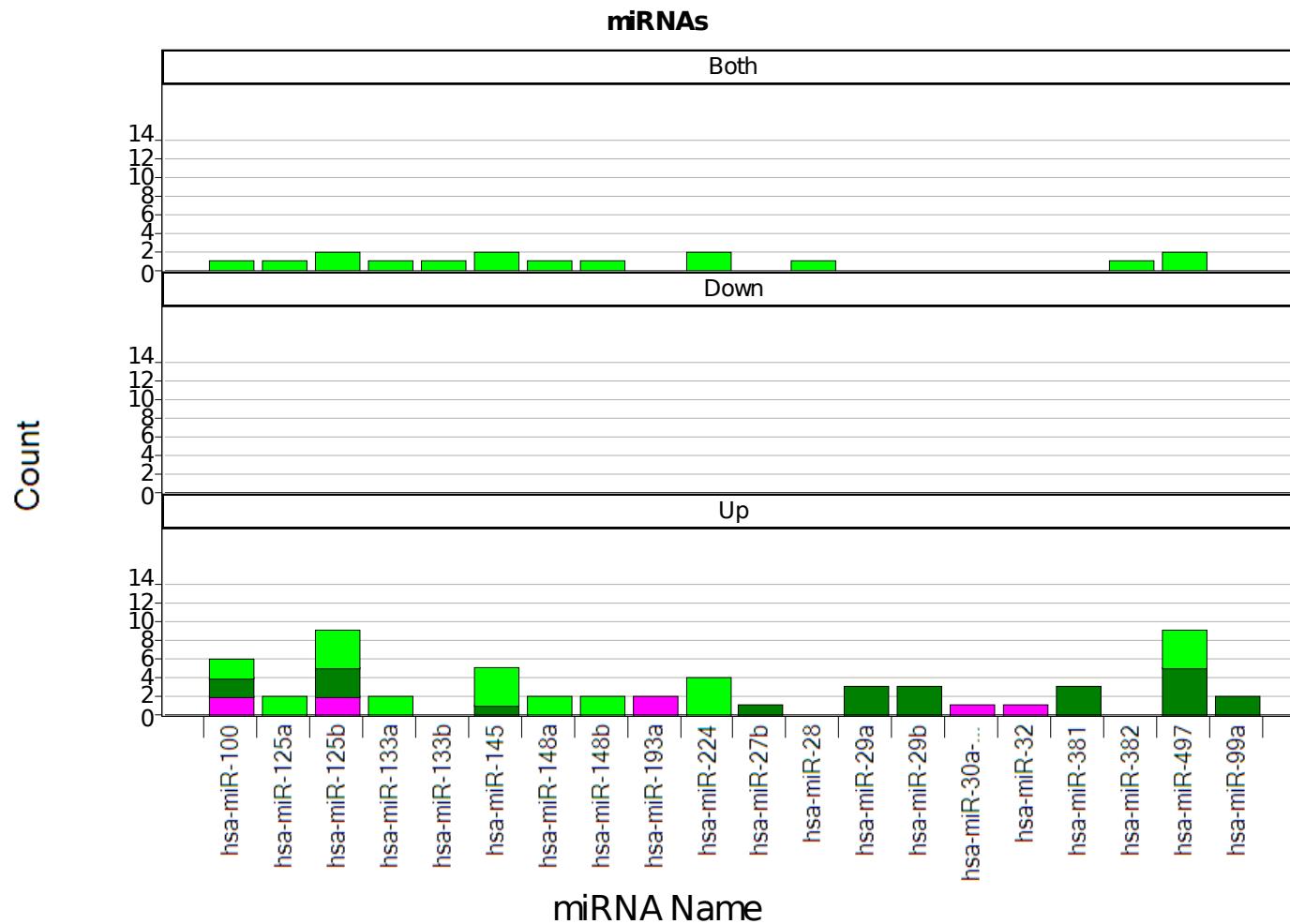
# AD Brain: miRNAs in Insulin Signaling Pathways



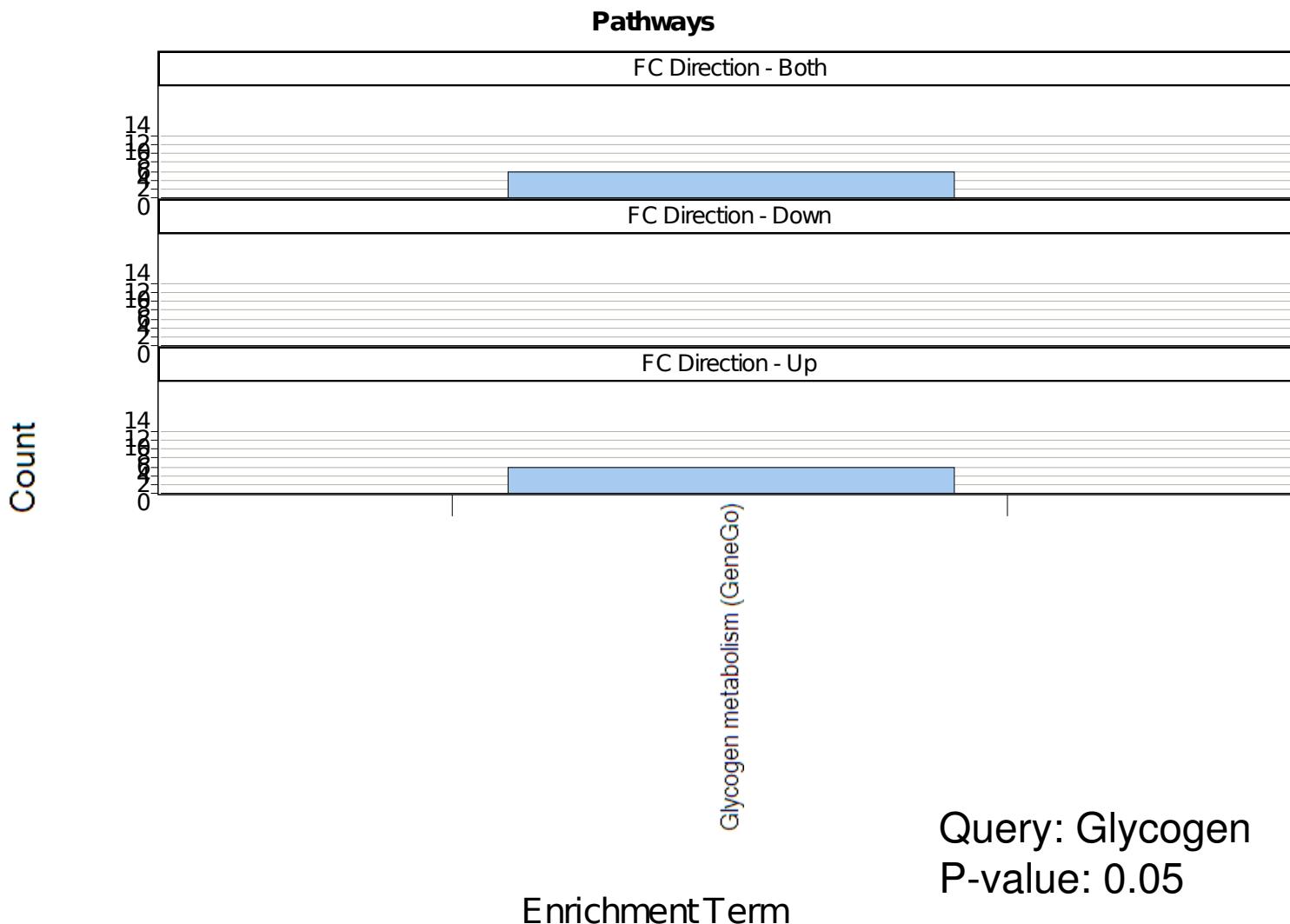
# AD Brain: miRNAs in Insulin Signaling Pathways



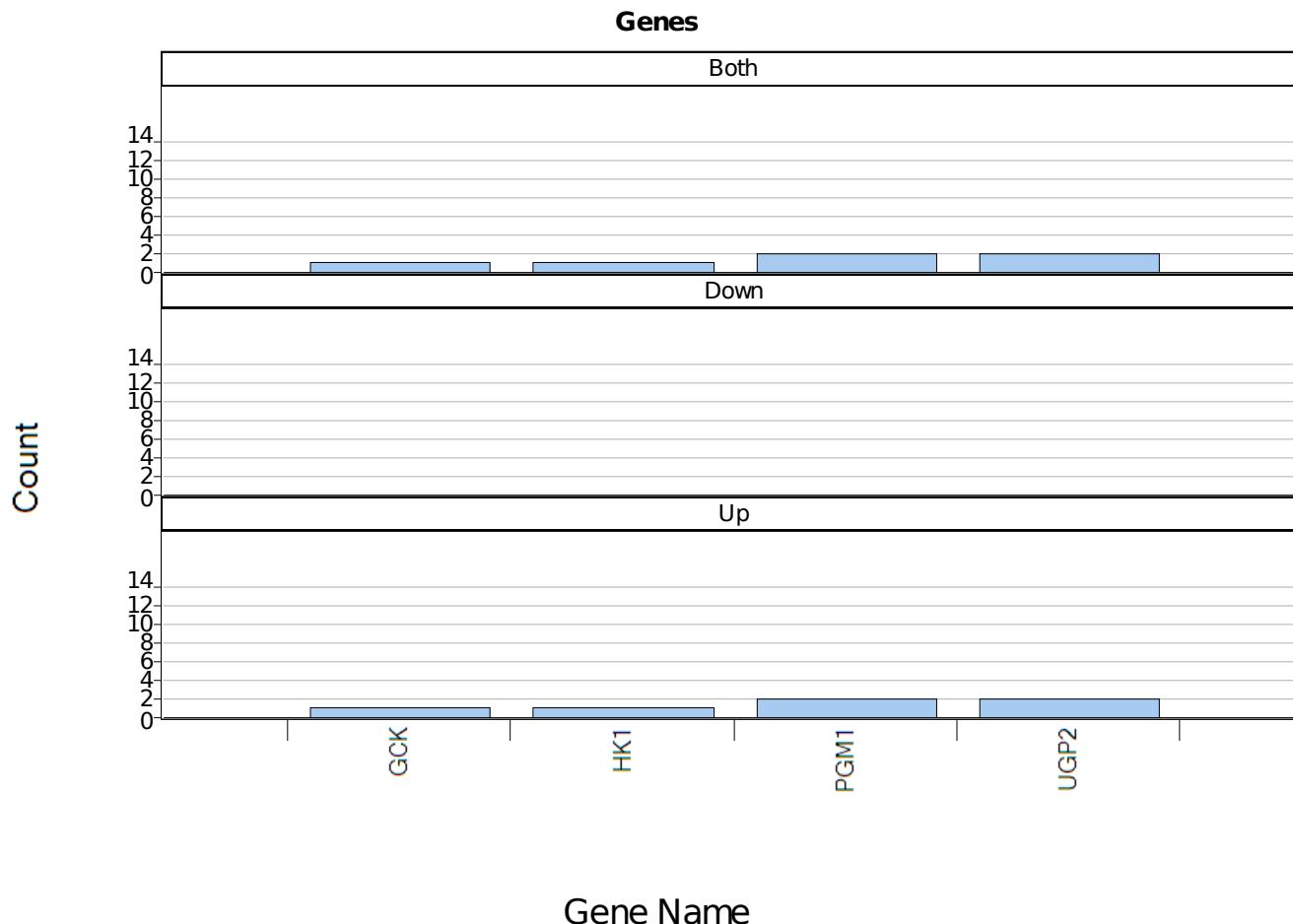
# AD Brain: miRNAs in Insulin Signaling Pathways



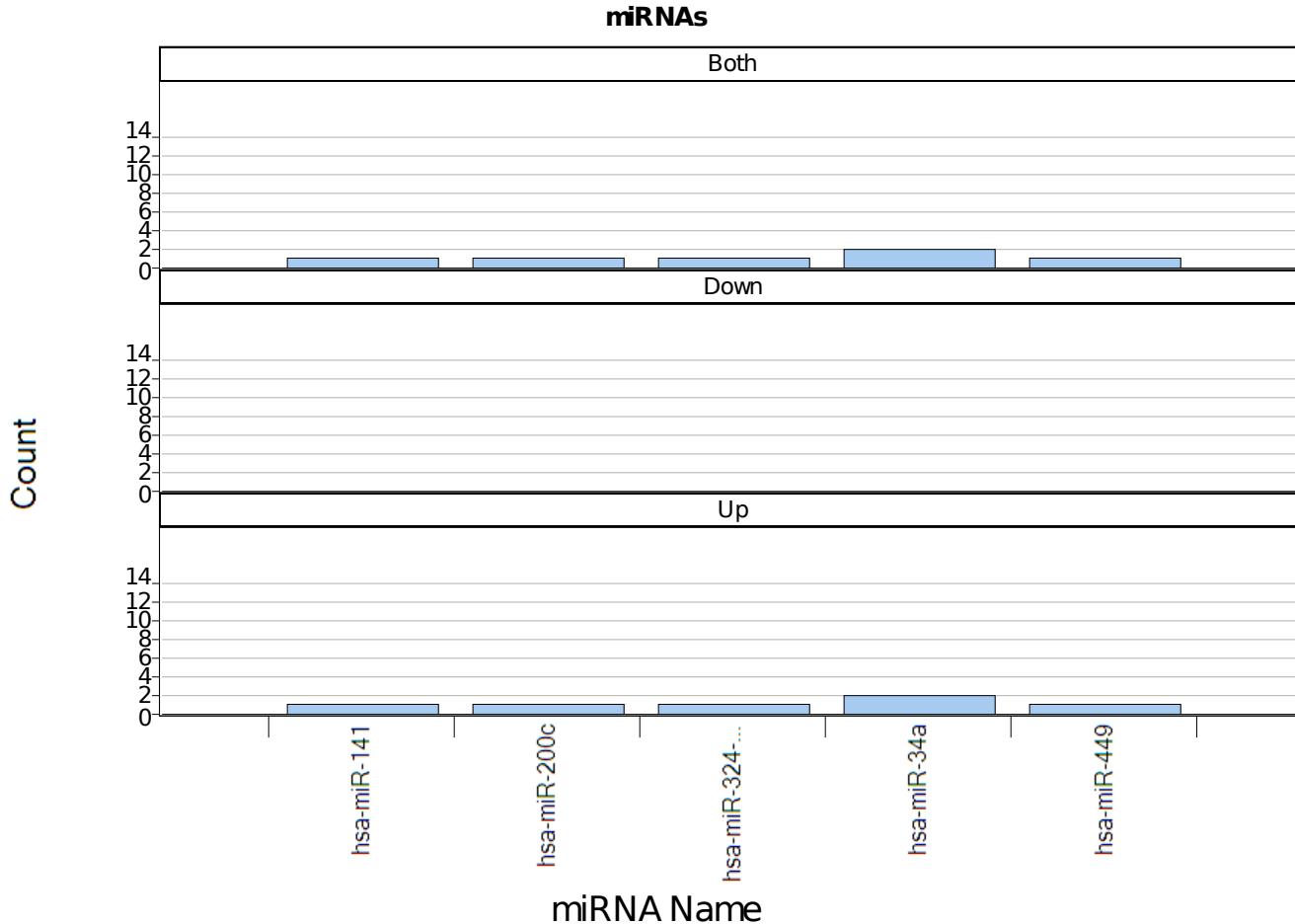
# AD Brain: miRNAs in Glycogen Metabolism



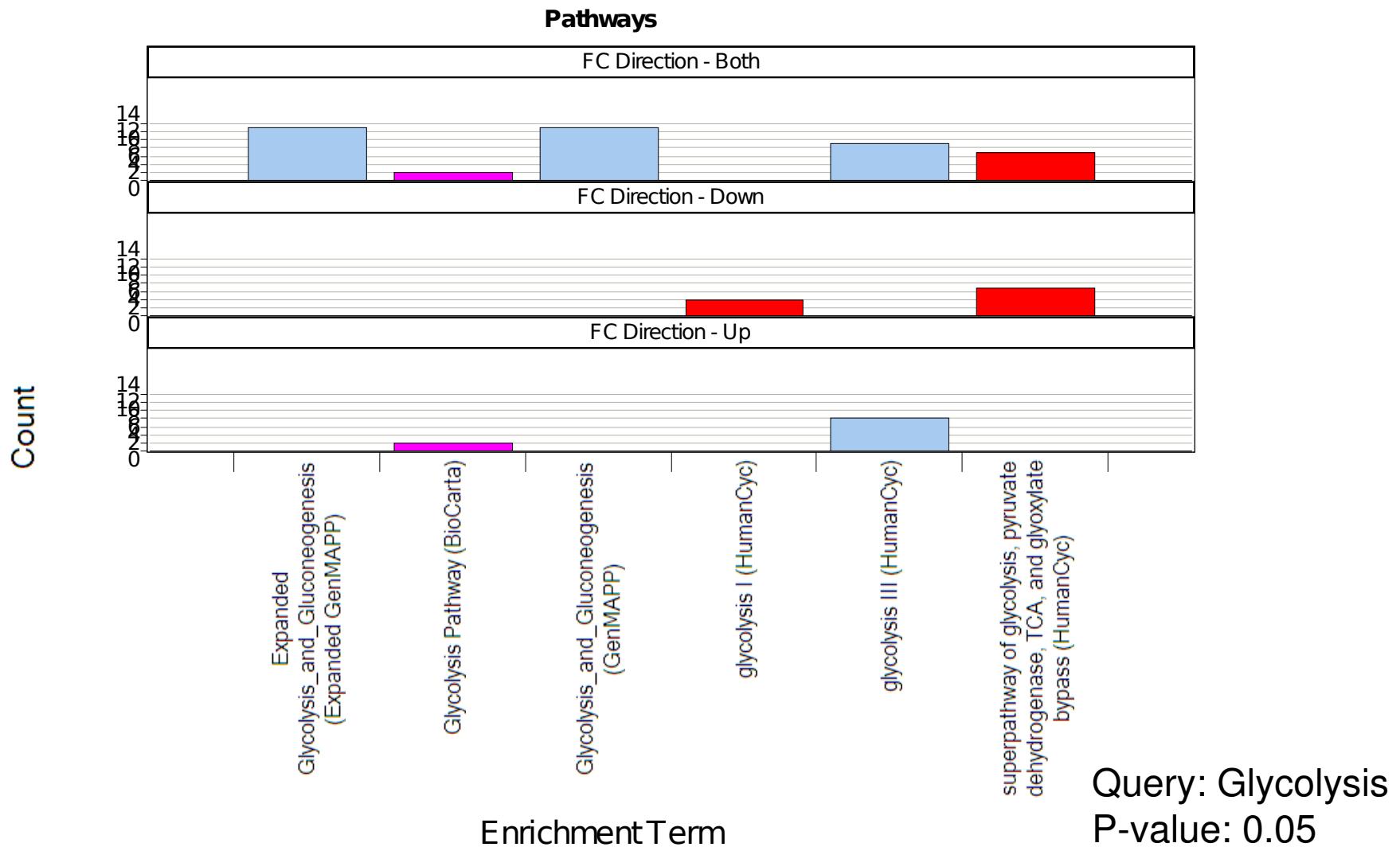
# AD Brain: miRNAs in Glycogen Metabolism



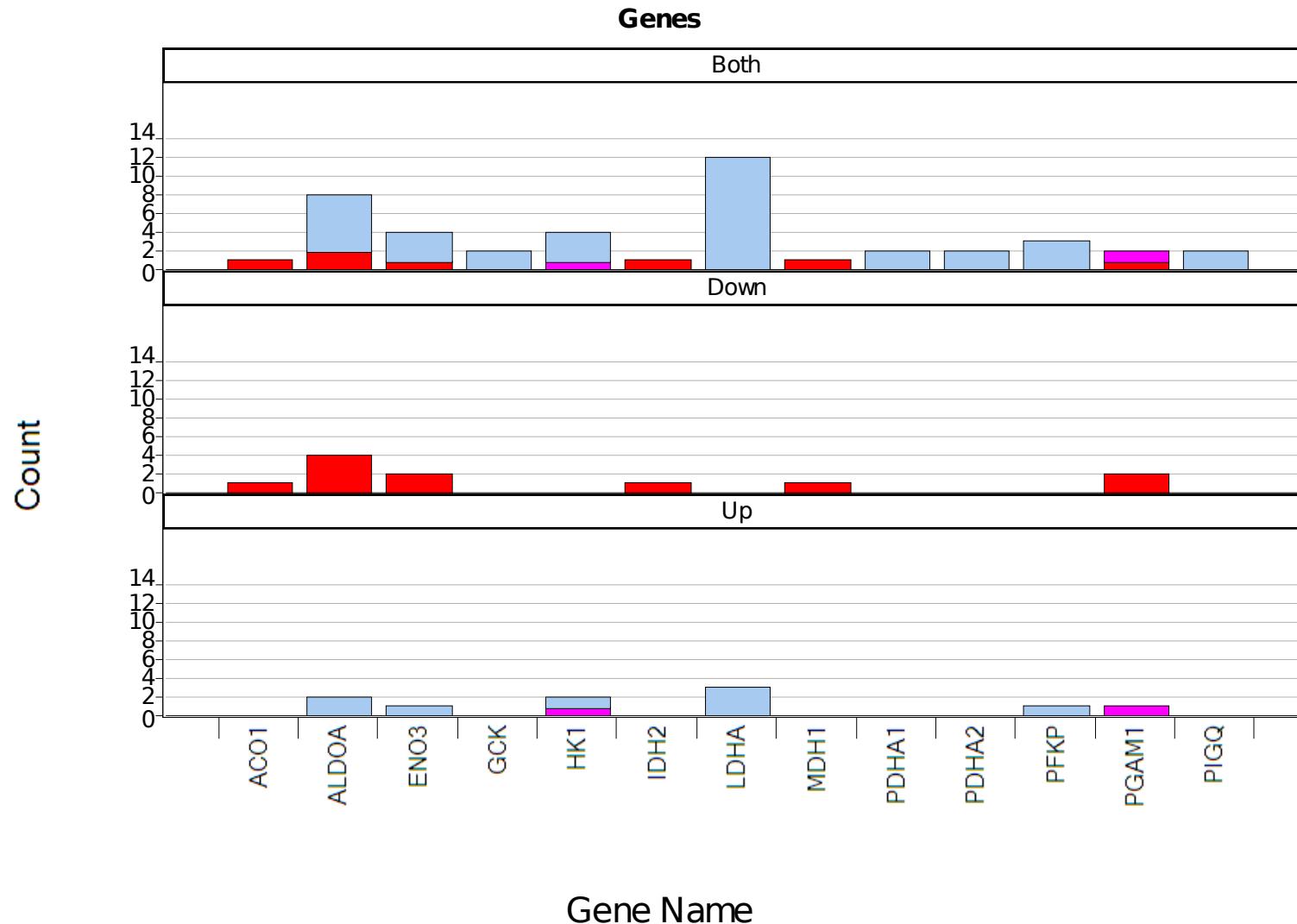
# AD Brain: miRNAs in Glycogen Metabolism



# AD Brain: miRNAs in Glycolysis Pathways



# AD Brain: miRNAs in Glycolysis Pathways



# AD Brain: miRNAs in Glycolysis Pathways

