Roaming Around AML

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Abstract

Some random words placeholding the abstract

1. Introduction

This article conducts an unplanned investigation of bunch of data and research results about AML decease. As indicated by ?, the fifth most affected site in body by cancer is the circulatory system, with about a hundred thousands estimated cases in total of a million cancer in the US, 2002. AML is one of the most suffered and fatal one with almost 35% of the cases resulting into death. Hence, piling up some results in this area will accelerate more experienced scientists toward a working solution. However, this field is not abandoned, and has been worked on extensively. Therefore, they are considered as rich sources of result validation. The process of the investigation begins with a beginner level differential expression analysis resulting in some gene lists with significant gene expression alterations. These lists are used as the input of the next pipeline component, gene enrichment. Extracting some pieces of information in this step will be the basis of some fact finding endeavors.

2. Differential Expression Analysis

This article has employed? data set and Limma tools?. The data set primarily consists of 170 samples having their expressions measured by 32321 probes and classified in two categories, having self-explanatory labels, "Normal" and "Leukemia." These samples were first examined in three prospectives, respectively independent of the cell source, cells of type T, cells of type B(Table 1.)

After having the data in this format, a basic visualization was employed to form an intuition about the comparability of the data. The data needed no normalization (figure 1.) However, the heatmap and its interensic hierarchical clustering failed to assure the validity of the data.

extract some genes with significantly different expression levels in normal versus affected cells.

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Table 1: The samples were studied dependant and independant of their source.

Source Dependancy	Normal Sources	Leukemia Sources
Independent	B Cells, CD34+HSPC, Granulocytes, Monocytes, T Cells	AML Cell Line, AML Patient, B ALL Cell Line, B ALL Patient, T ALL Cell Line, T ALL Patient
T cells	T Cells	T ALL Cell Line, T ALL Patient
B cells	B Cells	B ALL Cell Line, B ALL Patient

3. Gene Set Enrichment Analysis

As the gene sets extracted, they were passed to EnrichR to find possible causes and cures for AML. The GSEA results were truncated not to be represent ideas elusive to naked eyes. So only results with adjusted p-value less than .05 were selected. Moreover, at most 5 top cases extracted from each database were kept and the rest were eliminated. The results were sorted by combined score before the elimination process. The databases that were exploited for each purpose are listed in Table B.48 . Appendix A would explain the reason of each database choice.

Table 2: Databases in	n Use	for	GSEA
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Analysis	Databases
Transcription Factors	TRANSFAC and JASPAR PWMs, miRTarBase 2017
Pathways	KEGG2019 Human, NCI-Nature 2016, Reactome 2016
Gene Ontology	GO Biological Process 2018, GO Molecular Function 2018, GO
	Cellular Component 2018
Drugs	ARCHS4 IDG Coexp

The comprehensive genes outputed by GSEA can be viewed in Appendix B. Amongth the extracted genes, pathways, etc, some were more attractive and listed here with some details. The headers are represented in the format of ([Source_Change], Name, Adjusted P-value, Z-score, Combined Score, database.)

3.1. Transcription Factors (TF)

This section has been widely influenced by proteinatlas and KEGG.

3.1.1. ETV4

Transcription regulator, e.g is unfavorable to liver cancer while favorable to thyroid cancer.

3.1.2. E2F4

Transcription facator, significant role in cell cycle regulation.

3.1.3. E2F1

Transcription factor of the E2F TF family.

3.1.4. NRF1

Nuclear respiratory factor 1, TF regulating key metabolic genes.

3.1.5. NFIC

TF in cells and replication factor for adenovirus.

3.1.6. NFYA

TF associated with a subunit of a high specified and affined DNA binding complex.

3.1.7. POU2F1

Of the same group of oct4 i.e octamer transcription factors.

3.2. Pathways (PW)

3.2.1. Glycosaminoglycan degradation

Glycosaminoglycans are polar molecules found in extracellular matrix and interact with growth factors

3.2.2. Glycosphingolipid biosynthesis

Glycosphingolipids form a major part of lipid rafts in cell membrane, in which some receptors lay

3.2.3. RIG-I-like receptor signaling pathway

RIG-I-like receptors have accepted the role of pathogen sensing of RNA viruses

3.2.4. Autoimmune thyroid disease

A disease in which thyroid cells are defected and secrete antigens consequently, attracting B and T cells toward themselves, therefore, driven to necrosis/apoptosis.

3.2.5. Graft-versus-host disease

Allogeneic hematopoietic transplantations causes some disorders invoking donor cells against the host ones.

- 3.2.6. Endosomal/Vacuolar pathway
- 3.2.7. Interferon alpha/beta signaling_Homo sapiens_R-HSA-909733
- 3.2.8. Antigen Presentation: Folding, assembly and peptide loading of class I MHC_Homo sapiens_R-HSA-983170
- 3.2.9. Immune System_Homo sapiens_R-HSA-168256
- 3.2.10. Interferon gamma signaling_Homo sapiens_R-HSA-877300
- 3.2.11. DNA replication
- 3.2.12. Homologous recombination
- 3.2.13. Oocyte meiosis
- 3.2.14. Fanconi anemia pathway
- 3.2.15. Small cell lung cancer
- 3.2.16. E2F transcription factor network_Homo sapiens_bb4d0fd3-6191-11e5-8ac5-06603eb7f303
- 3.2.17. PLK1 signaling events_Homo sapiens_e5e87977-6194-11e5-8ac5-06603eb7f303
- 3.2.18. FOXM1 transcription factor network_Homo sapiens_c51cda49-6192-11e5-8ac5-06603eb7f303
- 3.2.19. Aurora B signaling_Homo sapiens_304a75af-618c-11e5-8ac5-06603eb7f303
- 3.2.20. ATR signaling pathway_Homo sapiens_8991cbac-618b-11e5-8ac5-06603eb7f303
- 3.2.21. Cell Cycle_Homo sapiens_R-HSA-1640170
- 3.2.22. Cell Cycle, Mitotic_Homo sapiens_R-HSA-69278
- 3.2.23. M Phase_Homo sapiens_R-HSA-68886
- 3.2.24. Cell Cycle Checkpoints_Homo sapiens_R-HSA-69620
- 3.2.25. G2/M Checkpoints Homo sapiens R-HSA-69481
- 3.2.26. Th17 cell differentiation
- 3.2.27. Allograft rejection
- $3.2.28.\ TNF\ receptor\ signaling\ pathway_Homo\ sapiens_316be05e-6196-11e5-8ac5-06603eb7f303$
- 3.2.29. TRAIL signaling pathway_Homo sapiens_3a79fddf-6196-11e5-8ac5-06603eb7f303
- 3.2.30. IL12-mediated signaling events_Homo sapiens_7acdea19-6193-11e5-8ac5-06603eb7f303
- 3.2.31. Downstream signaling in naive CD8+ T cells_Homo sapiens_92180cef-6191-11e5-8ac5-06603eb7f303
- 3.2.32. IL2-mediated signaling events_Homo sapiens_a2a1883c-6193-11e5-8ac5-06603eb7f303
- 3.2.33. Cytokine Signaling in Immune system_Homo sapiens_R-HSA-1280215
- 3.2.34. Glyoxylate and dicarboxylate metabolism
- 3.2.35. Glycosaminoglycan biosynthesis
- $3.2.36.~p73~transcription~factor~network_Homo~sapiens_a88c505e-6194-11e5-8ac5-06603eb7f303$
- 3.2.37. Mitotic Prometaphase_Homo sapiens_R-HSA-68877
- 3.2.38. Resolution of Sister Chromatid Cohesion Homo sapiens R-HSA-2500257
- 3.2.39. Intestinal immune network for IgA production

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- 3.2.40. TNF signaling pathway
- 3.2.41. Asthma
- 3.2.42. Immunoregulatory interactions between a Lymphoid and a non-Lymphoid cell_Homo sapiens_R-HSA-198933
- 3.2.43. Adaptive Immune System_Homo sapiens_R-HSA-1280218
- 3.2.44. CD22 mediated BCR regulation Homo sapiens R-HSA-5690714

Appendix A. GSEA Dataset Selection

place holder

Appendix B. GSEA Full Selected Genes

Table B.3: Databases in Use for GSEA

Name	Adj. value	P-	Z- score	Combined Score
ETV4 (human)	0.043		-1.74	15.24

Table B.4: Databases in Use for GSEA

Name	Adj. P- value	Z- score	Combined Score
E2F4 (human)	6.248e-9	-1.97	48.40
E2F1 (human)	0.00002605	-1.55	23.43
NRF1 (human) NFIC (human)	0.0001446 0.001465	-1.71 -1.69	22.44 18.00
NFYA (human)	0.002248	-1.60	16.03

Table B.5: Databases in Use for GSEA

Name	Adj. P	- Z-	Combined
	value	score	Score
hsa-miR-193b-3p	8.760050e-	-	973.9777
	64	6.3641	76
hsa-miR-192-5p	2.501043e-	-	522.3863
	31	6.7657	65
hsa-miR-215-5p	2.718386e-	-	437.7653
_	34	5.1843	328
hsa-miR-34a-5p	9.159129e-	-	269.1568
•	21	5.1171	.80
hsa-miR-92a-3p	1.886866e-	-	223.6528
-	08	9.3816	37

Author biography

Reza Asakereh A good guy with bad fruits

Name	Adj. P- value	Z- score	Combined Score
E2F4 (human)		-1.95	18.87
E2F1 (human)		-1.57	18.41
TIL D.7. D	A L CORA		
Name	atabases in Use for GSEA Adj. P-	Z-	Combined
	value	score	Score
hsa-miR-193b-3p	1.574e-35	-6.36	558.85
hsa-miR-192-5p	4.991e-29	-6.77	485.40
hsa-miR-215-5p	6.399e-30	-5.18	384.69
hsa-miR-34a-5p	1.639e-7	-5.12	112.13
hsa-let-7b-5p	0.008490	-7.90	85.63
Table B.8: D	atabases in Use for GSEA		
Name		Z-	Combined
	value	score	Score
POU2F1 (human)	0.03246	-1.74	15.68
Tabla P.O. D	atabases in Use for GSEA		
Name		Z-	Combined
	value	score	Score
E2F4 (human)	3.861e-8	-1.97	44.58
E2F1 (human)	0.005012	-1.55	15.12
NRF1 (human)	0.01787	-1.71	14.02
Table B 10: F	Patabases in Use for GSEA		
Name		<u>Z</u> -	Combined
2 TOURS	value	score	Score
hsa-miR-193b-3p	1.275e-39	-6.36	618.57
hsa-miR-192-5p	4.036e-18	-6.77	315.22
hsa-miR-215-5p	2.494e-19	-5.18	258.08
hsa-miR-34a-5p	5.211e-14	-5.12	188.50
hsa-let-7b-5p	0.00006949	-7.90	121.79

	Table	B.11:	Databases	in	Use	for	GSEA
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Name	1	Z- Combined score Score
Glycosaminoglycan degradation	0.02246	- 3210.85 613.78
Glycosphingolipid biosynthesis	0.4322	- 1321.91 900.86
RIG-I-like receptor signaling pathway		-97.67 704.97
Autoimmune thyroid disease Graft-versus-host disease		-78.88 653.40 -68.48 636.28

Table B.12: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
Endosomal/Vacuolar pathway_Homo sapiens_R-HSA-1236977	0.000001320	-1.98	37.94
Interferon alpha/beta signaling_Homo sapiens_R-HSA-909733	0.00001904	-1.86	29.37
Antigen Presentation: Folding, assembly and peptide loading of class I MHC_Homo sapiens_R-HSA-983170	0.00002778	-1.93	28.99
Immune System_Homo sapiens_R-HSA-168256	0.0001276	-2.20	28.61
Interferon gamma signaling_Homo sapiens_R-HSA-877300	0.00008257	-1.76	24.06

Table B.13: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
DNA replication 6.096e-17	-49.69	1855.26	
Homologous recombination	5.327e-9	-49.65	1100.04
Oocyte meiosis	1.312e-10	-26.82	704.47
Fanconi anemia pathway	0.000001224	-41.92	687.64
Small cell lung cancer	0.0006399	-63.94	602.21

Table	B 14.	Databases	in Use	for	GSEA

Name	Adj. P-value	Z- score	Combined Score
E2F transcription factor network Homo	1.616e-20	-1.73	85.49
sapiens_bb4d0fd3-6191-11e5-8ac5-06603eb7f303	1.0106-20	-1.70	00.49
PLK1 signaling events_Homo sapiens_e5e87977-6194-	1.127e-20	-1.66	83.87
11e5-8ac5-06603eb7f303	0.700 . 10	1 71	FO C7
FOXM1 transcription factor network_Homo sapiens_c51cda49-6192-11e5-8ac5-06603eb7f303	2.769e-16	-1.51	58.67
Aurora B signaling_Homo sapiens_304a75af-618c-11e5-	5.779e-18	-1.26	54.28
8ac5-06603eb7f303	. === =	4.40	10.00
ATR signaling pathway_Homo sapiens_8991cbac-618b-11e5-8ac5-06603eb7f303	4.751e-15	-1.18	42.36

Table B.15: Databases in Use for GSEA

Name	Adj. P	- Z-	Combined
	value	score	Score
Cell Cycle_Homo sapiens_R-HSA-1640170	1.426e-117	_	677.98
Cell Cycle, Mitotic_Homo sapiens_R-HSA-69278	1.227e-106		616.03
M Phase_Homo sapiens_R-HSA-68886 Cell Cycle Checkpoints_Homo sapiens_R-HSA-69620	2.552e-50	-2.43	290.68
	6.168e-44	-2.34	244.94
G2/M Checkpoints_Homo sapiens_R-HSA-69481	2.432e-40	-2.34 -2.32	222.45

Table B.16: Databases in Use for GSEA

J	Z-	Combined
value	score	Score
0.1424	-	2091.24
	585.89	
0.00003361	-80.82	1151.46
0.00002734	-69.21	1041.32
0.00004041	-68.48	923.26
0.00003361	-54.70	767.09
	value 0.1424 0.00003361 0.00002734 0.00004041	value score 0.1424 - 585.89 0.00003361 -80.82 0.00002734 -69.21 0.00004041 -68.48

Table	R 17	: Database	s in	Use	for	GSEA

Name	Adj. P-value	Z- score	Combined Score
TNF receptor signaling pathway_Homo	0.0001737	-1.53	19.39
apiens_316be05e-6196-11e5-8ac5-06603eb7f303 ΓRAIL signaling pathway_Homo sapiens_3a79fddf- 6196-11e5-8ac5-06603eb7f303	0.0001737	-1.41	18.49
L12-mediated signaling events_Homo	0.0008804	-1.54	16.39
apiens_7acdea19-6193-11e5-8ac5-06603eb7f303 Downstream signaling in naive CD8+ T cells_Homo	0.0009973	-1.43	14.59
apiens_92180cef-6191-11e5-8ac5-06603eb7f303 L2-mediated signaling events_Homo sapiens_a2a1883c-	0.002484	-1.55	14.07

Table B.18: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
Immune System_Homo sapiens_R-HSA-168256	0.00001975	-2.23	37.84
Cytokine Signaling in Immune system_Homo sapiens_R-HSA-1280215	0.00003972	-2.40	37.28
Endosomal/Vacuolar pathway_Homo sapiens_R-HSA-1236977	0.00004649	-1.90	28.15
Interferon alpha/beta signaling_Homo sapiens_R-HSA-909733	0.00004649	-1.84	27.00
Interferon gamma signaling_Homo sapiens_R-HSA-877300	0.00006090	-1.75	24.90

Table B.19: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
Glyoxylate and dicarboxylate metabolism	0.5917	- 701.48	867.99
Glycosaminoglycan biosynthesis	0.3410	- 406.31	856.52
Homologous recombination	0.00001242	-50.16	744.50
Small cell lung cancer	0.001191	-67.62	628.70
DNA replication	0.00006099	-49.02	624.51

Table B.20: Databases in Use for GSEA				
Name	Adj. P-	Z-	Combined	
	value	score	Score	
PLK1 signaling events_Homo sapiens_e5e87977-6194-	1.853e-13	-1.66	55.78	
11e5-8ac5-06603eb7f303				
FOXM1 transcription factor network_Homo	1.887e-11	-1.65	46.63	
$sapiens_c51cda49-6192-11e5-8ac5-06603eb7f303$				
E2F transcription factor network_Homo	3.319e-11	-1.68	45.91	
sapiens_bb4d0fd3-6191-11e5-8ac5-06603eb7f303				
p73 transcription factor network_Homo	8.821e-10	-1.38	32.22	
sapiens a88c505e-6194-11e5-8ac5-06603eb7f303				
ATR signaling pathway_Homo sapiens_8991cbac-618b-11e5-8ac5-06603eb7f303	1.914e-10	-1.24	31.11	
1169-0969-000096041909				

Table B.21: Databases in Use for GSEA				
Name	Adj. P-	Z-	Combined	
	value	score	Score	
Cell Cycle_Homo sapiens_R-HSA-1640170	3.743e-54	-2.46	317.68	
Cell Cycle, Mitotic_Homo sapiens_R-HSA-69278	5.027e-44	-2.47	259.31	
Mitotic Prometaphase_Homo sapiens_R-HSA-68877	1.905e-22	-2.03	111.73	
M Phase_Homo sapiens_R-HSA-68886	2.687e-18	-2.41	107.58	
Resolution of Sister Chromatid Cohesion_Homo	1.644e-20	-2.06	103.70	
sapiens_R-HSA- 2500257				
Intestinal immune network for IgA production	0.000003880	-99.15	1739.13	

Table B.22: Databases in Use for G	SEA			
Name	Adj. value	Р-	Z- score	Combined Score
			SCOTC	
Glycosphingolipid biosynthesis	0.8842		- 781.54	574.04
TNF signaling pathway	0.06974		-74.45	391.55
Autoimmune thyroid disease	0.08780		-77.59	383.98
Asthma	0.09760		-81.53	76.60

Table B.23: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
Immune System_Homo sapiens_R-HSA-168256	0.0003718	-2.23	31.24
Immunoregulatory interactions between a Lymphoid and a non-Lymphoid cell_Homo sapiens_R-HSA-198933	0.0006682	-2.00	25.45
Adaptive Immune System_Homo sapiens_R-HSA-1280218	0.006699	2.27	22.67
CD22 mediated BCR regulation_Homo sapiens_R-HSA-5690714	0.009758	-1.94	17.65
TNF receptor superfamily (TNFSF) members mediating non-canonical NF-kB pathway_Homo sapiens_R-HSA-5676594	0.009595	-1.88	17.61

Table B.24: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
Small cell lung cancer DNA replication	0.002421	-64.35	525.51
	0.0004052	-48.35	515.36
Homologous recombination Fanconi anemia pathway	0.0004032 0.0005634 0.001818	-48.79 -40.98	488.48 355.37
Human T-cell leukemia virus 1 infection	0.0003497	-40.98 -18.10	242.43

Table B.25: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
E2F transcription factor network_Homo sapiens bb4d0fd3-6191-11e5-8ac5-06603eb7f303	6.635e-13	-1.78	57.97
FOXM1 transcription factor network_Homo sapiens c51cda49-6192-11e5-8ac5-06603eb7f303	6.102e-9	-1.65	37.63
• —	2.976e-7	-1.47	26.78
Aurora B signaling_Homo sapiens_304a75af-618c-11e5-8ac5-06603eb7f303	9.910e-8	-1.26	24.67
Regulation of retinoblastoma protein_Homo sapiens_407a3468-6195-11e5-8ac5-06603eb7f303	0.000003482	-1.52	23.55

Table B.26: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
Cell Cycle_Homo sapiens_R-HSA-1640170	1.430e-40	-2.46	241.02
Cell Cycle, Mitotic_Homo sapiens_R-HSA-69278	7.756e-36	-2.47	213.08
Mitotic G1-G1/S phases_Homo sapiens_R-HSA-453279	1.077e-18	-2.11	98.22
G1/S Transition_Homo sapiens_R-HSA-69206	2.942e-15	-2.11	80.21
Mitotic Prometaphase_Homo sapiens_R-HSA-68877	1.569e-15	-2.02	78.71

Table B.27: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
antigen processing and presentation of exogenous peptide antigen via MHC class I, TAP-independent (GO:0002480)	0.0001851	-2.33	35.76
type I interferon signaling pathway (GO:0060337)	0.0002905	-2.32	30.11
retrograde transport, vesicle recycling within Golgi (GO:0000301)	0.02414	-4.03	29.32
regulation of necrotic cell death (GO:0010939) regulation of necroptotic process (GO:0060544)	$\begin{array}{c} 0.003712 \\ 0.0002905 \end{array}$	-2.48 -1.62	23.91 20.96

Table B.28: Databases in Use for GSEA

Name	Adj. P- value	Z- score	Combined Score
death-inducing signaling complex (GO:0031264)	0.01076	-3.30	22.26
recycling endosome membrane (GO:0055038)	0.00003093	-1.46	21.80
integral component of lumenal side of endoplasmic retic-	0.001205	-1.95	20.60
ulum membrane (GO:0071556)			
COPII-coated ER to Golgi transport vesicle	0.001228	-2.06	19.47
(GO:0030134)			
MHC protein complex (GO:0042611)	0.001980	-2.05	17.74

	Table B.29:	Databases	in	Use	for	GSEA
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Name	Adj. P-	Z-	Combined
	value	score	Score
DNA metabolic process (GO:0006259) DNA replication (GO:0006260) DNA repair (GO:0006281) mitotic cell cycle phase transition (GO:0044772) chromatin remodeling at centromere (GO:0031055)	1.366e-44	-1.36	147.85
	1.493e-35	-1.56	135.24
	3.977e-26	-1.71	109.74
	2.146e-32	-1.17	92.67
	1.246e-15	-2.28	88.85

Table B.30: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
DNA helicase activity (GO:0003678) ATPase activity (GO:0016887) DNA-dependent ATPase activity (GO:0008094)	7.421e-10	-2.66	66.98
	4.563e-9	-2.31	53.57
	3.454e-13	-1.53	51.82
DNA binding (GO:0003677)	1.223e-16	-1.20	50.97
3'-5' DNA helicase activity (GO:0043138)	0.000003135	-2.82	44.63

Table B.31: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
nuclear chromosome part (GO:0044454)	4.133e-37	-1.25	111.11
chromosome, centromeric region (GO:0000775)	5.429e-20	-2.25	108.81
microtubule organizing center (GO:0005815)	1.986e-18	-2.02	88.76
spindle (GO:0005819)	1.837e-29	-1.18	83.06
chromatin (GO:0000785)	1.158e-19	-1.66	78.42

Table B.32: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
retrograde transport, vesicle recycling within Golgi (GO:0000301)	0.009160	-4.03	37.61
cytokine-mediated signaling pathway (GO:0019221)	6.018e-9	-1.35	35.45
type I interferon signaling pathway (GO:0060337)	0.001496	-2.32	29.10
regulation of lymphocyte activation (GO:0051249)	0.005443	-2.70	27.84
antigen processing and presentation of exogenous peptide antigen via MHC class I, TAP-independent (GO:0002480)	0.001988	-2.33	27.55

Table B.33: Databases in Use for GSEA

Name	Adj. P- value	- Z- score	Combined Score
cytokine receptor activity (GO:0004896)	0.0002861	-1.84	25.68
tumor necrosis factor-activated receptor activi (GO:0005031)	ty 0.04099	-2.56	20.28

Table B.34: Databases in Use for GSEA

Name	Adj.	P-	Z-	Combined
	value		score	Score
clathrin vesicle coat (GO:0030125)	0.03999		-3.52	19.54
integral component of lumenal side of endoplasmic retic-	0.002593	3	-1.94	19.08
ulum membrane $(GO:0071556)$				
MHC protein complex (GO:0042611)	0.003483	3	-2.08	18.73
T cell receptor complex (GO:0042101)	0.02267		-2.55	16.46
clathrin coat of trans-Golgi network vesicle (GO:0030130)	0.05913		-3.12	15.73

Table B.35: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
strand displacement (GO:0000732)	1.727e-11	-2.58	75.45
kinetochore organization (GO:0051383)	2.163e-11	-2.56	73.81
mitotic nuclear division (GO:0140014)	1.157e-11	-2.22	65.92
DNA biosynthetic process (GO:0071897)	1.520e-12	-1.97	62.81
DNA replication (GO:0006260)	2.687e-15	-1.55	61.43

Table B.36: Databases in Use for GSEA

Name	Adj.	P-	Z-	Combined
	value		score	Score
DNA helicase activity (GO:0003678)	0.00000	9750	-2.62	38.96
microtubule motor activity (GO:0003777)	1.938e-	7	-1.85	37.43
ATPase activity (GO:0016887)	0.00000	3774	-2.31	37.41
motor activity (GO:0003774)	2.992e-	7	-1.58	30.62
DNA-dependent ATPase activity (GO:0008094)	4.744e-	7	-1.52	28.12

Table B.37: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
chromosome, centromeric region (GO:0000775) condensed chromosome, centromeric region (GO:0000779)	4.185e-16	-2.26	89.52
	5.224e-12	-2.09	61.06
spindle microtubule (GO:0005876)	5.953e-12	-1.95	56.22
condensed chromosome kinetochore (GO:0000777)	3.066e-10	-2.22	54.12
spindle (GO:0005819)	1.758e-16	-1.18	48.64

Table B.38: Databases in Use for GSEA

Name	Adj. P-		Combined
	value	score	Score
renal filtration (GO:0097205)	0.002162	-3.50	38.03
phagocytosis, engulfment (GO:0006911)	0.0001266	-2.39	35.59
regulation of immune effector process (GO:0002697)	0.001193	-2.89	34.48
glomerular filtration (GO:0003094)	0.001895	-3.00	33.73
positive regulation of lymphocyte activation (GO:0051251	0.0001266	-2.21	32.93

Table B.39: Databases in Use for GSEA

Name	Adj. value	P-	Z- score	Combined Score
immunoglobulin receptor binding (GO:0034987)	0.00008	611	-2.41	36.35

Table B.40: Databases in Use for GSEA

Name	Adj. P-value	Z- score	Combined Score
DNA metabolic process (GO:0006259) microtubule cytoskeleton organization involved in mitosis (GO:1902850)	1.976e-20 4.370e-12	-1.36 -1.65	71.65 53.05
sister chromatid segregation (GO:0000819) mitotic spindle organization (GO:0007052) DNA replication (GO:0006260)	3.095e-9 3.027e-13 1.211e-11	-2.04 -1.39 -1.55	50.33 49.10 47.77

Table B.41:	Databagag	in Hao	for	CCEA
Table D.41:	Databases	in Use	IOI.	CODEA

Name	Adj. P- value	Z- score	Combined Score
histone kinase activity (GO:0035173)	0.0001513	-3.09	41.13
DNA helicase activity (GO:0003678)	0.0001706	-2.67	34.36
DNA polymerase binding (GO:0070182)	0.0004986	-2.31	25.27
3'-5' DNA helicase activity (GO:0043138)	0.004400	-2.80	22.77
DNA-dependent ATPase activity (GO:0008094)	0.00006878	-1.53	22.22

Table B.42: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
microtubule organizing center (GO:0005815) chromosome, centromeric region (GO:0000775) nuclear chromosome part (GO:0044454) centrosome (GO:0005813), 1.411e-10 condensed chromosome kinetochore (GO:0000777)	1.411e-10 2.555e-8 1.751e-13 -1.61 5.286e-7	-2.07 -2.21 -1.25 42.58 -2.27	54.44 45.55 42.91

Table B.43: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
STK17A_IDG_kinase_ARCHS4_coexpression P2RY10_IDG_GPCR_ARCHS4_coexpression MAP3K14_IDG_kinase_ARCHS4_coexpression GPR25_IDG_GPCR_ARCHS4_coexpression GPR174_IDG_GPCR_ARCHS4_coexpression	5.174e-28	-1.62	110.43
	1.710e-25	-1.55	95.41
	9.806e-18	-1.54	66.84
	1.197e-15	-1.77	66.61
	8.520e-17	-1.58	63.96

Table B.44: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
UCK2_IDG_kinase_ARCHS4_coexpression PKMYT1_IDG_kinase_ARCHS4_coexpression CHRNA9_IDG_ionchannel_ARCHS4_coexpression RIOK1_IDG_kinase_ARCHS4_coexpression	9.315e-75	-1.61	283.39
	2.303e-65	-1.53	235.44
	2.957e-46	-1.54	168.06
	7.258e-36	-1.60	136.09
PKN3_IDG_kinase_ARCHS4_coexpression	7.575e-35	-1.55	127.31

T-1-1- D	45.	Databases	: TT	c	CCEA	
Lable B	45.	Databases	ın Use	tor	C-SFA	

Name	Adj. P- value	Z- score	Combined Score
STK17A_IDG_kinase_ARCHS4_coexpression	6.729e-22	-1.62	88.38
GPR174_IDG_GPCR_ARCHS4_coexpression	2.546e-21	-1.62	84.65
P2RY10_IDG_GPCR_ARCHS4_coexpression	2.546e-21	-1.54	80.07
DYRK2_IDG_kinase_ARCHS4_coexpression	8.547e-12	-1.60	47.72
${\rm GPR171_IDG_GPCR_ARCHS4_coexpression}$	3.522e-10	-1.72	44.50

Table B.46: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
UCK2_IDG_kinase_ARCHS4_coexpression PKMYT1_IDG_kinase_ARCHS4_coexpression CHRNA9_IDG_ionchannel_ARCHS4_coexpression PKN3_IDG_kinase_ARCHS4_coexpression CSNK2A2_IDG_kinase_ARCHS4_coexpression	1.384e-31	-1.60	121.31
	1.384e-31	-1.54	117.12
	1.934e-23	-1.54	87.25
	3.962e-15	-1.55	57.22
	3.962e-15	-1.47	54.49

Table B.47: Databases in Use for GSEA

Name	Adj. P-	Z- Co	mbined
	value	score Sco	ore
P2RY10_IDG_GPCR_ARCHS4_coexpression MAP3K14_IDG_kinase_ARCHS4_coexpression STK38L_IDG_kinase_ARCHS4_coexpression GPR174_IDG_GPCR_ARCHS4_coexpression GPR152_IDG_GPCR_ARCHS4_coexpression	3.895e-7 0.000007491 0.00002773	-1.56 79. -1.55 30. -1.61 26. -1.60 23. -1.98 17.	67 45 70

Table B.48: Databases in Use for GSEA

Name	Adj. P-	Z-	Combined
	value	score	Score
PKMYT1_IDG_kinase_ARCHS4_coexpression	8.882e-32	-1.54	118.82
UCK2_IDG_kinase_ARCHS4_coexpression	1.443e-24	-1.60	95.40
CHRNA9_IDG_ionchannel_ARCHS4_coexpression	9.312e-17	-1.54	63.51
PKN3_IDG_kinase_ARCHS4_coexpression	9.496e-14	-1.56	52.99
CSNK2A2_IDG_kinase_ARCHS4_coexpression	6.583e-11	-1.46	39.83