Frederick National Laboratory for Cancer Research

sponsored by the National Cancer Institute



Cancer Type/Site Classification using Deep-Learning

(Preliminary presentation slides)

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Acknowledgements

NCI-DOE Pilot-1 Team

BIDS

- Drs. George Zaki, Andrew Weissman, Mark Jensen and Eric Stahlberg
- Amar Khalsa, Dr. Deb Hope
- Colleagues who reviewed the material



Feel free to follow-along

Github

• https://github.com/ravichas/ML-TC1

Introduction

This is part of the NCI-DOE knowledge/capability transfer efforts

 Share tools/techniques/solutions for cancer related problems. We often take a testcase and show how it works

- You would be able to take the test-case (code/scripts) and tune it to your needs
- We want to hear from you, please send us your feed-back



Motivation: Cancer Prediction vs Cancer Detection

- Cancer <u>Prediction</u> has been the major focus
 - Prognosis, Recurrence, Susceptibility

- Cancer <u>Detection</u> (classification of tumors/cancers) is lagging behind <u>Prediction</u> and we would like to share an application that might be useful
 - Detect/Identify cancer type at an early stage

Goal(s)/Questions

 Take genomic expression data from tumor/cancer samples and apply Deep-Learning to create cancer types/site(s) classifier models

Are the expression profiles unique?

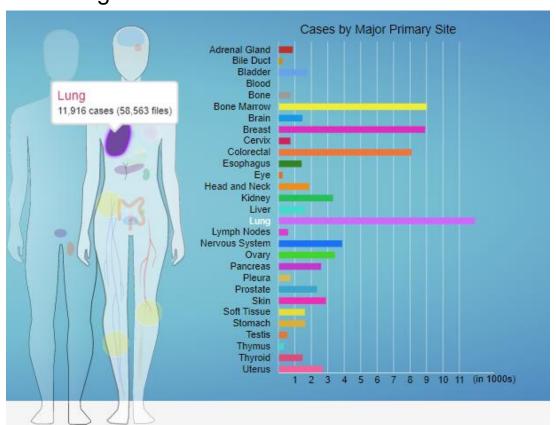
- Can we use the model as early cancer type detection
 - Improving chance of early detection cure/survival?



Cancer Burden

- Cancer is a group of diseases with worldwide risk
- Acquired or somatic changes causes 90-95% of caner (all types)
 - Source TCGA
- ~ 200 forms of cancer
 - DOI: 10.5114/wo.2014.47136
- For 2020
 - ~1.8M new cancer cases are expected
 - ~600K deaths will occur

Figure from Genomic Data Commons



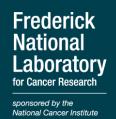


Expected New Cases/Deaths in 2020

New Cancer Cases

Between 2010 and 2020, we expect the number of new cancer cases in the United States to go up about 24% in men to more than 1 million cases per year, and by about 21% in women to more than 900,000 cases per year.

US population gender	Cancers that are expected to increase
Men	Prostrate, Kidney, Liver and Bladder
Women	Lung, Breast, Uterine and Thyroid



Dynamic genomic changes result in Cancer

Somatic alterations in oncogenes are the source of Transcript alterations

Article				
Genomi	ic basis for	RNA alte	rations ir	cancer

https://doi.org/10.1038/s41586-020-1970-0

Received: 29 March 2018

Accepted: 11 December 2019

Published online: 5 February 2020

Transcript alterations often result from somatic changes in cancer genomes. Various forms of RNA alterations have been described in cancer, including overexpression, altered splicing and gene fusions; however, it is difficult to attribute these to underlying genomic changes owing to heterogeneity among patients and tumor types, and the relatively small cohorts of patients for whom samples have been analyzed by both transcriptome and whole-genome sequencing.

Somatic alterations in the human cancer genome

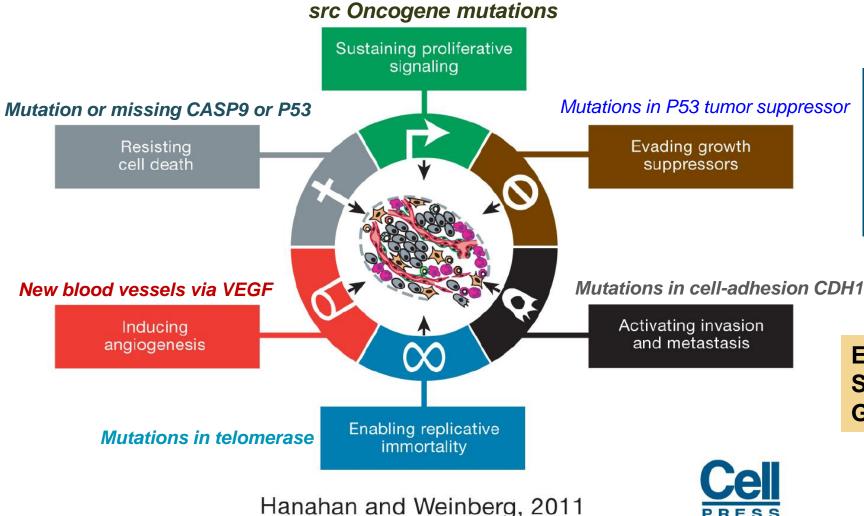
Barbara Weir, Xiaojun Zhao, and Matthew Meyerson*

CANCER CELL: NOVEMBER 2004 · VOL. 6 · COPYRIGHT © 2004 CELL PRESS

Hallmarks of cancer: Acquired capabilities (mutations) that drive cancer



Hallmarks of Cancer: The Next Generation



REVIEW | VOLUME 100, ISSUE 1, P57-70, JANUARY 07, 2000

The Hallmarks of Cancer

Douglas Hanahan A ≥ Robert A Weinberg

Open Archive • DOI: https://doi.org/10.1016/S0092-8674(00)81683-9

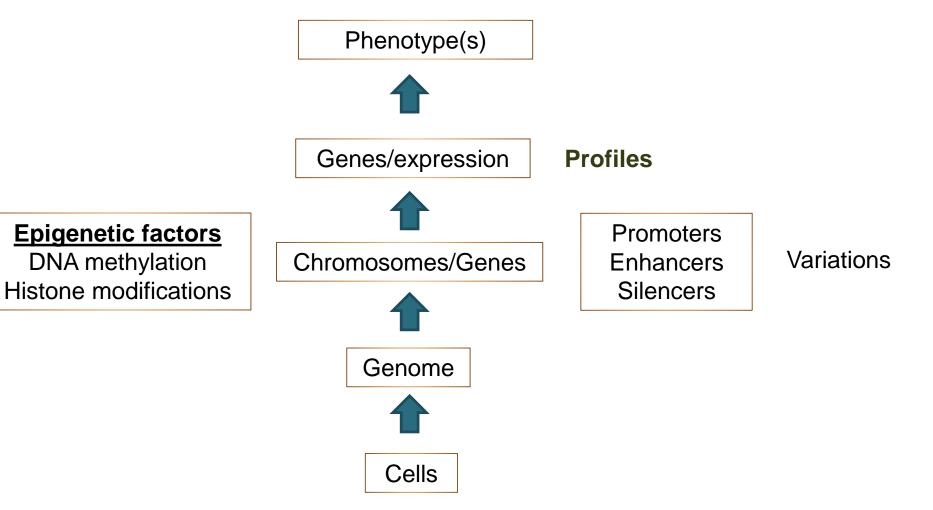
Expression changes in oncogenes; Six capabilities; Overview of **Genotype/phenotypes?**



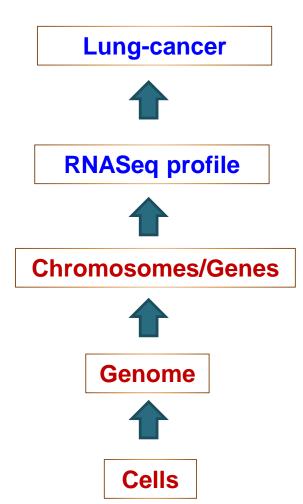


Influence of genomic features on phenotypes: An overview

DNA methylation



Influence of genomic features on phenotypes: An overview



Diagnosis/treatment vs Prediction





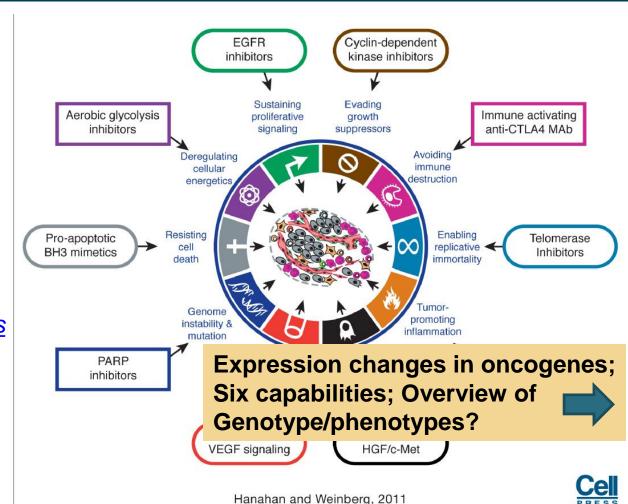
Treatment vs Type-Prediction

Treatment

- Gene-centric (or a slice of pathway)
- Imatinib targeting BCR/KIT

Detecting Type

- "The architecture of <u>occurring genetic aberrations</u> such as somatic mutations, CNVs, changed gene expression profiles, and different epigenetic alterations, is <u>unique</u> for each <u>type of cancer</u>.", DOI: 10.5114/wo.2014.47136
- Complex
- Multi-gene centric



Type-Prediction



The architecture of occurring genetic aberrations such as somatic mutations, CNV, changed gene expression profiles, and different epigenetic alterations, is unique for each type of cancer

DOI: 10.5114/wo.2014.47136

PERSPECTIVE

Understanding Genotype-Phenotype Effects in Cancer via Network Approaches

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Author Summary

Cancer is now increasingly studied from the perspective of dysregulated pathways, rather than as a disease resulting from mutations of individual genes. A pathway-centric view acknowledges the heterogeneity between genomic profiles from different cancer patients while assuming that the mutated genes are likely to belong to the same pathway and cause similar disease phenotypes. Indeed, network-centric approaches have proven to be helpful for finding genotypic causes of diseases, classifying disease subtypes, and identifying drug targets. In this review, we discuss how networks can be used to help understand patient-to-patient variations and how one can leverage this variability to elucidate interactions between cancer drivers.

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What kind of data do we need?

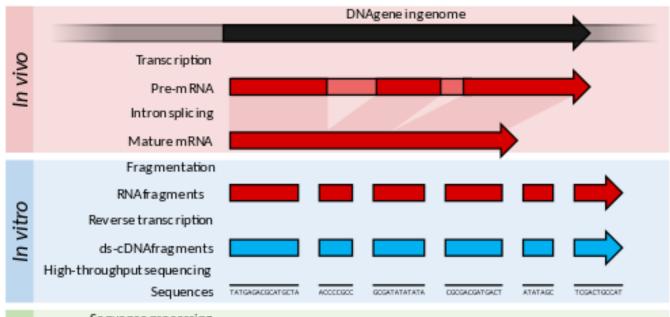
ta do we need?

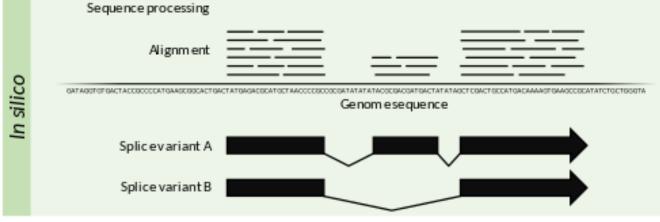
DNAgene ingenome

Transc ription

NGS

NGS





READS



Data source: The Cancer Genome Atlas (TCGA)

- NIH launched TCGA Pilot Project a public funded project
- Goal of creating a comprehensive "atlas" of cancer genomic profiles.
- Large cohorts of over <u>30 human tumors</u> through large-scale genome sequencing and integrated multi-dimensional analyses.
- Contains Microarray and NGS data
 - RNASeq
 - miRNA seq
 - SNP based platforms
 - **–**
- TCGA data is available via GDC

https://www.cancer.gov/about-nci/organization/ccg/research/structural-genomics/tcga

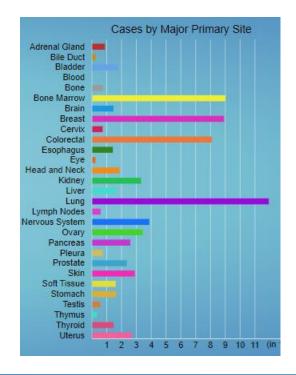


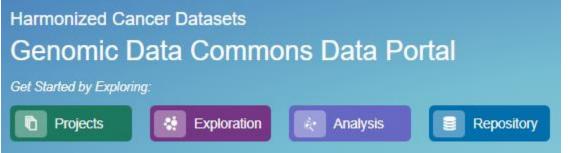
Data Harmonization: GDC (https://gdc.cancer.gov/)

 Data and metadata is submitted to the GDC in standard data types and file formats. Other data sources (Ex. TCGA) are also included

 Data are harmonized against a common reference genome (GRCh38)

 For this workshop, we will focus on TCGA Genomic expression data from GDC





Expression Data Quantification

- RC_g: Number of reads mapped to the gene
- RC_{g75}: The 75th percentile read count value for genes in the sample
- L: Length of the gene in base pairs;
 Calculated as the sum of all exons in a gene

$$FPKM-UQ = \frac{RC_g \times 10^9}{RC_{g75} \times L}$$

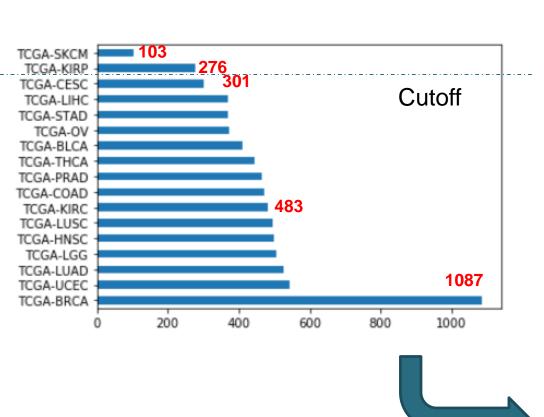
FASTQ Alignment to Ref Genome (SAM/BAM) **Quantification HTSeq** Gene Expression (FPKM-UQ)

Fragments Per Kilobase of transcript per Million mapped reads

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How much data for modeling?



CODE	Cancer Site/Type
BRCA	Breast invasive carcinoma
UCEC	Uterine Corpus Endometrial Carcinoma
LUAD	Lung adenocarcinoma
LGG	Brain Lower Grade Glioma
HNSC	Head and Neck squamous cell carcinoma
LUHSC	Lung squamous cell carcinoma
KIRC	Kidney renal clear cell carcinoma
PRAD	Prostate adenocarcinoma
COAD	Colon adenocarcinoma
THCA	Thyroid carcinoma
BLCA	Bladder Urothelial Carcinoma
OV	Ovarian serous cystadenocarcinoma
STAD	Stomach adenocarcinoma
LIHC	Liver hepatocellular carcinoma
CEC	Cervical squamous cell carcinoma and endocervical adenocarcinoma

300 samples each

sponsored by the National Cancer Institute

Expression data from a sample

Gene: AC090241.2 ENSG00000270112

Description novel transcript, antisense to ST8SIA5

Location Chromosome 18: 46,756,487-46,802,449 forward strand.

GRCh38:CM000680.2

About this gene This gene has 8 transcripts (splice variants)

Transcripts Hide transcript table

Gene: DNAH3 ENSG00000158486

Description dynein axonemal heavy chain 3 [Source:HGNC Symbol;Acc:HGNC:2949 &]

Gene Synonyms DKFZp434N074, DLP3, Dnahc3b, Hsadhc3

Location Chromosome 16: 20,933,111-21,159,441 reverse strand.

GRCh38:CM000678.2

About this gene This gene has 6 transcripts (splice variants), 371 orthologues, 14 paralogues and is a member of 1 Ensembl protein family.

Transcripts Hide transcript table

TCGA-BRCA

	Genes	¥	Expression	¥
	ENSG00000242268.2		1658.464179	
4	ENSG00000270112.3		460.2343433	
	ENSG00000167578.15		52440.10096	
	ENSG00000273842.1		0	
	ENSG00000078237.5		68165.45626	
	ENSG00000146083.10		255959.2351	
	ENSG00000225275.4		0	
	ENSG00000158486.12		104.9473768	
	ENSG00000198242.12		4968556.658	
	ENSG00000259883.1		6108.999052	
	ENSG00000231981.3		0	
	ENSG00000269475.2		0	
	ENSG00000201788.1		0	
	ENSG00000134108.11		957330.2056	
	ENSG00000263089.1		3484.027373	
	ENSG00000172137.17		41485.9507	
	ENSG00000167700.7		226717.4208	
	ENSG00000234943.2		2082.245035	
	ENSG00000240423.1		310.5246749	
	ENSG00000060642.9		155863.9216	
	ENSG00000271616.1		0	
	ENSG00000234881.1		0	
	ENSG00000236040.1		394.4755669	
	ENSG00000231105.1		1583.312582	
	ENSG00000243044.1		0	
	ENSG00000182141.8		45538.60648	
	ENSG00000269416.4		119.0847054	
	ENSG00000264981.1		0	

60,483 transcripts



Sample300

Data Preparation

Sample1

Breast Cancer • • • • • • • • •

Sample4

60,484 transcripts

Genes 💌 Expression	Genes
0242268.2 1658.464179	ENSG00000242268.2
0270112.3 460.2343433	ENSG00000270112.3
0167578.15 52440.10096	ENSG00000167578.15
0273842.1 0	ENSG00000273842.1
0078237.5 68165.45626	ENSG00000078237.5
0146083.10 255959.2351	ENSG00000146083.10
0225275.4 0	ENSG00000225275.4
0158486.12 104.9473768	ENSG00000158486.12
0198242.12 4968556.658	ENSG00000198242.12
0259883.1 6108.999052	ENSG00000259883.1
0231981 3 0	ENSG00000231981 3
0269475 2 0	ENSG00000269475 2
	ENSG00000203773.2
0134108 11 957330 2056	ENSG00000134108 11
	ENSG000000254280.11
	ENSG00000172137 17
	ENSG000001/2201.21
	ENSG00000237103.7
	ENSG000002340423.1
	ENSG00000240423.1
	ENSG000000771616.1
	ENSG00000271616.1
	ENSG00000234661.1
	ENSG00000236040.1
	ENSG00000231105.1
	ENSG00000243044.1 ENSG00000182141.8
	ENSG00000269416.4 ENSG00000264981.1
0264981.1 0	ENSGUUUUU264981.1

Sample2

Genes	▼ Expression ▼
ENSG00000242268.2	1658.464179
ENSG00000270112.3	460.2343433
ENSG00000167578.15	52440.10096
ENSG00000273842.1	0
ENSG00000078237.5	68165.45626
ENSG00000146083.10	255959.2351
ENSG00000225275.4	0
ENSG00000158486.12	104.9473768
ENSG00000198242.12	4968556.658
ENSG00000259883.1	6108.999052
ENSG00000231981.3	0
ENSG00000269475.2	0
ENSG00000201788.1	0
ENSG00000134108.11	957330.2056
ENSG00000263089.1	3484.027373
ENSG00000172137.17	41485.9507
ENSG00000167700.7	226717.4208
ENSG00000234943.2	2082.245035
ENSG00000240423.1	310.5246749
ENSG00000060642.9	155863.9216
ENSG00000271616.1	0
ENSG00000234881.1	0
ENSG00000236040.1	394.4755669
ENSG00000231105.1	1583.312582
ENSG00000243044.1	0
ENSG00000182141.8	45538.60648
ENSG00000269416.4	119.0847054
ENSG00000264981.1	0

Sample3

ENSG00000270112.3	460.2343433
ENSG00000167578.15	52440.10096
ENSG00000273842.1	0
ENSG00000078237.5	68165.45626
ENSG00000146083.10	255959.2351
ENSG00000225275.4	0
ENSG00000158486.12	104.9473768
ENSG00000198242.12	4968556.658
ENSG00000259883.1	6108.999052
ENSG00000231981.3	0
ENSG00000269475.2	0
ENSG00000201788.1	0
ENSG00000134108.11	957330.2056
ENSG00000263089.1	3484.027373
ENSG00000172137.17	41485.9507
ENSG00000167700.7	226717.4208
ENSG00000234943.2	2082.245035
ENSG00000240423.1	310.5246749
ENSG00000060642.9	155863.9216
ENSG00000271616.1	0
ENSG00000234881.1	0
ENSG00000236040.1	394.4755669
ENSG00000231105.1	1583.312582
ENSG00000243044.1	0
ENSG00000182141.8	45538.60648
ENSG00000269416.4	119.0847054
ENSG00000264981.1	0

	Expression *
	1658.464179
	460.2343433
5	52440.10096
	0
	68165.45626
0	255959.2351
	0
2	104.9473768
2	4968556.658
	6108.999052
	0
	0
	0
1	957330.2056
	3484.027373
7	41485.9507
	226717.4208
	2082.245035
	310.5246749
	155863.9216
	0
	0
	394.4755669
	1583.312582
	0
	45538.60648
	119.0847054

Sample297 Sample298

Genes	 Expression 	¥
SG00000242268.2	1658.464179	
SG00000270112.3	460.2343433	
SG00000167578.15	52440.10096	
SG00000273842.1	0	
SG00000078237.5	68165.45626	
SG00000146083.10	255959.2351	
SG00000225275.4	0	
SG00000158486.12	104.9473768	
SG00000198242.12	4968556.658	
SG00000259883.1	6108.999052	
SG00000231981.3	0	
SG00000269475.2	0	
SG00000201788.1	0	
SG00000134108.11	957330.2056	
SG00000263089.1	3484.027373	
SG00000172137.17	41485.9507	
SG00000167700.7	226717.4208	
SG00000234943.2	2082.245035	
SG00000240423.1	310.5246749	
SG00000060642.9	155863.9216	
SG00000271616.1	0	
SG00000234881.1	0	
SG00000236040.1	394.4755669	
SG00000231105.1	1583.312582	
SG00000243044.1	0	
SG00000182141.8	45538.60648	
SG00000269416.4	119.0847054	
SG00000264981.1	0	

Sample299



Data Preparation

	Sample1	Sample2	Sample3	Sample4	Sample297	Sample298	Sample299	Sample300
Breast Cancer	Cemes Convention Convent	Comparison Co	Comment Com		Service	Common Common		
Lung Cancer	Series				Descondent Des	Common Common		Series
					Genes	Genes	Genes Expression ENSGROUND42268.2 1558.464179	Genes
Kidney Cancer		Genetal Comparison Compar		Comparison Com	Decicionocida 1.0 2.0	INCOGNOCOPICE 3	NewGOODCOMPATION 1 - 30-340-410	MONOCONDONIAL MAN MAN

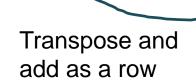


Merged Sample Expression Data

Genes

		0	1	2	3	4	5	6	7	8	9	6	0474	60475	60476	60477	60478	60479	60480	60481	60482	submitter_id
	0	574548	2263.14	983212	69718	54834.9	19718.1	175853	735123	38662.4	233190		0	0	0	0	0	0	0	0	0	TCGA-04-1331-01A-01R-1569-13
	1	352295	4592.37	663107	39745.4	36553.5	41147.1	241313	396423	37567	128693		0	0	0	0	0	0	0	0	0	TCGA-04-1332-01A-01R-1564-13
Щ	2	295162	649.026	1.21115e+06	57385.5	33097.4	58051.8	228615	346066	105567	408267		0	0	0	0	0	0	0	0	0	TCGA-04-1338-01A-01R-1564-13
7	3	329580	1835.59	1.08437e+06	33812.3	24516.1	22330.6	42134.4	895558	56178	83847.3		0	0	0	0	0	0	0	0	0	TCGA-04-1341-01A-01R-1564-13
#	4	289269	40061.7	2.44837e+06	26399.5	18248	49610	74761.1	571992	71951.9	98726.4		0	0	0	0	0	0	0	0	0	TCGA-04-1343-01A-01R-1564-13
SAMP																						
Z	4495	1.18093e+06	0	1.01139e+06	67877.2	15005.7	50527.3	6.21536e+06	1.47373e+06	459656	167488		0	0	0	0	0	0	0	0	0	TCGA-ZS-A9CD-01A-11R-A37K-07
	4496	929228	0	869800	95607.5	17188.6	9352.12	7.61121e+06	196838	354465	138074		0	0	0	0	0	0	0	0	0	TCGA-ZS-A9CE-01A-11R-A37K-07
	4497	469276	476.683	516938	110051	34469.4	37334.7	5.95811e+06	427832	323833	154861		0	0	0	0	0	0	0	0	0	TCGA-ZS-A9CF-01A-11R-A38B-07
	4498	2.44119e+06	18282.7	853547	79288.7	106926	42593.9	4.80111e+06	955338	331924	177020		0	0	0	0	0	0	0	0	0	TCGA-ZS-A9CG-01A-11R-A37K-07
	4499	259853	505,488	591328	74253.7	42553.5	118772	148978	508465	153862	170412		0	0	0	0	0	0	0	0	0	TCGA-ZX-AA5X-01A-11R-A42T-07

4500 rows × 60484 columns



Quantifying mRNA abundance and Scaling

- GDC harmonization data is provided in FPKM-UQ
- In our code, FPKM-UQ is rescaled to TPM using the following formula.

$$\mathsf{TPM}_i = \left(\frac{\mathsf{FPKM}_i}{\Sigma_j \mathsf{FPKM}_j}\right) \cdot 10^6$$

TPM has nice mathematical properties and a stable entity

https://docs.gdc.cancer.gov/Encyclopedia/pages/HTSeq-FPKM-UQ/

Mapping and quantifying mammalian transcriptomes by RNA-Seq



One-hot encoding to convert Cancer types to numbers

- Convenient to transform categorical variables into a numerical quantity for computations
 - BRCA to 0 ; LUAD to 1 etc.
 - 0, 1, 2, 3, ..., 13, 14, 15

```
>>> encoded
[0., 0., 0., 0., 0., 0., 0., 1., 0., 0., 0., 0., 0., 0., 0., 0.]
 [0., 0., 0., 0., 0., 0., 0., 0., 1., 0., 0., 0., 0., 0., 0., 0., 0.]
 [0., 0., 0., 0., 0., 0., 0., 0., 0., 1., 0., 0., 0., 0., 0., 0., 0.]
 dtvpe=float32)
```

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Data preparation steps summary

GDC dataset; https://portal.gdc.cancer.gov/

GDC • Filter Program/Projects (Ex. TCGA-BRCA) • Filter to retain only RNA-Seq data; FPKM-UQ scaling **GDC RNASeq** Primary Tumor samples only Tumor • Download final list and meta files **Extract** Restrict sample size >= 300 and extract each class to create a balanced dataset (each class = 300 samples) Samples • Download/merge individual expression data 4500 files ~ 2.5 GB; carried out in Biowulf Create Coding • Look in Clinical file to extract labels and attach them; one-hot encoding Prep

▼ Modeling

• Final dataset in the previous set will be used as input



Before we break for hands-on

 Python as the programming language for this workshop, but similar libraries are available in R or other languages





- Will use Jupyter Notebook for sharing the code
 - With little effort one can convert the Python code into R and still use Jupyter Notebook



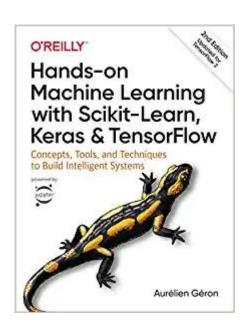
To be continued after hands-on

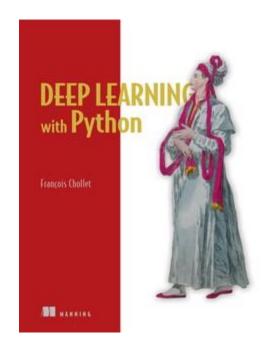
https://github.com/ravichas/ML-TC1



Before we begin the modeling section ...

Due to lack of time, I wont be covering the basics of Neural Network





These are good books for beginners and up

Keras is a high-level NN package that is built on top of popular high-level libraries (TF, Theano). Works well with CPU/GPU

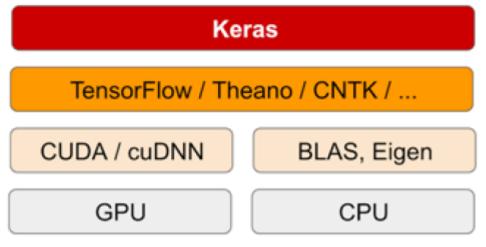


Figure from Deep Learning with Python

Supervised Learning

- Goal
 - Construct a model that takes in input features/target pair to return a prediction for target/outcome
- Train a machine learning
 - Model refers to learning its parameters, which typically involves minimizing a loss function on training data with the aim of making accurate predictions on unseen (test) data

Supervised Learning:

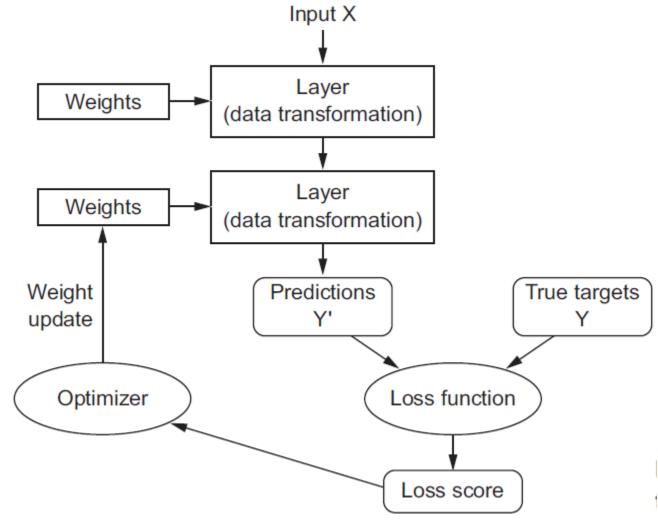
Data: (x,y); where x is the genomic expression profile; y is the cancer classes

Goal? Learn the function that maps $x \rightarrow y$



Deep Learning Procedure

Taken from Deep Learning with Keras book



Terminology

								_			_		50475	50475	50477	50470	50470	50400	50404	50402	
	0	1	2	3	4	5	6	/	8	9	6	004/4	604/5	60476	604//	60478	60479	60480	60481	60482	submitter_i
0	574548	2263.14	983212	69718	54834.9	19718.1	175853	735123	38662.4	233190	***	0	0	0	0	0	0	0	0	0	TCGA-04-1331-01A-01R-1569-1
1	352295	4592.37	663107	39745.4	36553.5	41147.1	241313	396423	37567	128693	•••	0	0	0	0	0	0	0	0	0	TCGA-04-1332-01A-01R-1564-1
2	295162	649.026	1.21115e+06	57385.5	33097.4	58051.8	228615	346066	105567	408267	***	0	0	0	0	0	0	0	0	0	TCGA-04-1338-01A-01R-1564-1
3	329580	1835.59	1.08437e+06	33812.3	24516.1	22330.6	42134.4	895558	56178	83847.3	***	0	0	0	0	0	0	0	0	0	TCGA-04-1341-01A-01R-1564-1
4	289269	40061.7	2.44837e+06	26399.5	18248	49610	74761.1	571992	71951.9	98726.4	***	0	0	0	0	0	0	0	0	0	TCGA-04-1343-01A-01R-1564-

Columns

input variables or features or attributes

Outcome column

Outcome variables or targets

Rows

- Training example or instance
- Whole table Training data set



A Simple Network

Input	Output
0.125	0.39
0.25	0.40
0.5	0.43
1	0.48
2	0.58
3	???

Data based on Mary Attenborough, in Mathematics for Electrical Engineering and Computing, 2003

What is different about Neural Network?

If you know the equation (algorithm), then you feed in the input and you get the output.
 You can code the function yourself

```
def function(m):

L = 0.1 * m + 0.38

return(L)
```

You can choose to use linear modeling and use the data to figure the relationship

```
Model ← Im( L ~ m)
```

Neural Network using the data learn the algorithm.

INPUT

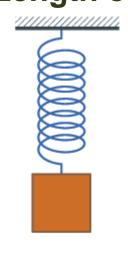
ALGORITHM

OUTPUT



A Simple Network

Input: Mass or M (kg)
Output: Length or L (m)



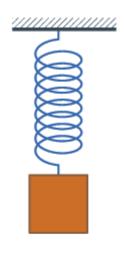
M L

Input	Output
0.125	0.39
0.25	0.40
0.5	0.43
1	0.48
2	0.58
3	???

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A Simple Network



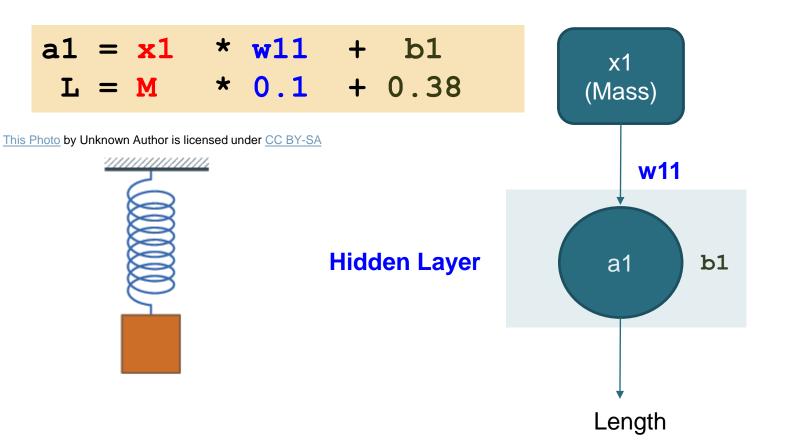
M	L
0.125	0.39
0.25	0.40
0.5	0.43
1	0.48
2	0.58
3	0.68

$$L = 0.1 * Mass + 0.38$$

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A Simple Network



M	L
0.125	0.39
0.25	0.40
0.5	0.43
1	0.48
2	0.58
3	0.68

These are the model variables: [array([[0.10058284]], dtype=float32), array([0.37793916], dtype=float32)]

Based on Mary Attenporougn, in Mathematics for Electrical Engineering and Computing, 2003



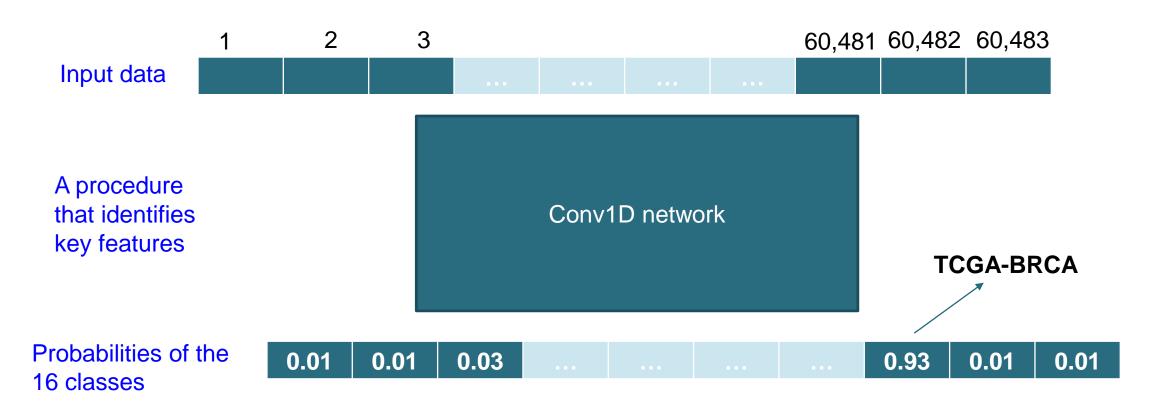
Error minimization

- Goal is to choose Ws such that predictions of the network should be close to y
- Error function or cost function a measure how good our predictions are
- Eventually, we wan to pick a set of w that minimizes the error function



Convolutional Neural Network

- We are going to take a vector of genomic expression values and feed them into a network with a series of operations to create a model
- Model is what we call convolutional-1D network



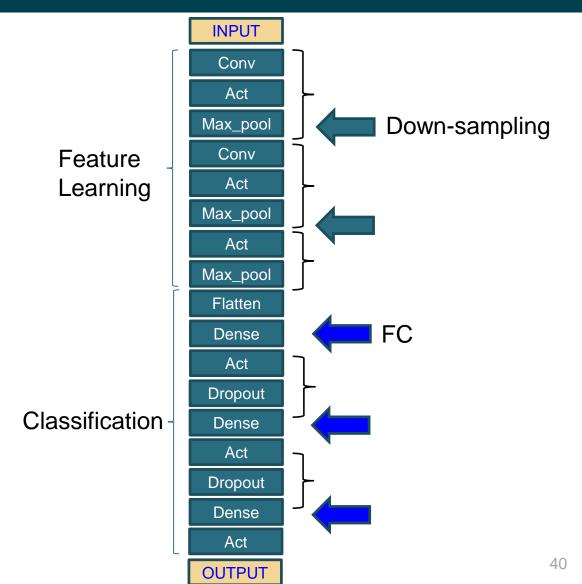


Components of conv1D

- 1. Activation (Act)
- 2. Convolution (Conv)
- 3. Maxpooling (Max_pool)
- 4. Flatten
- 5. Dense
- 6. Dropout

Topology of a network defines a "hypothesis space"

Choosing a specific topology is usually not straightforward and comes with practice.

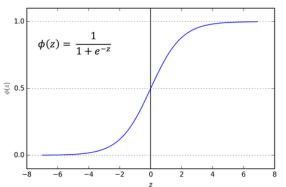




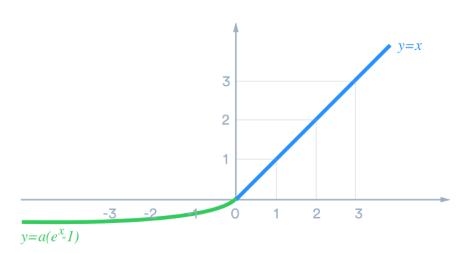
1. Activation Function

Activation functions are included to create non-linearity

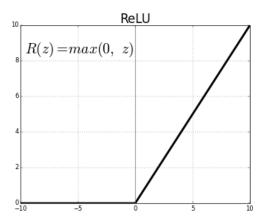
- Sigmoid
- ReLU
- Leaky ReLU
- ELU
- Maxout
- Tanh







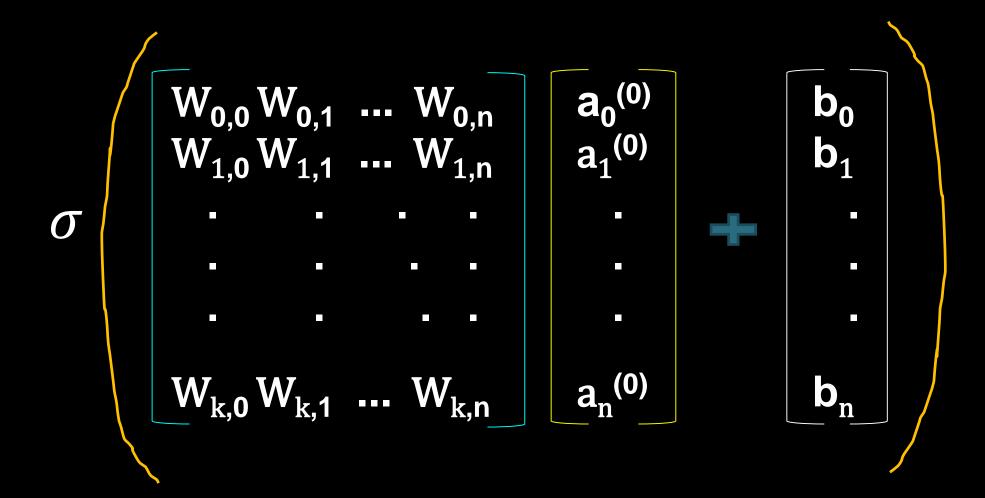
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$$a^{(L)} = \sigma(w^{(L)}a^{(L-1)} + b^{(L)})$$



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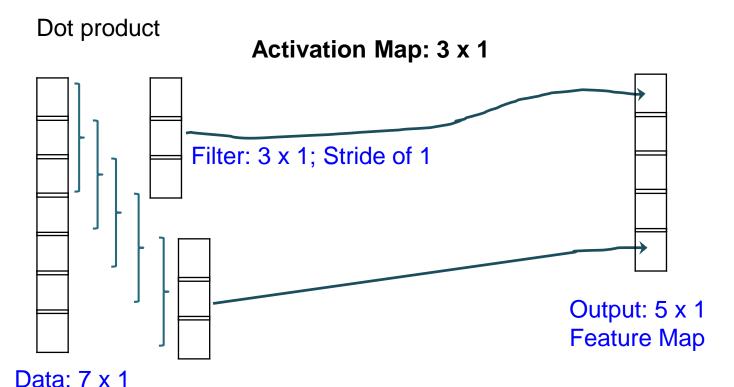




2. Convolution

Process of applying filter (<u>kernel</u>) to the data for the purpose of subsampling. Kernel is a matrix that has a smaller dimension than the input data creates chunks

Reduces the number of parameters and allow creation of deeper networks



Convolutional Layer

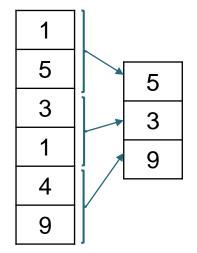
of HP

- # of filters
- Spatial Extent
- Stride
- Amount of zero padding



3. Pooling

- Pooling makes the representations smaller/manageable (downsampling) by retaining only important features; creates smaller clusters of manageable size
- Each activation map will be pooled separately.
- Common approach is Max Pooling



Max-pooling with filter size of 2x1 and stride of 2

Max Pooling Intuition:

Enhancing the signals by looking at a region and pick the maximum activation value

Each of these are activation and we are looking for

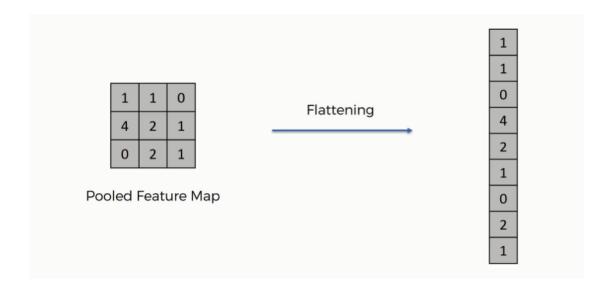
Research shows that zero-padding is not followed.

Because we are interested in down-sampling



4. Flatten

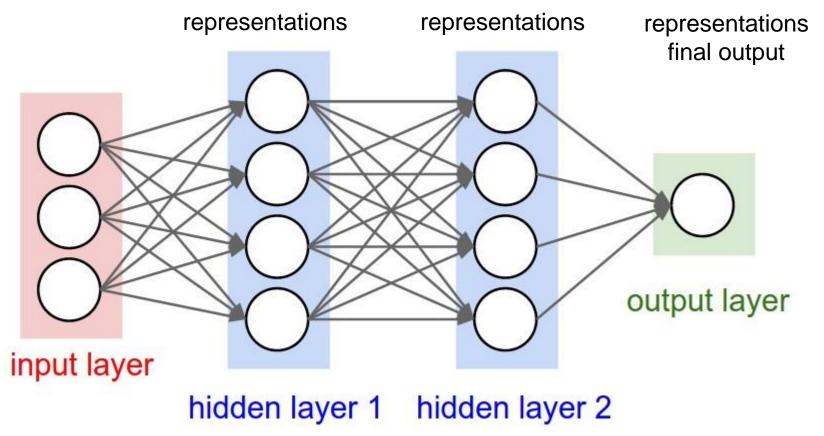
Procedure to transform a 2D matrix (features) to a 1D vector which in turn can be fed into a fully-connected layer (dense)





5. Dense

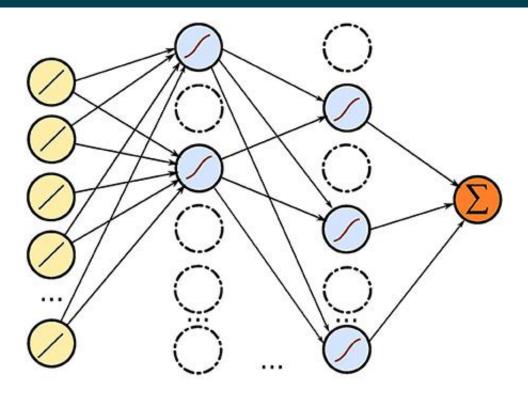
Each neuron receives input from all the neurons in the previous layer (densely connected)



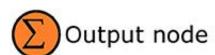
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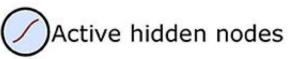
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6. Dropout



// Input nodes





()Inactive hidden nodes

Imbalance in the weights among the nodes can lead to some node weights not contributing to the learning

One solution:

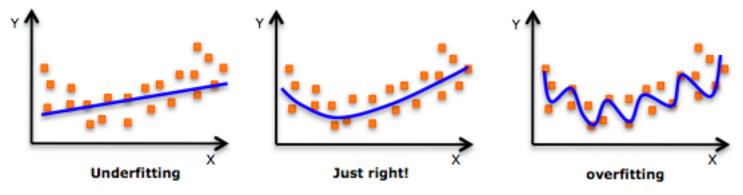
Remove a random proportion of selection of neurons in a neural network during training

Can help weak learners become strong learners

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6. Dropout



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Model Summary

1 0 128 10 1

1.0 128 10 1 Model: "sequential_1"			
Layer (type)	Output	Shape	Param #
convld_1 (ConvlD)	(None,	60464, 128)	2688
activation_1 (Activation)	(None,	60464, 128)	0
max_pooling1d_1 (MaxPooling1	(None,	60464, 128)	0
conv1d_2 (Conv1D)	(None,	60455, 128)	163968
activation_2 (Activation)	(None,	60455, 128)	0
max_pooling1d_2 (MaxPooling1	(None,	6045, 128)	0
flatten_1 (Flatten)	(None,	773760)	0
dense_1 (Dense)	(None,	200)	154752200
activation_3 (Activation)	(None,	200)	0
dropout_1 (Dropout)	(None,	200)	0
dense_2 (Dense)	(None,	20)	4020
activation_4 (Activation)	(None,	20)	0
dropout_2 (Dropout)	(None,	20)	0
dense_3 (Dense)	(None,	15)	315
activation_5 (Activation)	(None,	15)	0
Total params: 154,923,191 Trainable params: 154,923,191 Non-trainable params: 0	l		

INPUT Conv Act Max_pool Conv Act Max_pool Act Max_pool Flatten Dense Act Dropout Dense Act Dropout Dense

OUTPUT

~ 154 M parameters

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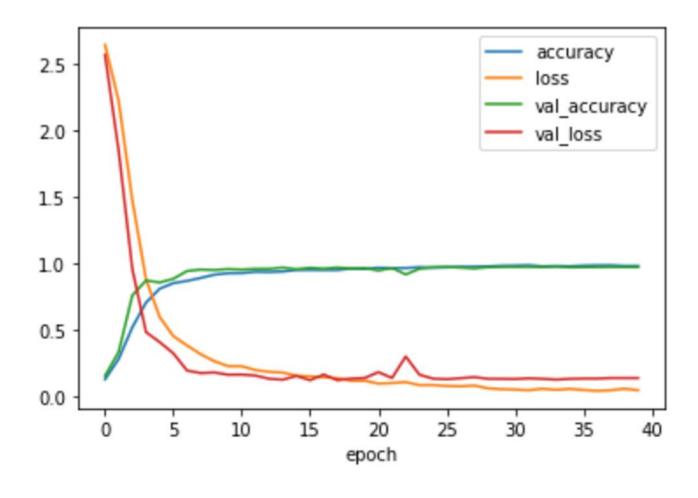
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Code execution and progress

```
Epoch 00001: val loss improved from inf to 2.56791, saving model to Pilotl.h5
Epoch 2/400
Epoch 00002: val loss improved from 2.56791 to 1.84441, saving model to Pmodel.h5
Epoch 3/400
Epoch 00003: val loss improved from 1.84441 to 0.95540, saving model to Pmodel.h5
Epoch 4/400
Epoch 00004: val loss improved from 0.95540 to 0.48347, saving model to Pmodel.h5
Epoch 5/400
Epoch 00005: val loss improved from 0.48347 to 0.40829, saving model to Pmodel.h5
Epoch 6/400
Epoch 00006: val loss improved from 0.40829 to 0.32363, saving model to Pmodel.h5
Epoch 7/400
Epoch 00007: val loss improved from 0.32363 to 0.19439, saving model to Pmodel.h5
Epoch 8/400
3375/3375 [========================== ] - 228s 68ms/step - loss: 0.3170 - accuracy: 0.8910 - val loss: 0.1754 - val acc:
Epoch 00008: val loss improved from 0.19439 to 0.17536, saving model to Pmodel.h5
Epoch 9/400
Epoch 00009: val loss did not improve from 0.17536
Epoch 10/400
Epoch 00010: val loss improved from 0.17536 to 0.16323, saving model to Pmodel.h5
```

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Model Performance





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Questions/Comments

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