



An Overview on the role of Artificial Intelligence (AI) and Deep Neural Networks (DNN) in cancers prediction, diagnosis, grading and prognosis

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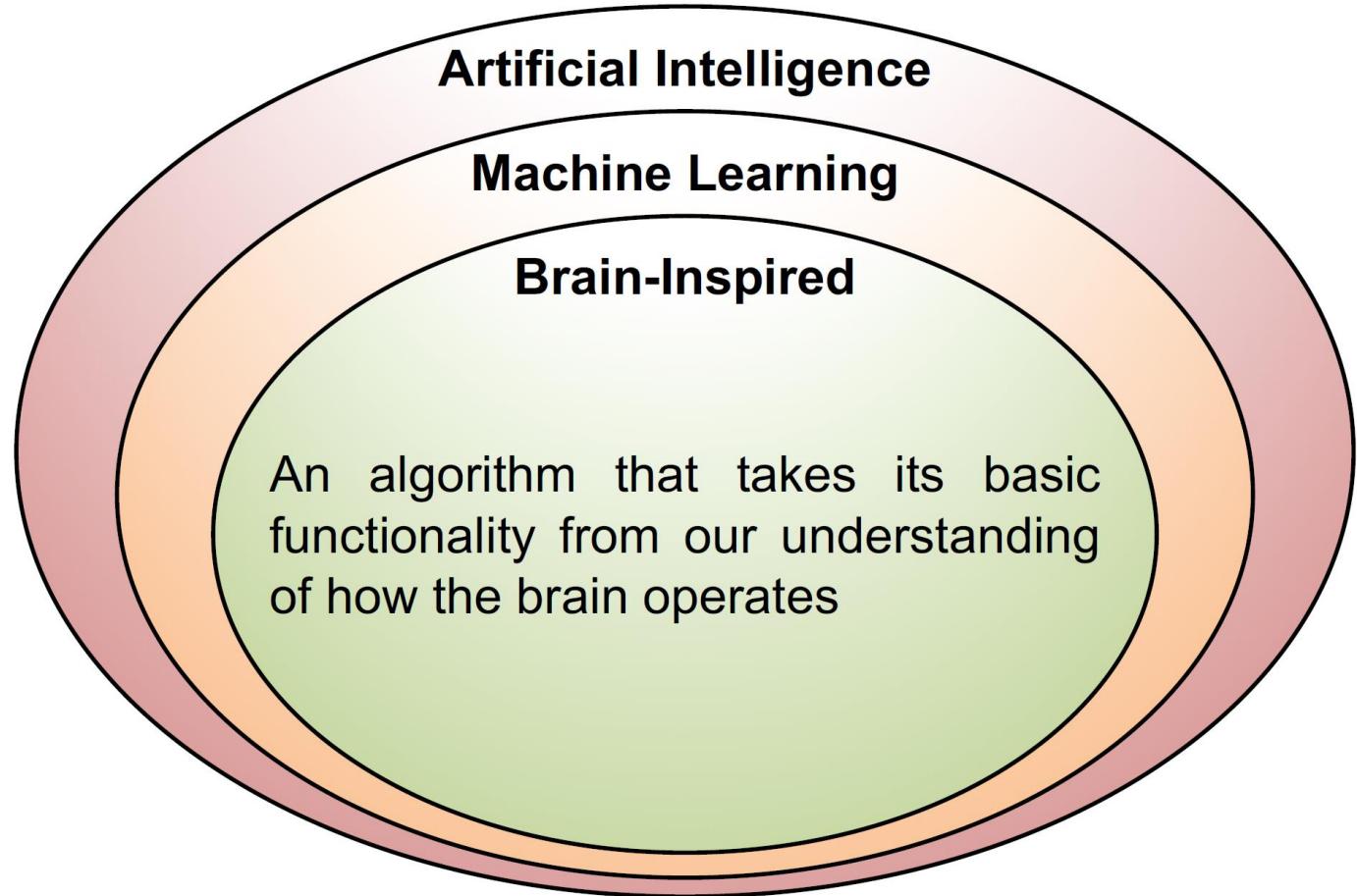
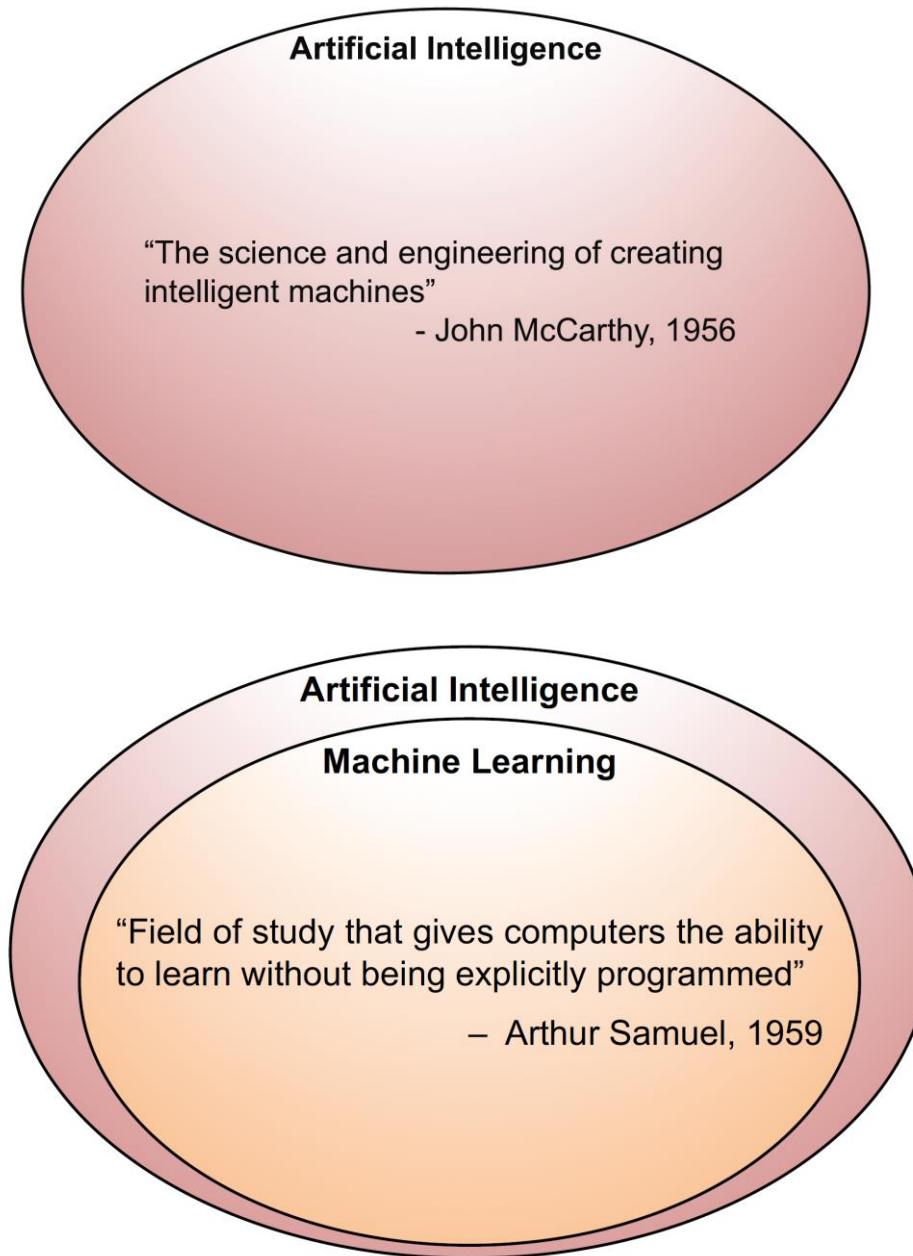
Springer Nature / Elsevier Scientific Reviewer.

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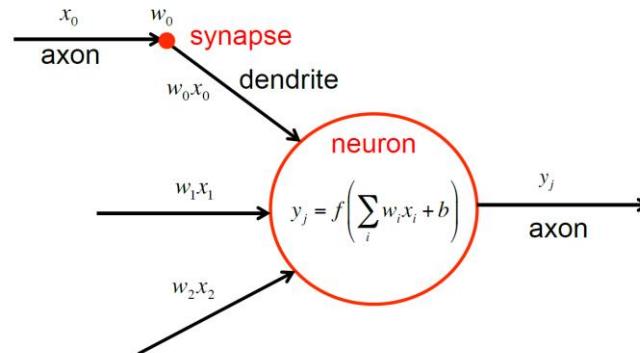
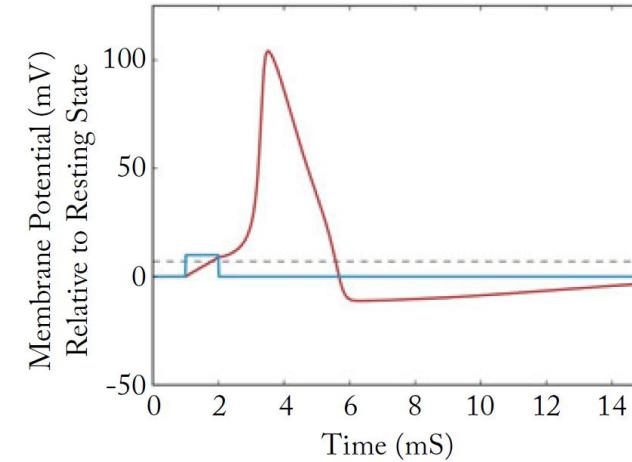
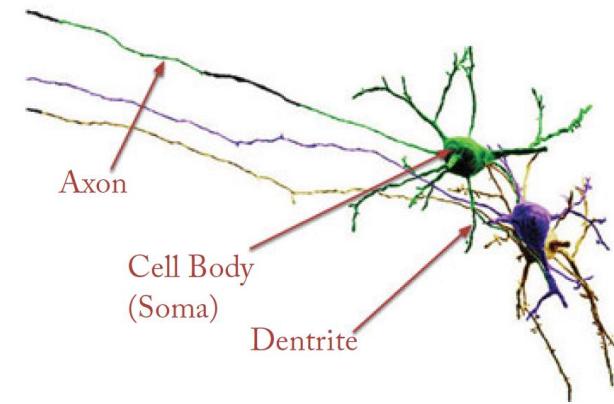
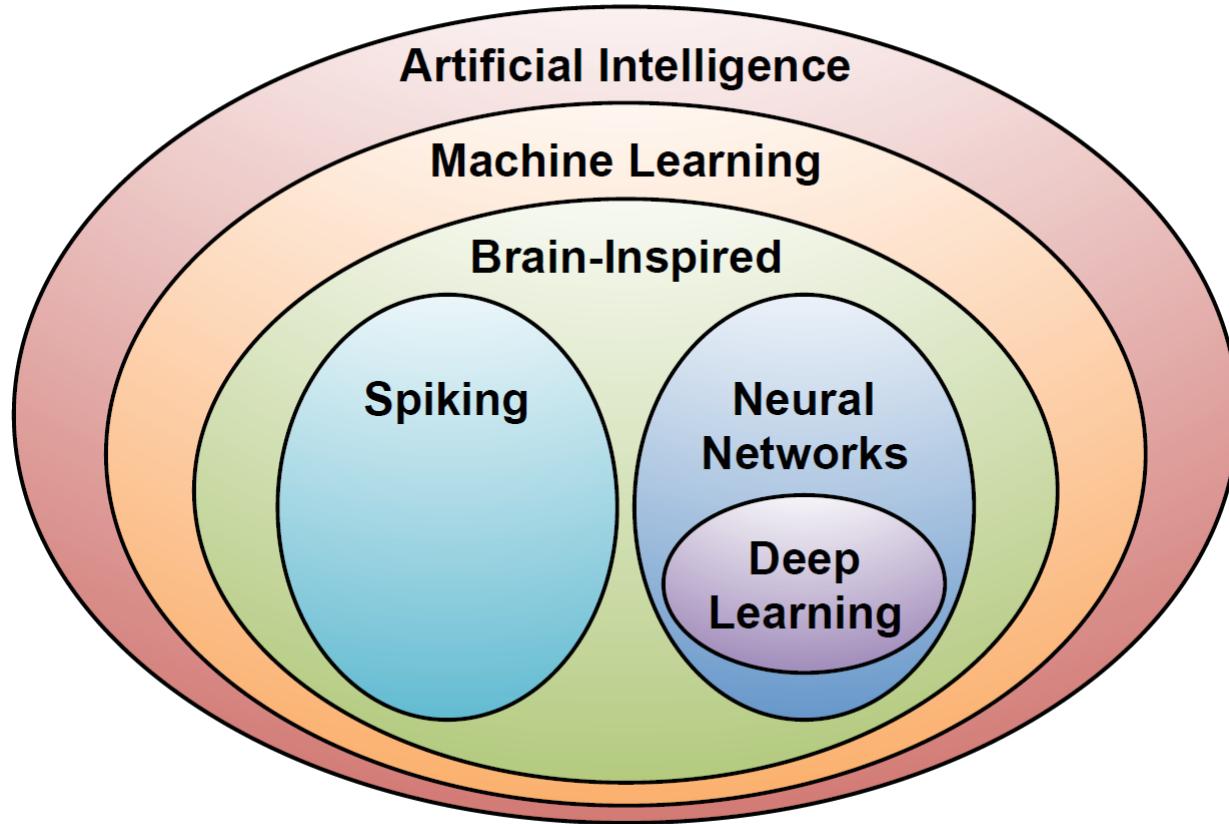
Nov 2021.



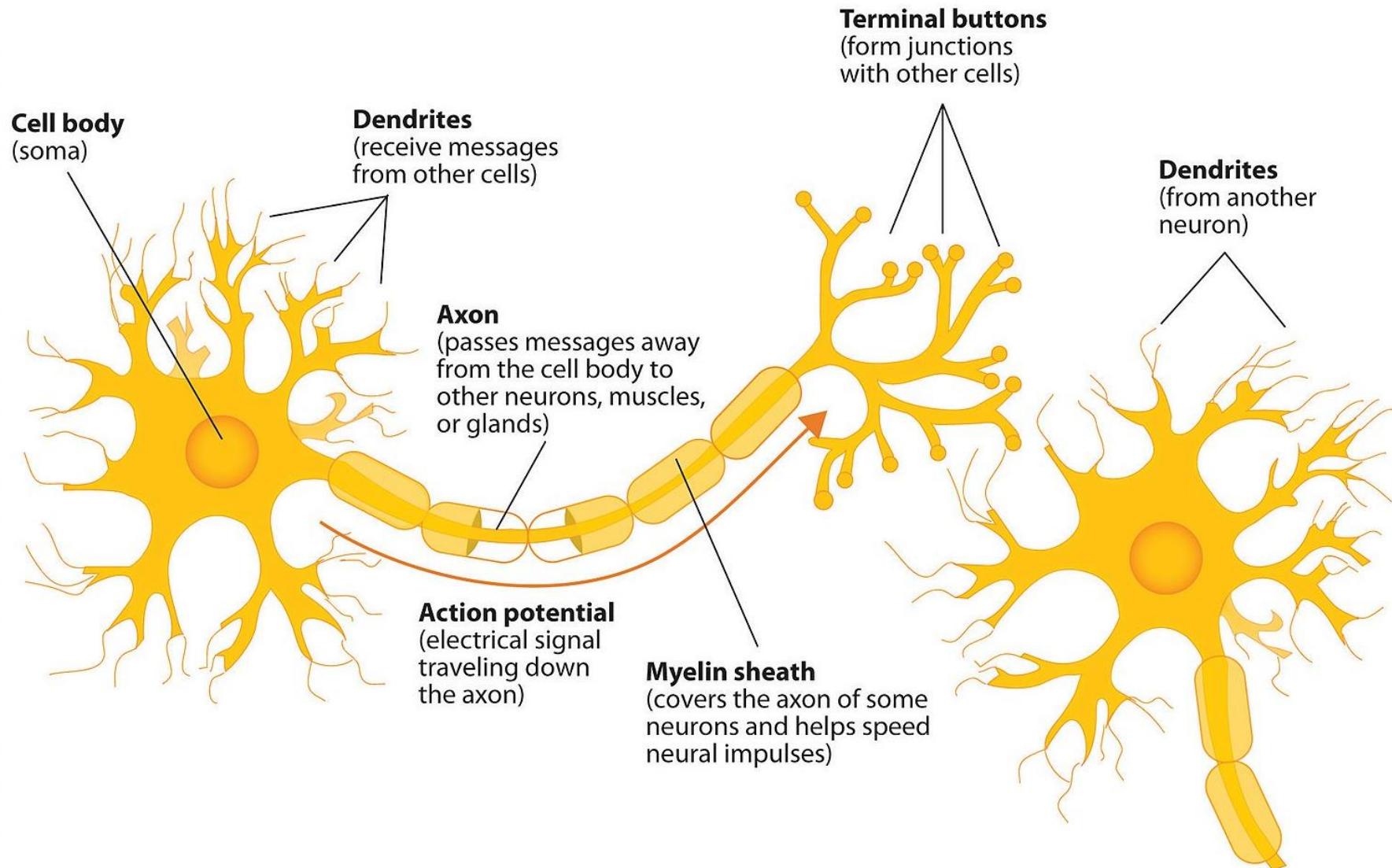
Artificial Intelligence (AI)



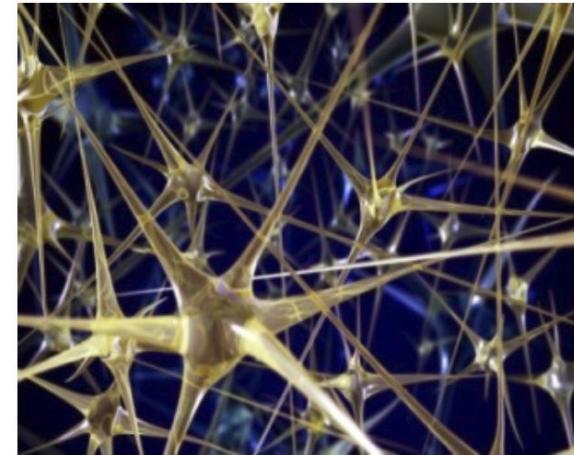
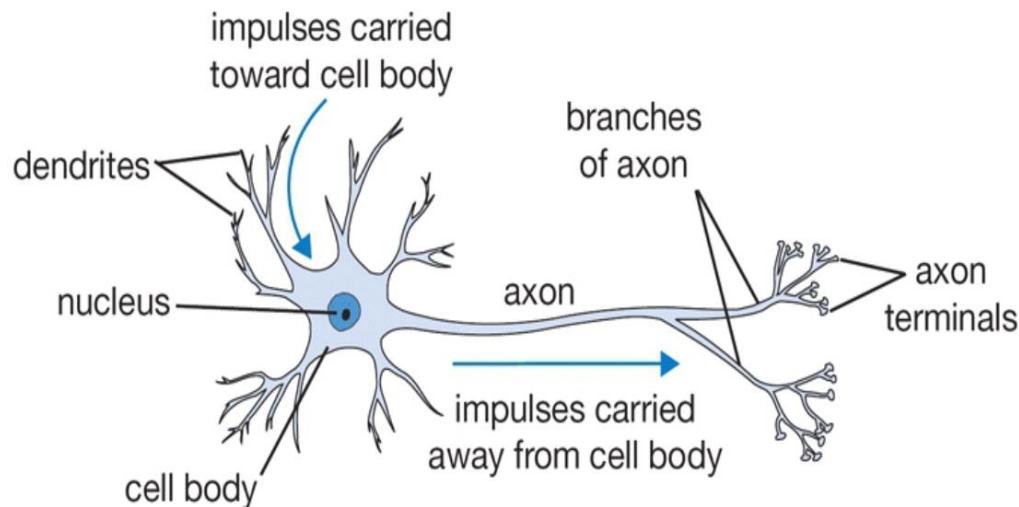
Deep Neural Networks (DNN)



Biological Neuron

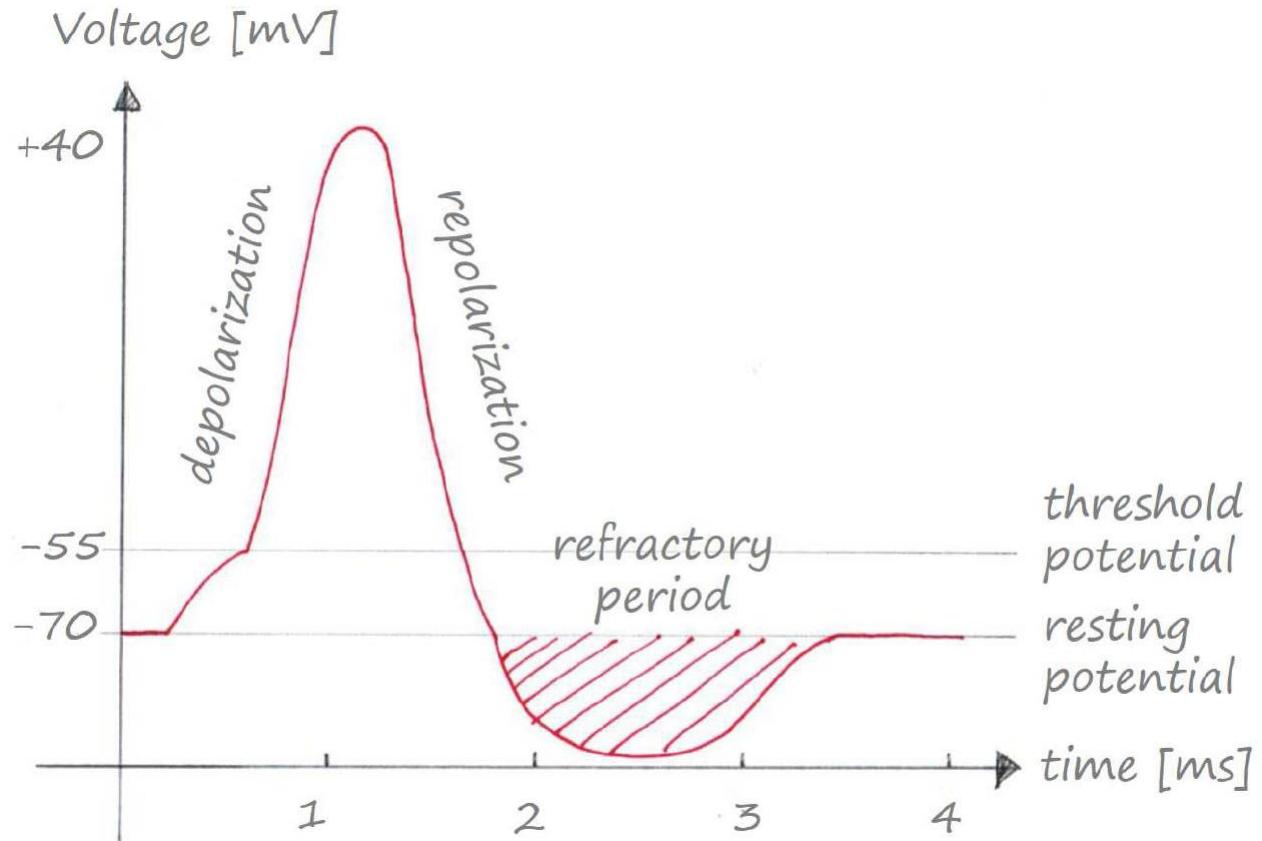
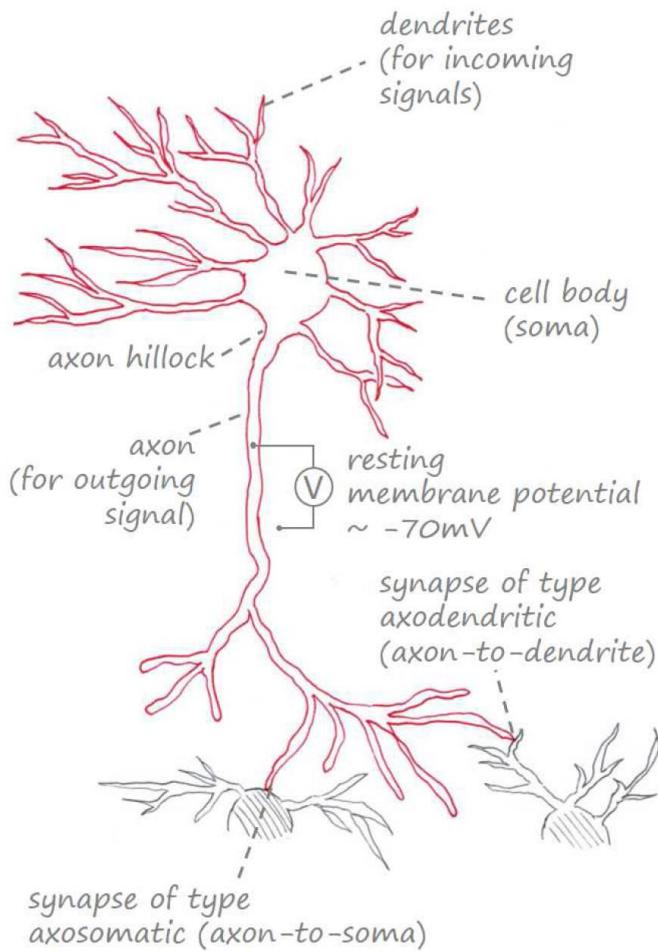


Biological Neuron

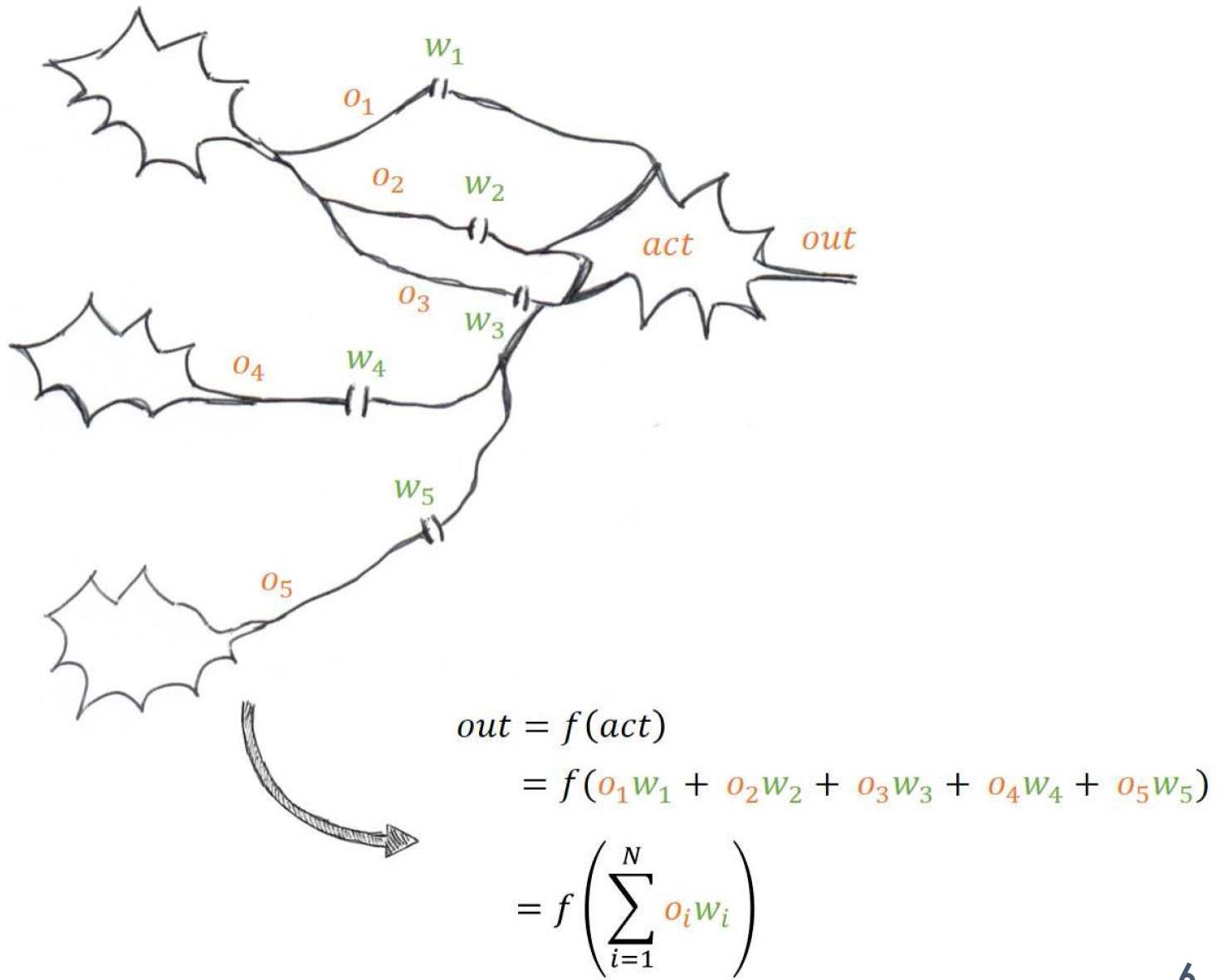
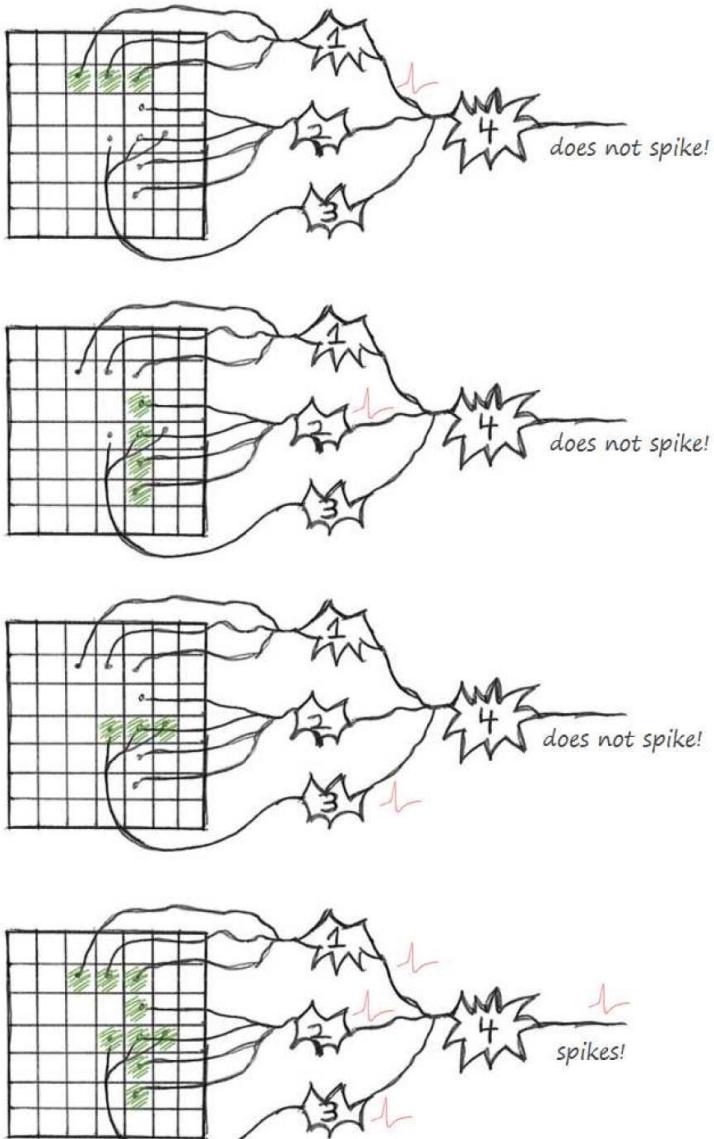


- The basic computational unit of the brain is a **neuron**
→ 86B neurons in the brain
- Neurons are connected with nearly **$10^{14} – 10^{15}$ synapses**
- Neurons receive input signal from **dendrites** and produce output signal along **axon**, which interact with the dendrites of other neurons via **synaptic weights**
- Synaptic weights – learnable & control influence strength

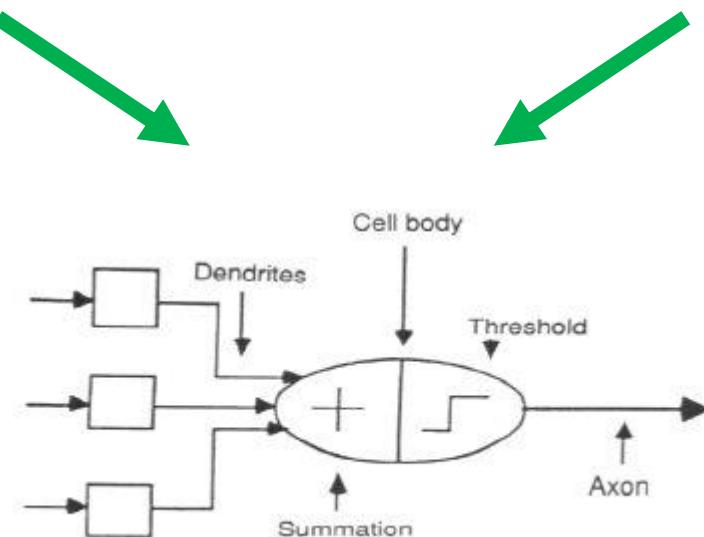
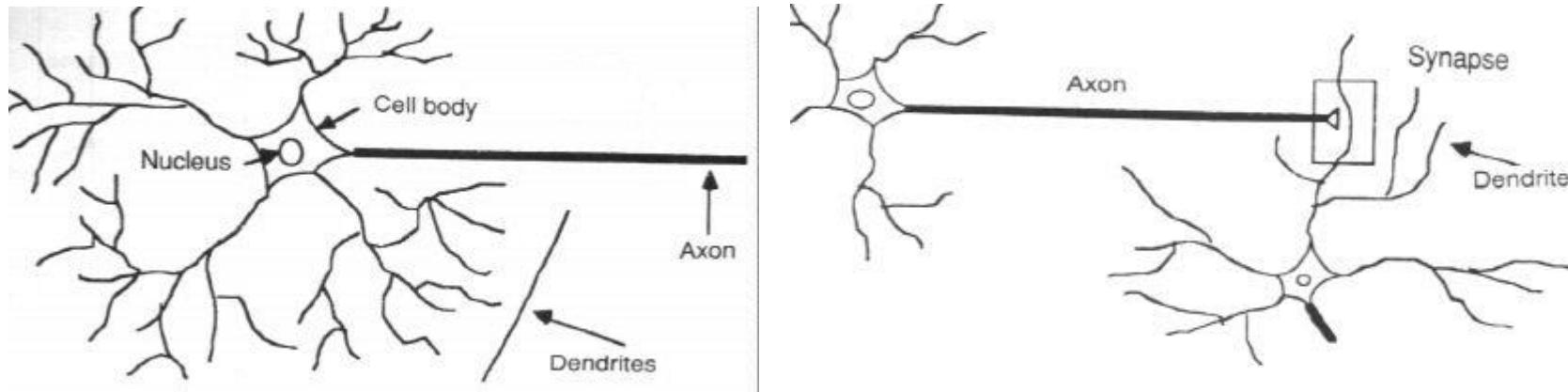
Biological Neuron



Artificial Neural Networks (ANN)



Artificial Neural Networks (ANN)



Artificial Neural Networks (ANN)

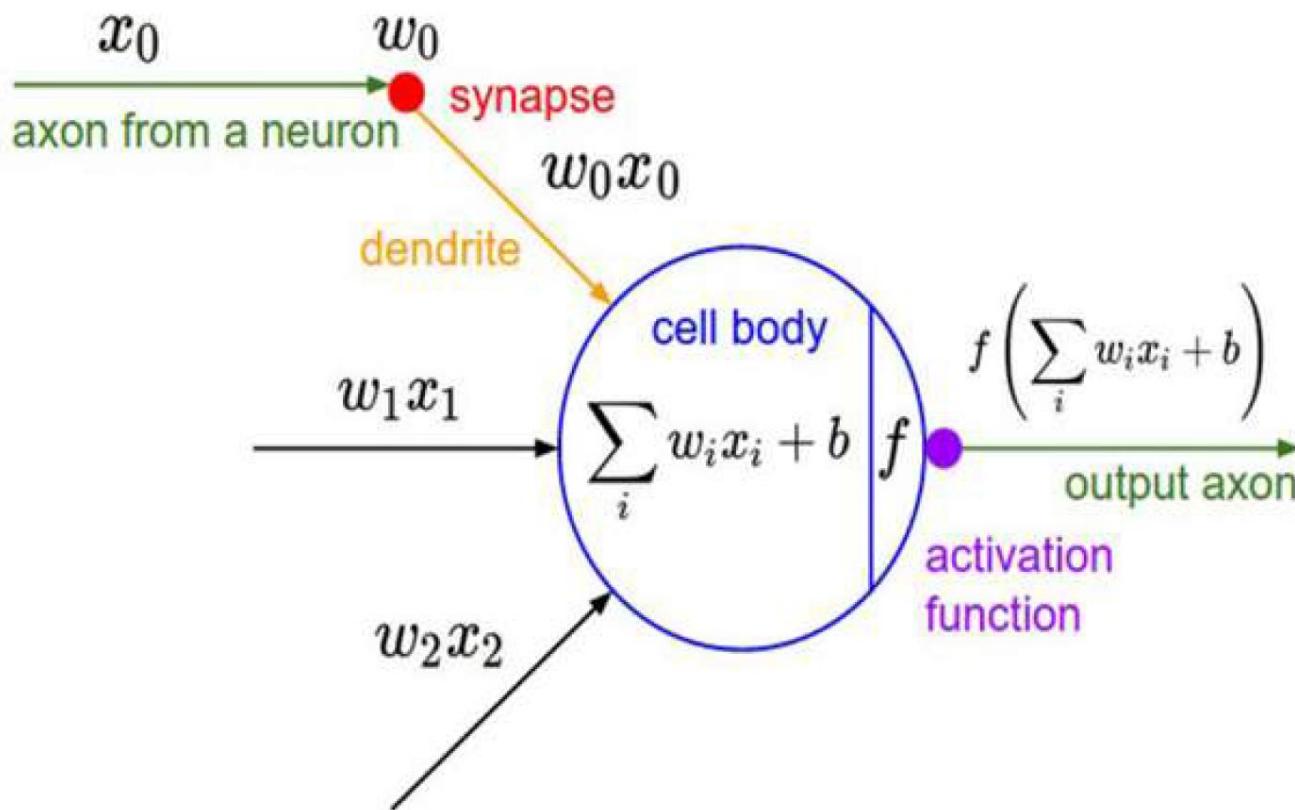
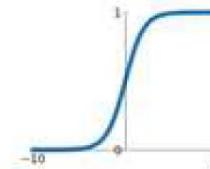


Image Source: Stanford

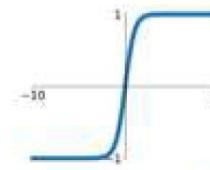
Sigmoid

$$\sigma(x) = \frac{1}{1+e^{-x}}$$



tanh

$$\tanh(x)$$



ReLU

$$\max(0, x)$$



Leaky ReLU

$$\max(0.1x, x)$$



Maxout

$$\max(w_1^T x + b_1, w_2^T x + b_2)$$

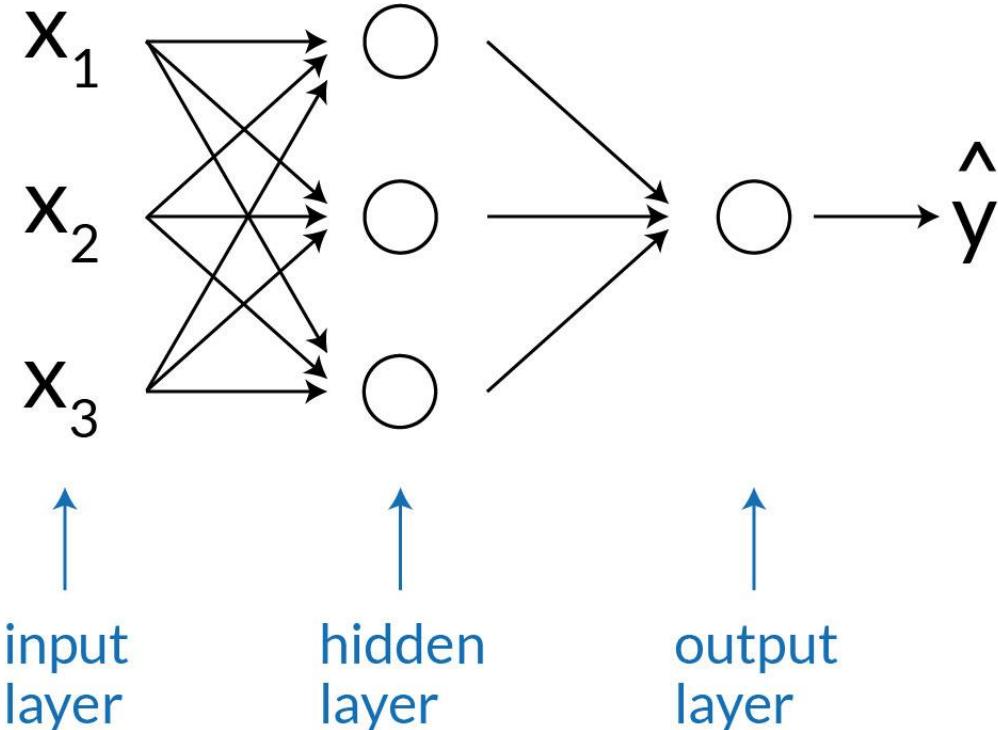
ELU

$$\begin{cases} x & x \geq 0 \\ \alpha(e^x - 1) & x < 0 \end{cases}$$

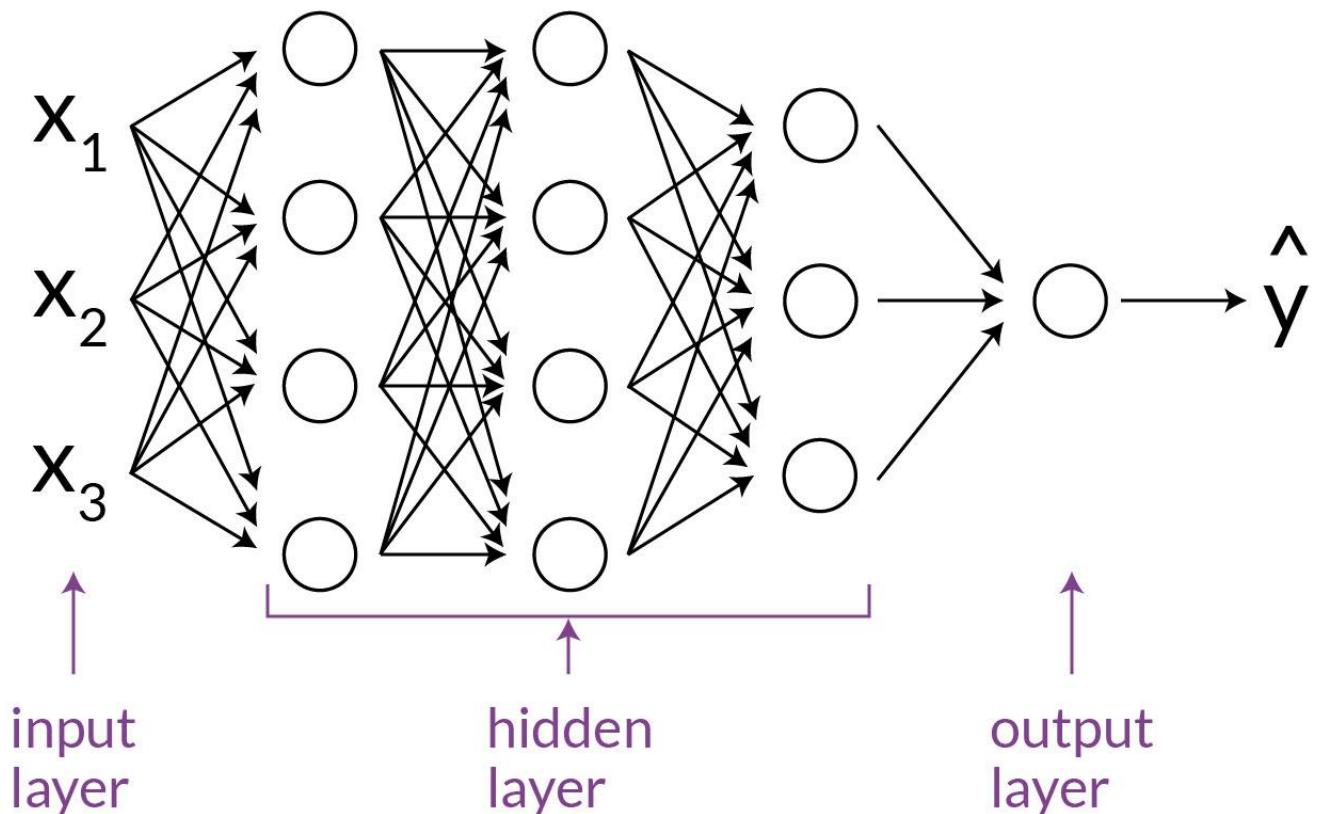


Deep Neural Networks

Shallow Neural Network

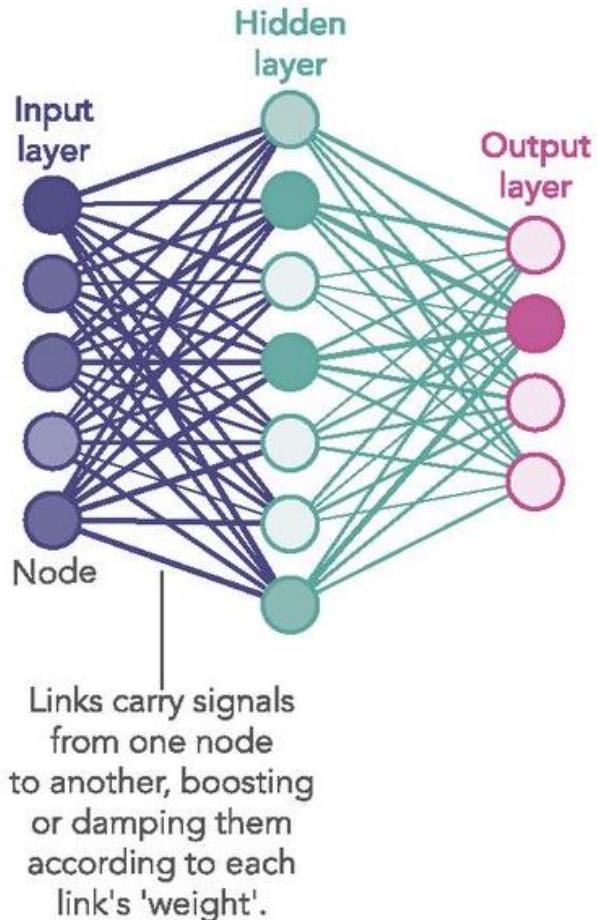


Deep Neural Network

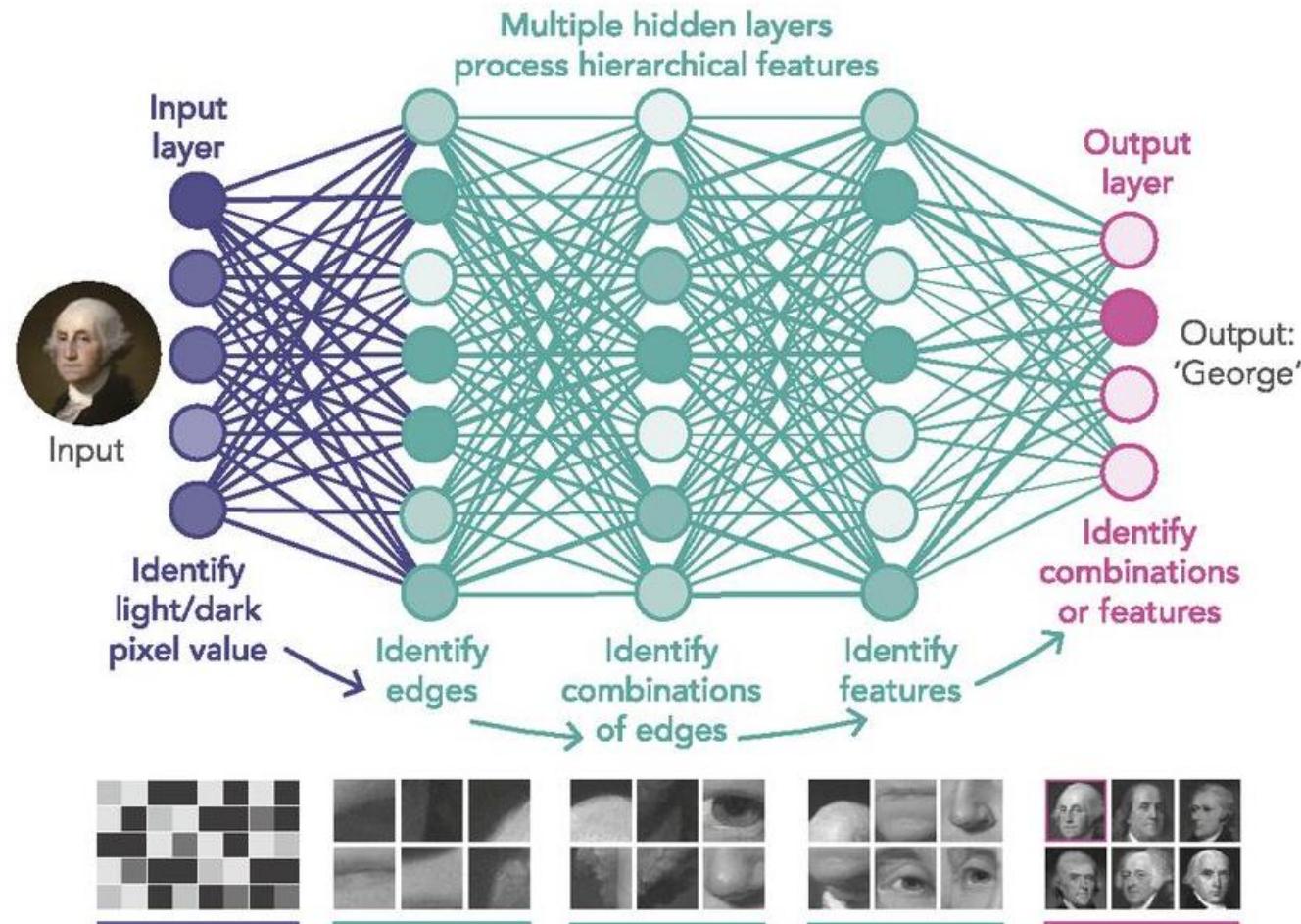


Deep Neural Networks

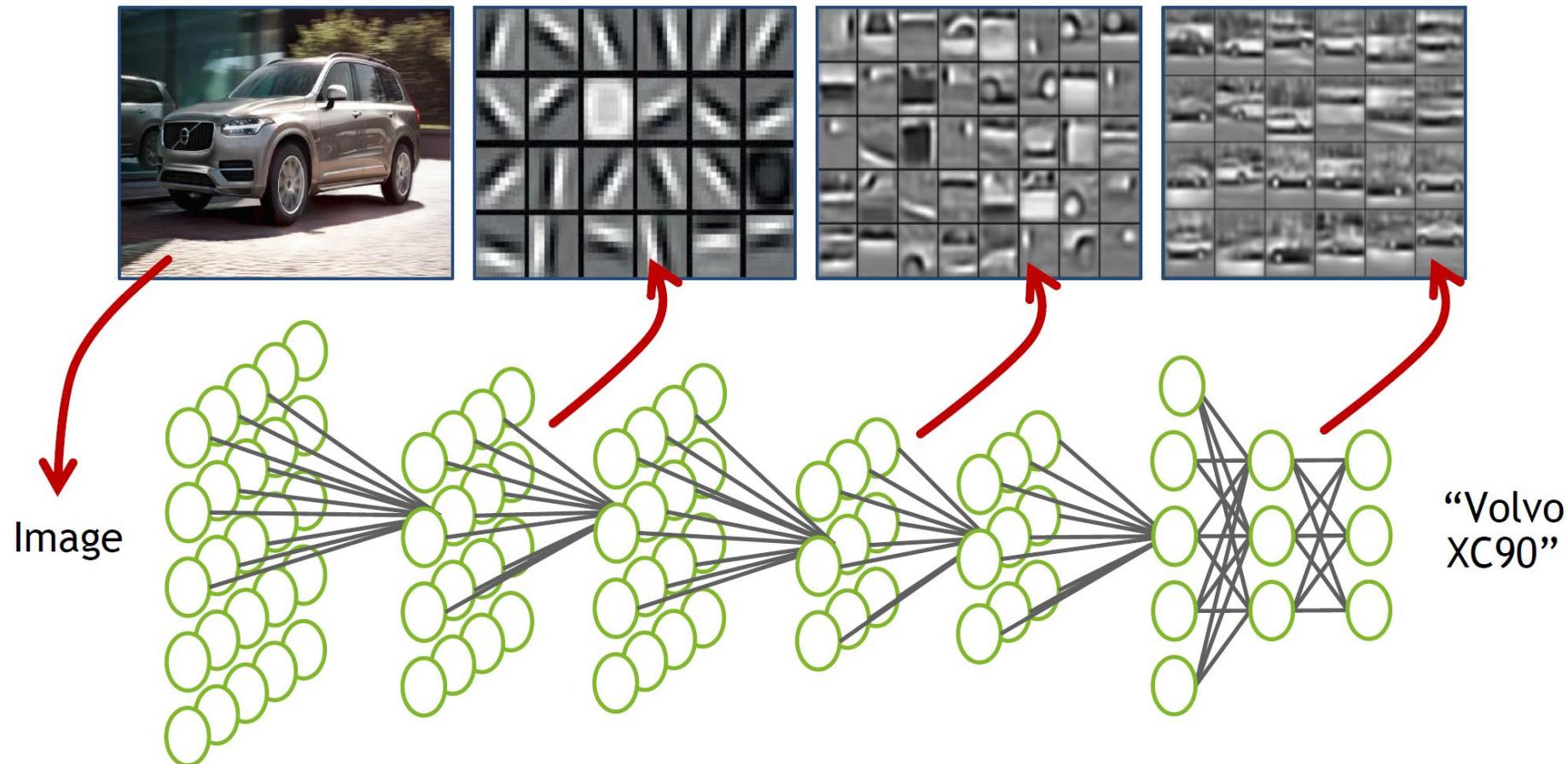
1980S-ERA NEURAL NETWORK



DEEP LEARNING NEURAL NETWORK



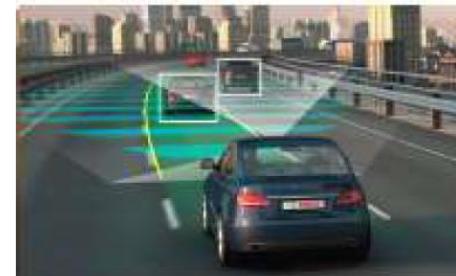
What is Deep Learning?



Deep Learning: Anywhere!



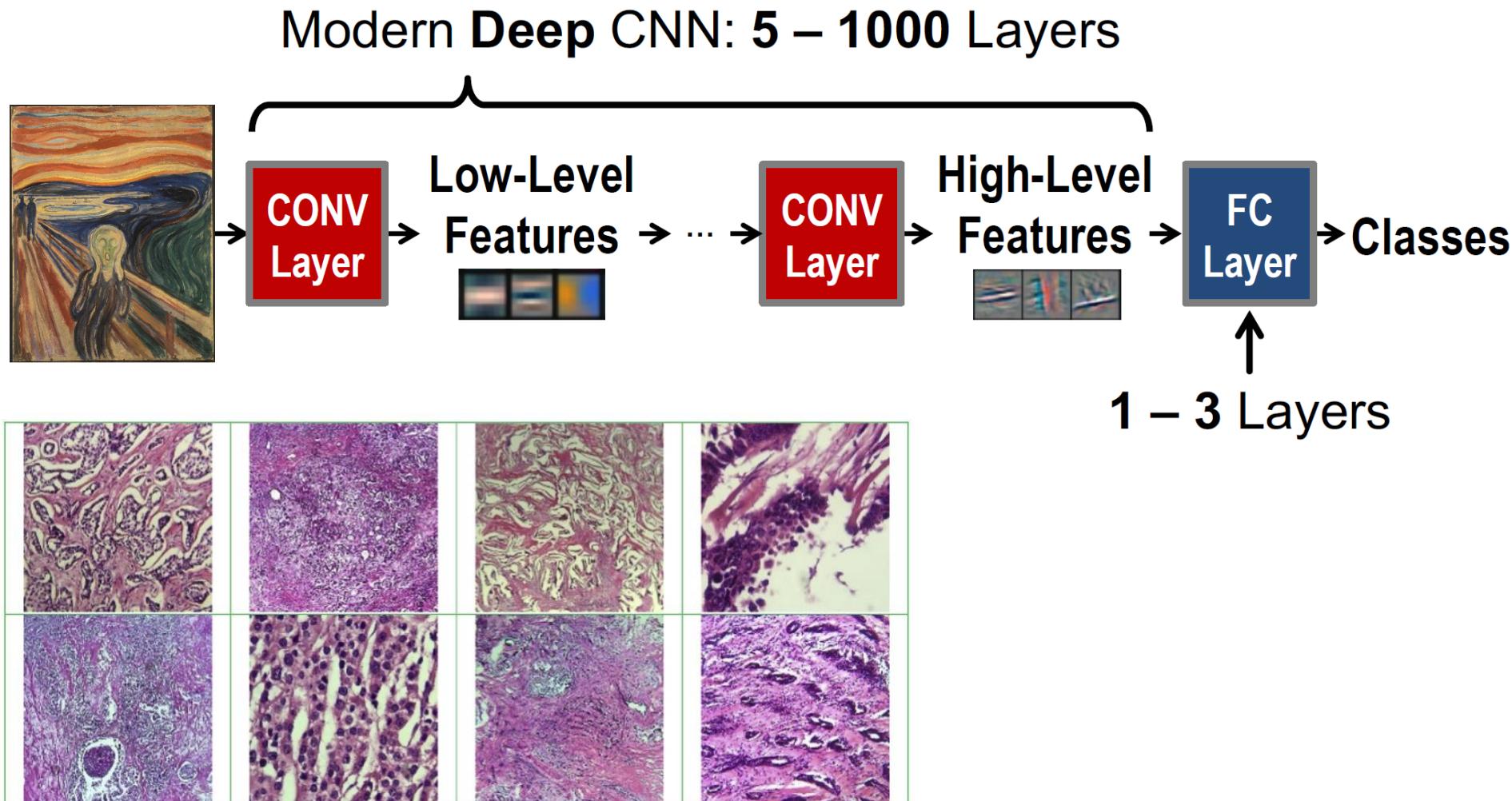
Google
Translate



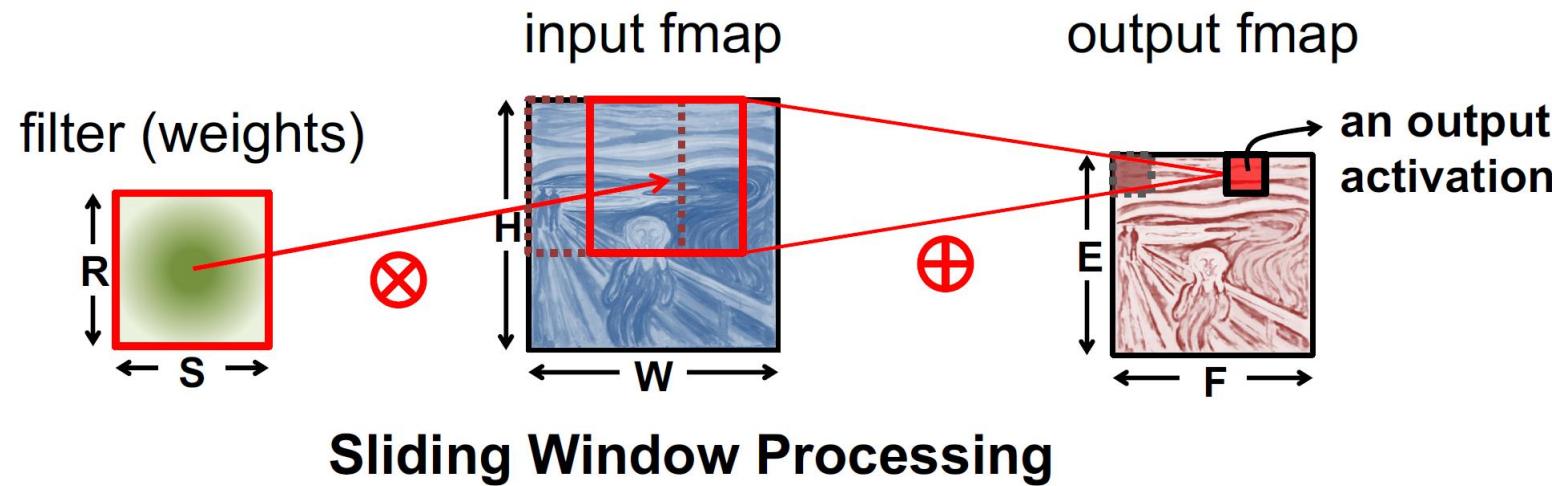
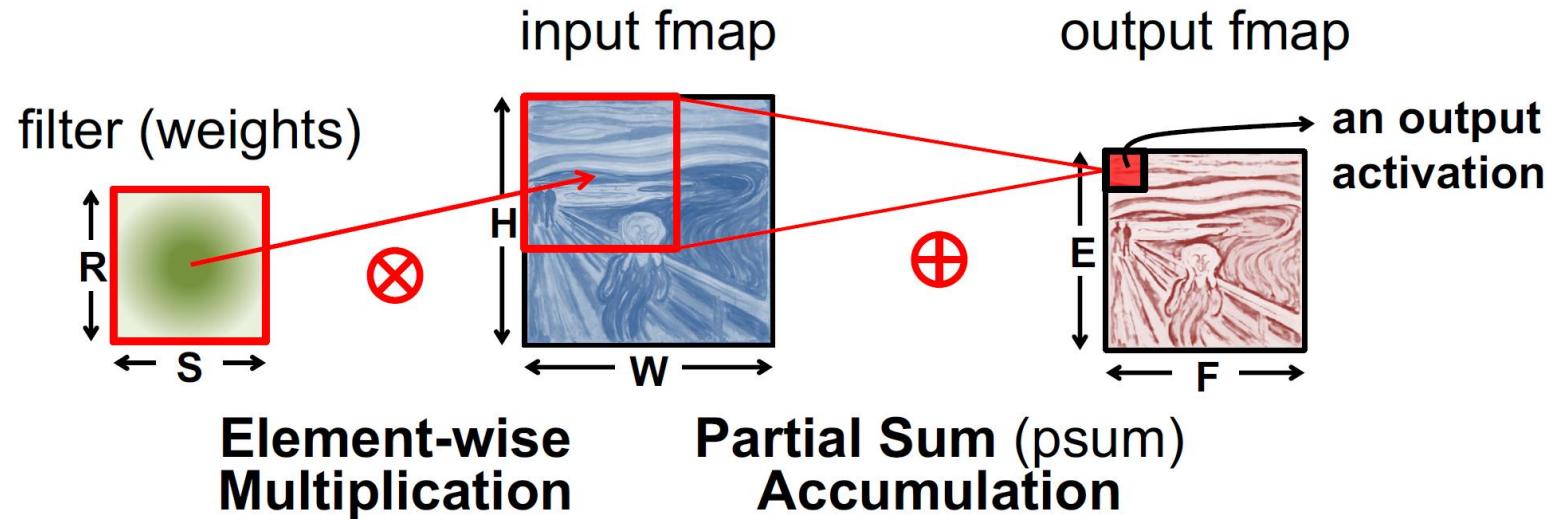
Popular Types of DNNs

- **Fully-Connected NN**
 - feed forward, a.k.a. multilayer perceptron (MLP)
- **Convolutional NN (CNN)**
 - feed forward, sparsely-connected w/ weight sharing
- **Recurrent NN (RNN)**
 - feedback
- **Long Short-Term Memory (LSTM)**
 - feedback + storage

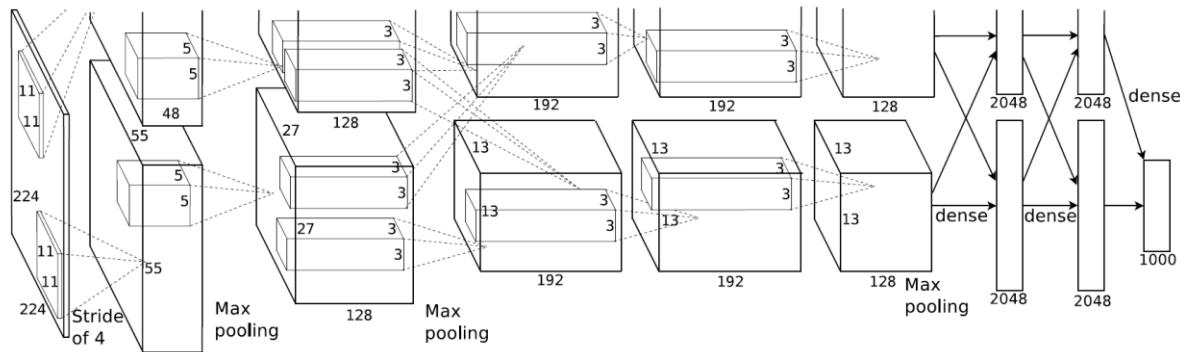
Deep Convolutional Neural Networks



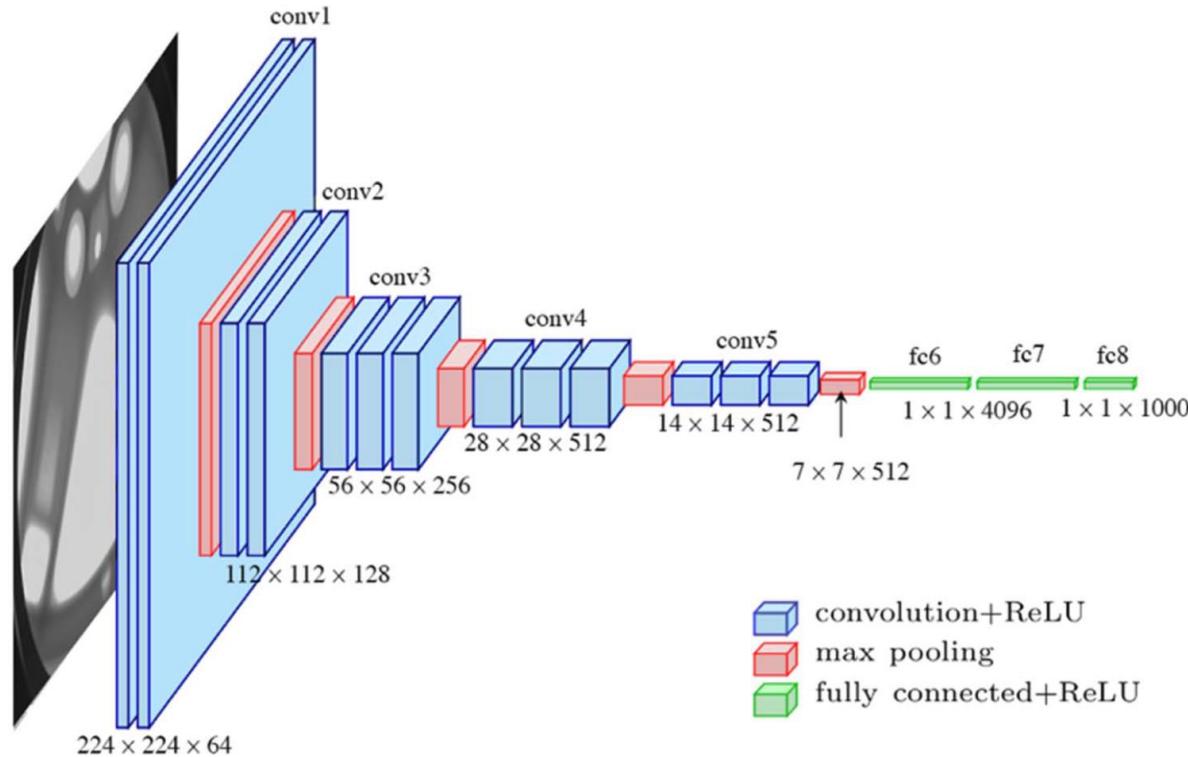
Convolutional Neural Networks (CNNs)



Convolutional Neural Networks (CNNs)



AlexNet



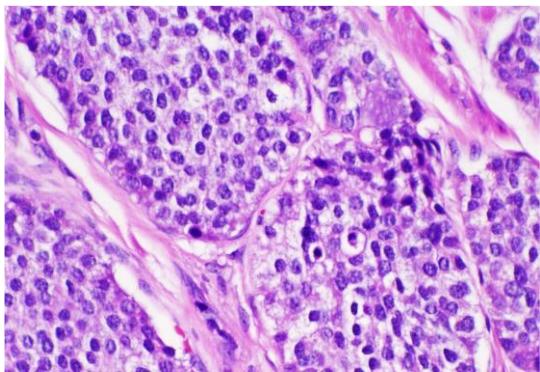
VGG16

Ref. K. Simonyan and A. Zisserman, "Very deep convolutional networks for large-scale image recognition," in

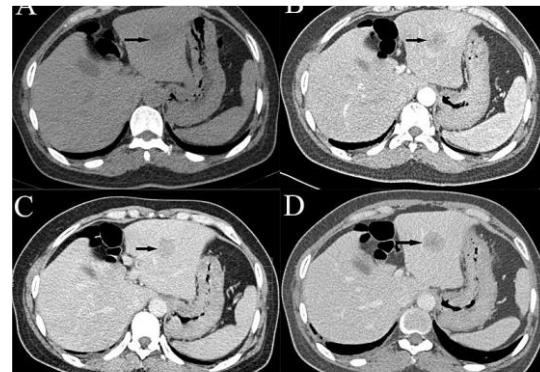
International Conference on Learning Representations (ICLR), 2015.

What does a Deep Neural Network needs to be Trained?

BIG DATA



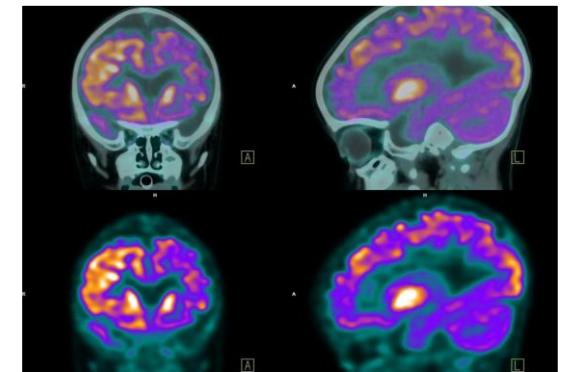
Histopathologic Images



CT-Scan



MRI



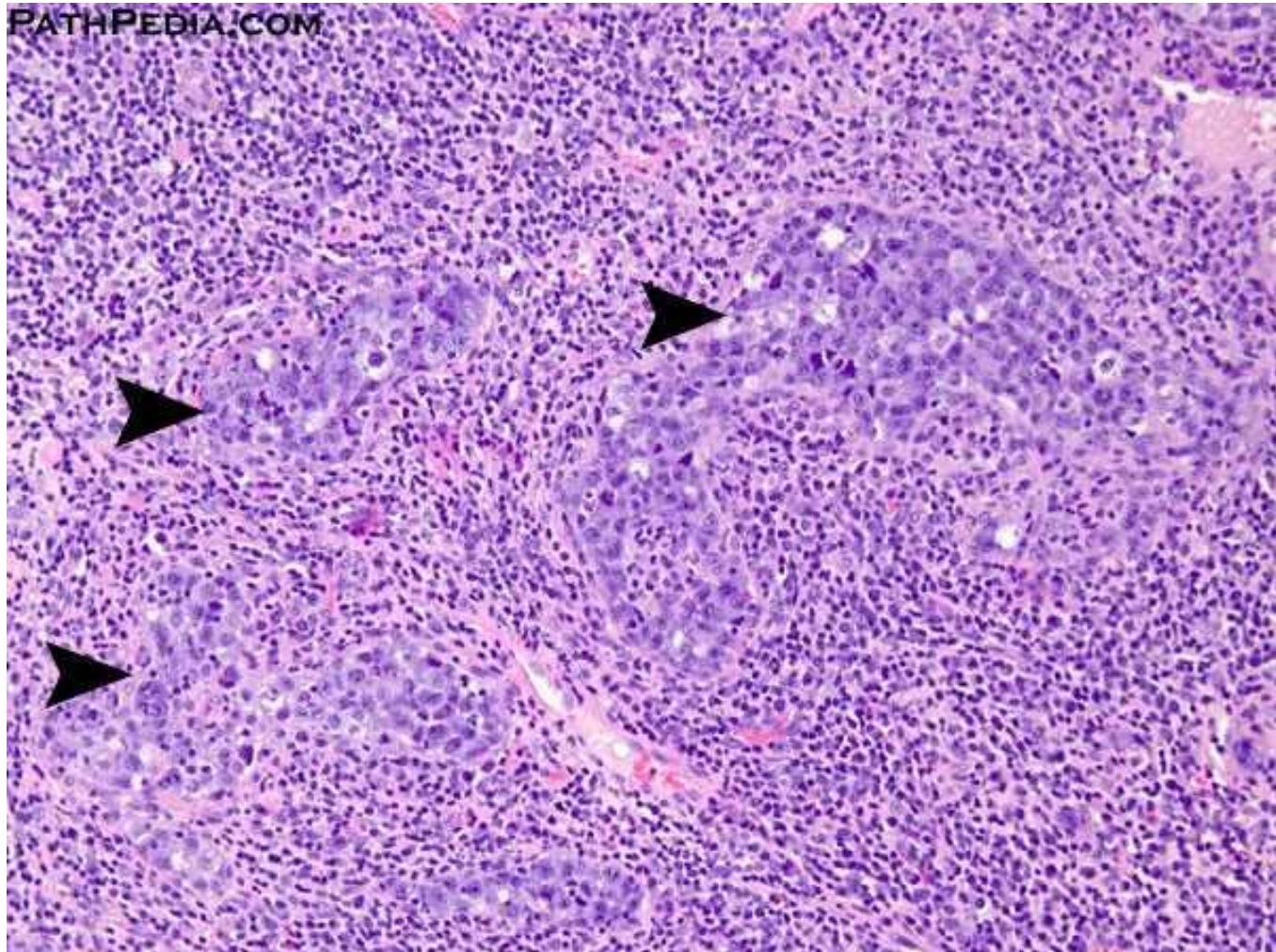
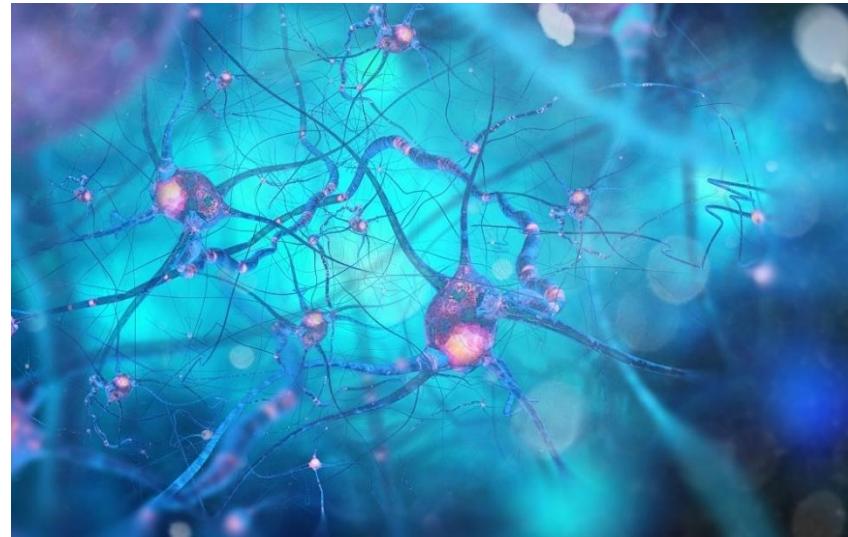
PET-Scan

How much data?
So much that we get our desired results!

Our First Contribution

DataBioX founded in 2020 for sharing biomedical datasets.

DataBioX first dataset: Invasive Ductal Carcinoma Grading Dataset





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Dr. Maryam Tabatabaeian, MD.

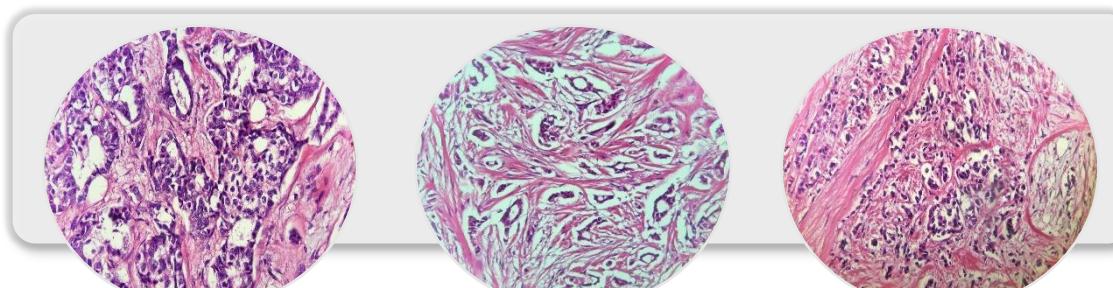
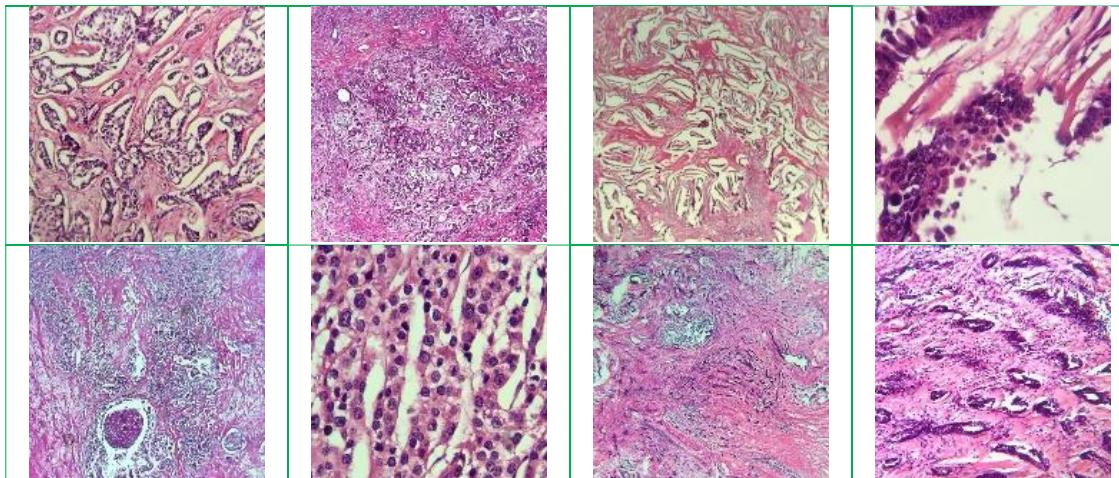
Founder, Anahid Breast Diseases Clinic.

General Surgeon.



Invasive Ductal Carcinoma Grading Dataset

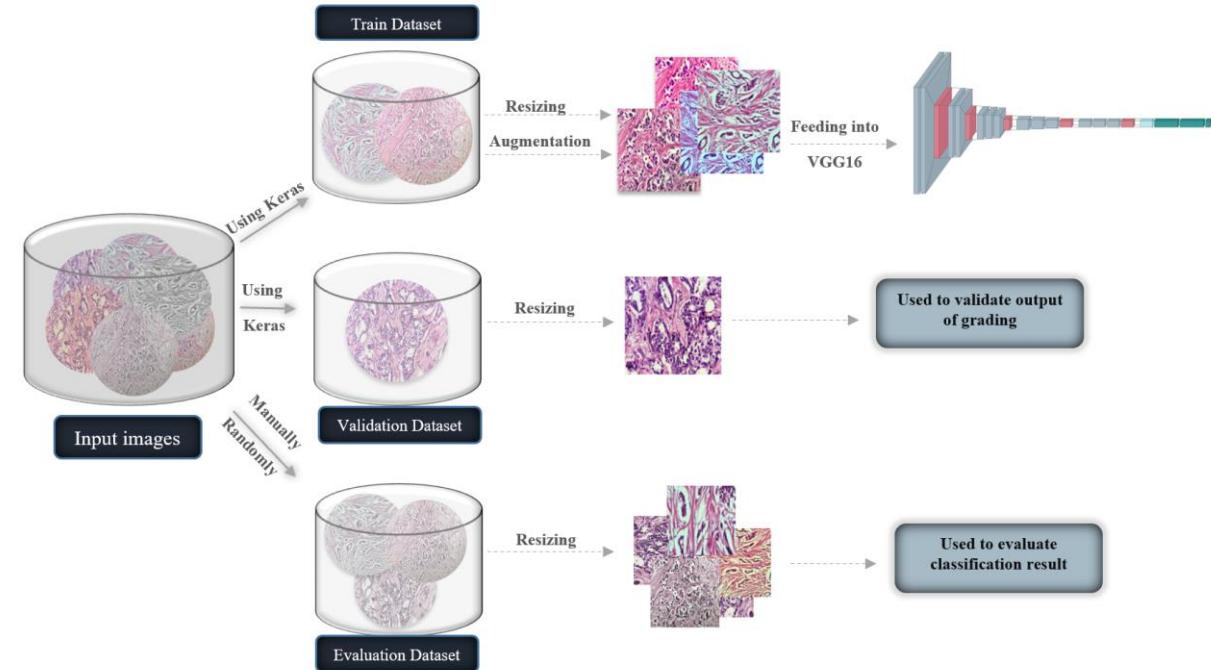
DataBioX: Training dataset for Supervised Learning



Grade I

Grade II

Grade III



Invasive Ductal Carcinoma Grading Dataset

DataBioX: Training dataset for Supervised Learning



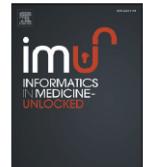
	Numbers	4x	10x	20x	40x
Grade I	37	45	40	43	131
Grade II	43	59	64	63	180
Grade III	44	56	49	49	143
Total	124	160	153	155	454





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A histopathological image dataset for grading breast invasive ductal carcinomas

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ARTICLE INFO

Keywords:
 Breast cancer
 Invasive ductal carcinoma
 Histopathology
 Digital pathology
 Grading
 Image dataset

ABSTRACT

Breast cancer is a common cancer in women, and one of the major causes of death among women around the world. Invasive ductal carcinoma (IDC) is the most widespread type of breast cancer with about 80% of all diagnosed cases. Early accurate diagnosis plays an important role in choosing the right treatment plan and improving survival rate among the patients. In recent years, efforts have been made to predict and detect all types of cancers by employing artificial intelligence. An appropriate dataset is the first essential step to achieve such a goal. This paper introduces a histopathological microscopy image dataset of 922 images related to 124 patients with IDC. The dataset has been published and is accessible through the web at: <http://databiox.com>. The distinctive feature of this dataset as compared to similar ones is that it contains an equal number of specimens from each of three grades of IDC, which leads to approximately 50 specimens for each grade.

1. Introduction

Cancer is a serious public health issue worldwide and the second leading cause of death in the United States [1]. According to the International Agency for Research on Cancer (IARC), about 18.1 million new cases and 9.6 million deaths caused by cancer were reported in 2018 [2].

As shown in Fig. 1, breast cancer is a common cancer and one of the major causes of death worldwide with 627,000 deaths among 2.1 million diagnosed cases in 2018 [2–6].

About 80% of all diagnosed breast cancers are invasive ductal carcinoma (IDC) which makes it the most widespread phenotypic subtype of all breast cancers [7]. It grows in milk ducts and invades fibrous tissue of the breast outside the duct. Pathologists recognize the cancer type and grade through visual investigation of tissue stained by hematoxylin and eosin (H&E) [17] as shown in Fig. 2. Accurate identification of IDC grade is essential in choosing the proper treatment plan and the outcome of the patient accordingly.

Histologic grade is a prognostic factor and an indicator of response to chemotherapy. It has been shown to be closely related to the frequency of recurrence and death due to IDC, and to disease-free interval and to longer life after mastectomy.

Several studies show that patients with a high grade IDC, and

mastectomy treated, had a remarkably higher frequency of auxiliary lymph node (ALN) and mortality rate than the patients with lower grade. Histologic grade has been shown to be significantly related not only to recurrence and death due to invasive ductal carcinoma, but also to the disease-free interval and overall length of survival after mastectomy regardless of clinical stage. High-grade carcinomas result in early treatment failures, whereas later recurrences are more often observed among low-grade tumors [8,9].

In the recent years, efforts have been made to predict and detect all types of cancers, employing artificial intelligence and its sub categories like machine learning and deep learning [10–15]. The appropriate dataset is the first essential step to achieve this goal. There are some available datasets for IDC [11–13] but our findings indicate that there is no published dataset for histologic grading of IDC. Therefore, this research aimed at providing a well-organized histopathological microscopy image dataset for grading IDC. The images have been taken from breast tissues stained with H&E and are labeled based on their grade and magnification level.

This paper has been organized as follows: Related works are presented in section 2. Section 3 covers the main contribution of this research, i.e. the prepared image dataset: databiox. Methods of collecting images and doing the statistics are presented in this section.

Published Paper in 2020

Elsevier Publication

Officially requested by many prestigious Universities around the world like MIT and University of Johns Hopkins.



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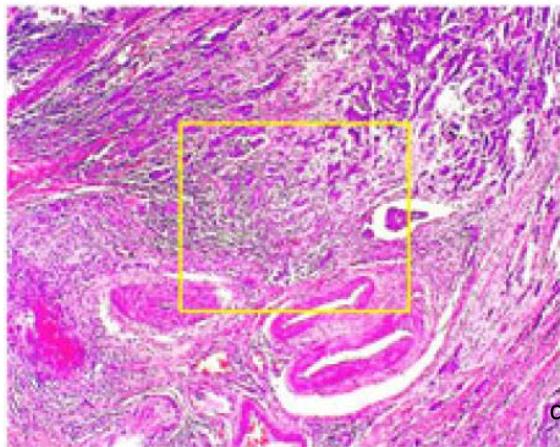
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Required Dataset

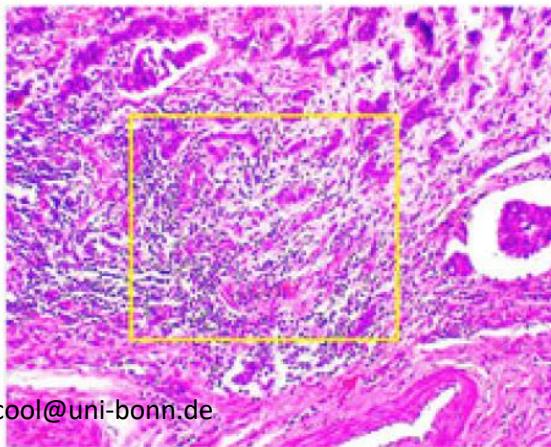
Breast Cancer Grading Dataset

Breast Cancer Classification Dataset

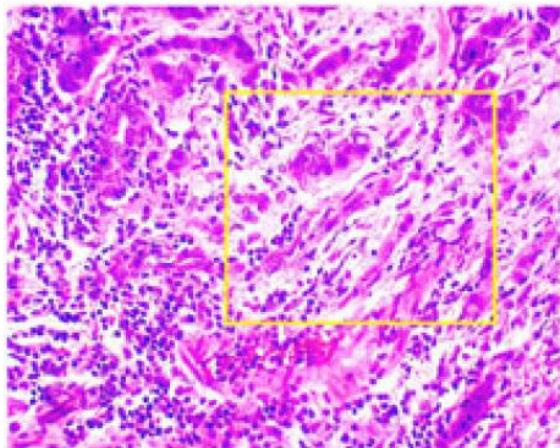
BreaKHis: Training dataset for Supervised Learning



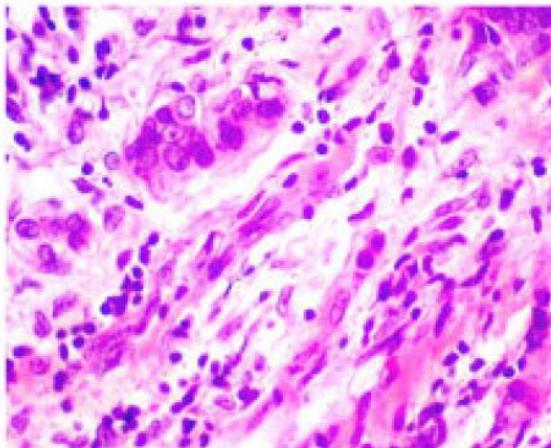
(a)



(b)



(c)



(d)

Magnification	Benign	Malignant	Total
40×	625	1370	1995
100×	644	1437	2081
200×	623	1390	2013
400×	588	1232	1820
Total	2480	5429	7909
# Patients	24	58	82

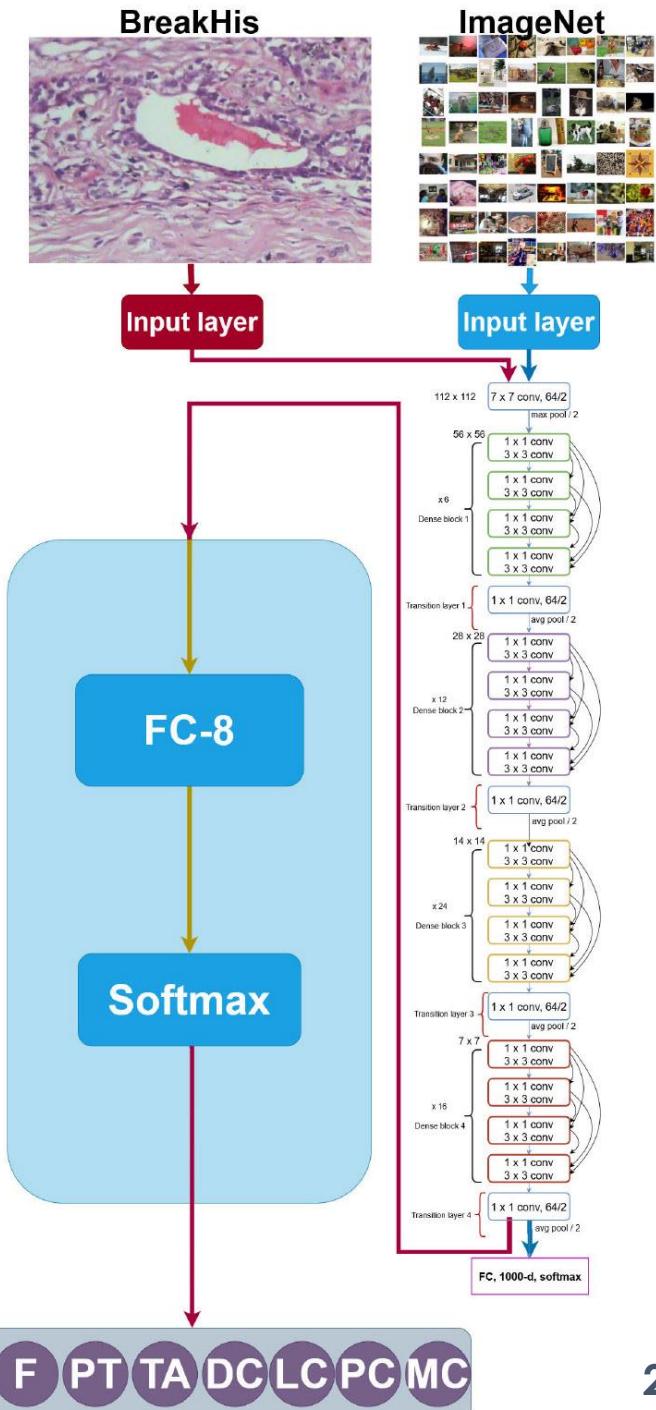
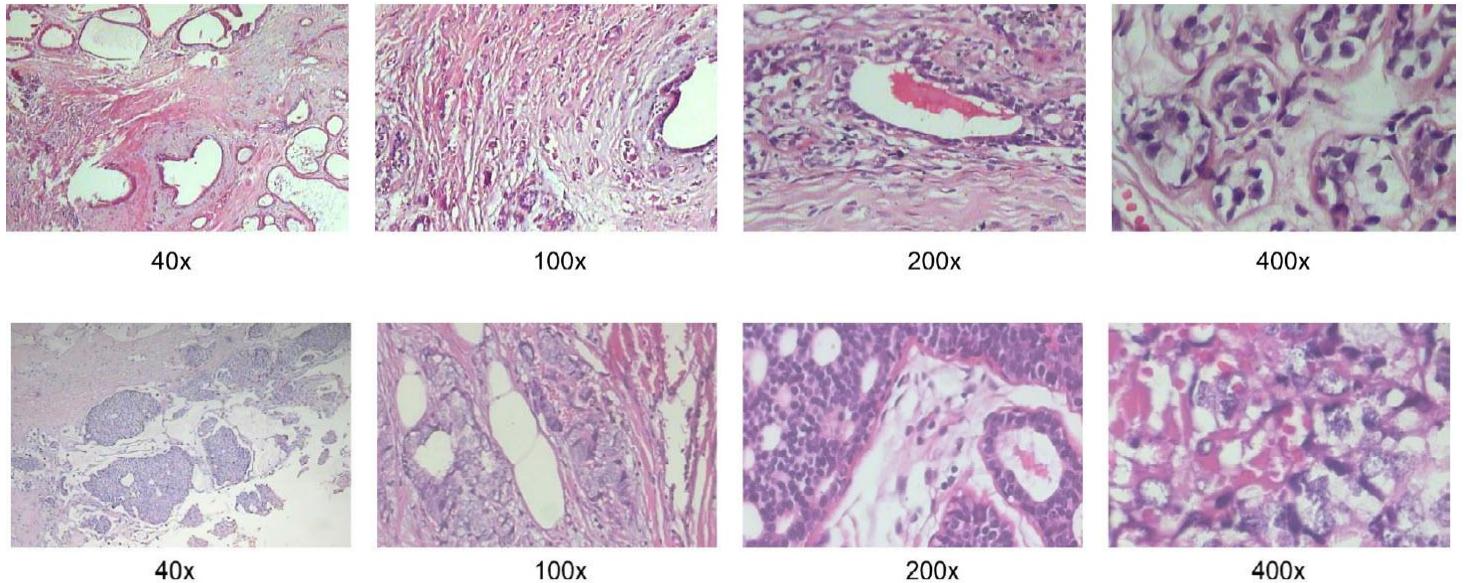
Magnification	A	F	TA	PT	Total
40×	114	253	109	149	598
100×	113	260	121	150	614
200×	111	264	108	140	594
400×	106	237	115	130	562
Total	444	1014	453	569	2368
# Patients	4	10	3	7	24

Breast Cancer Detection

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Deep Learning Applied for Histological Diagnosis of Breast Cancer (2020)

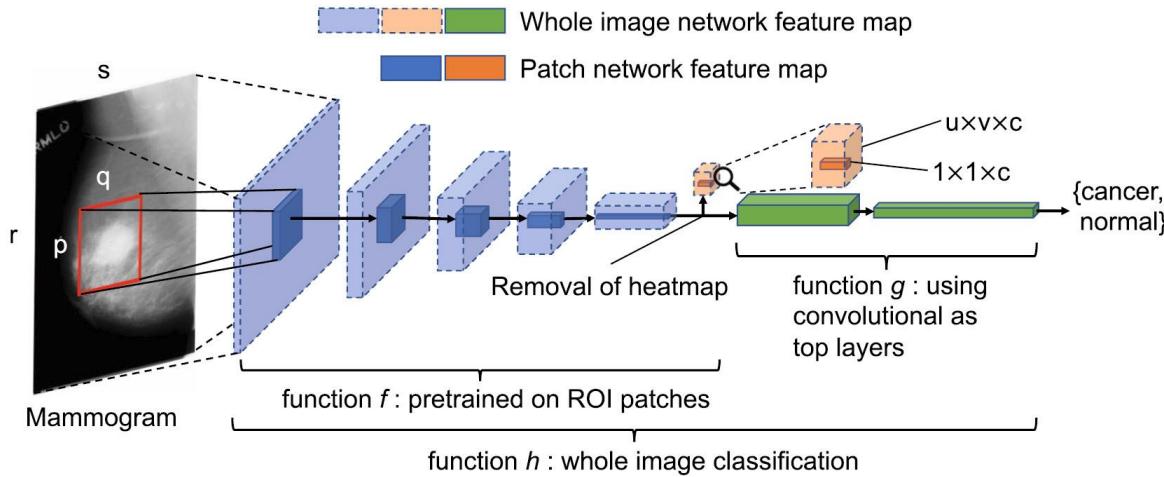


Model details	Model 1	Model 2	Model 3	Model 4	Model 5	Model 6
Transfer learning	-	-	ResNet50	DenseNet121	ResNet50	Densenet121
CNN layers setting	2 setting 3	2 setting 3	40 setting 3	117 setting 3	40 setting 4	117 setting 4
Data Augmentation	-	✓	✓	✓	✓	✓
Results						
Misclassified images	20	18	1	0	3	2
Accuracy(ILA)	79.59%	81.63%	98.98%	100%	96.94%	97.97%
Precision	78.20%	82.85%	98.38%	100%	95.31%	96.82%
Recall	95.31%	90.62%	100%	100%	100%	100%
F1 score	85.91%	86.57%	99.18%	100%	97.59%	98.38%

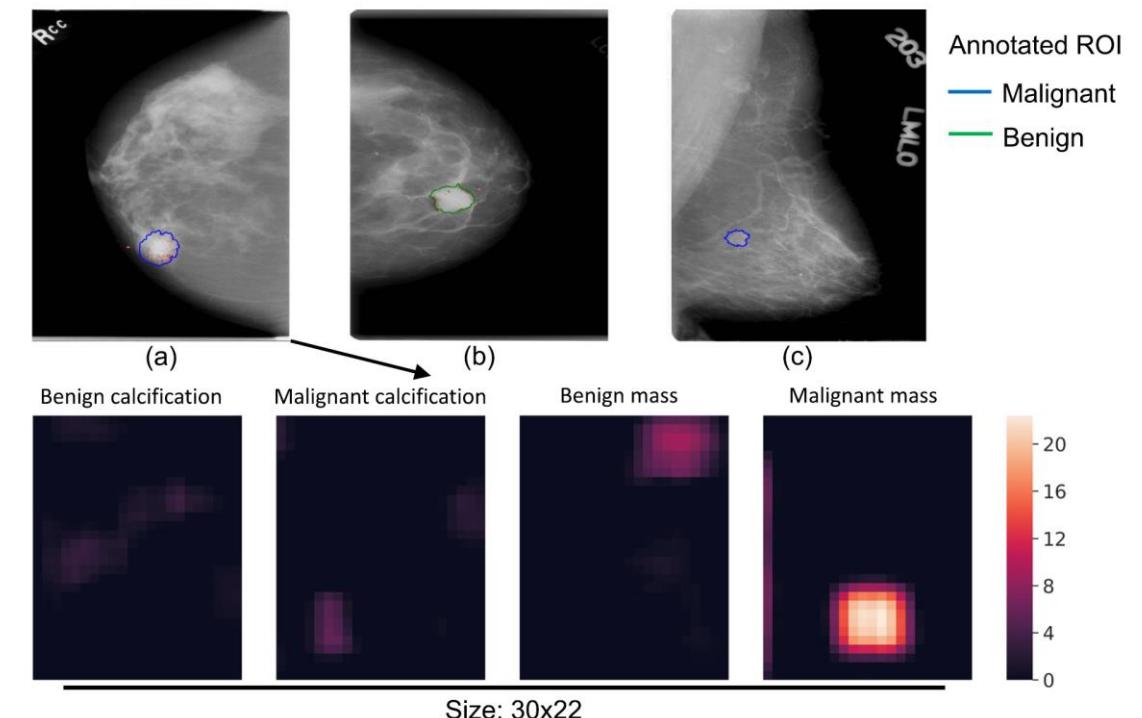
A F PT TA DCLC PCMC

Breast Cancer Detection

Deep Learning to Improve Breast Cancer Detection on Screening Mammography (2019)



#Patients	#Images	Resnet-Resnet	Resnet-VGG	VGG-VGG	VGG-Resnet
20	79	0.92	0.88	0.87	0.89
30	117	0.93	0.94	0.93	0.90
40	159	0.93	0.95	0.93	0.93
50	199	0.94	0.95	0.94	0.93
60	239	0.95	0.95	0.95	0.94
72 (All)	280	0.95	0.95	0.95	0.95

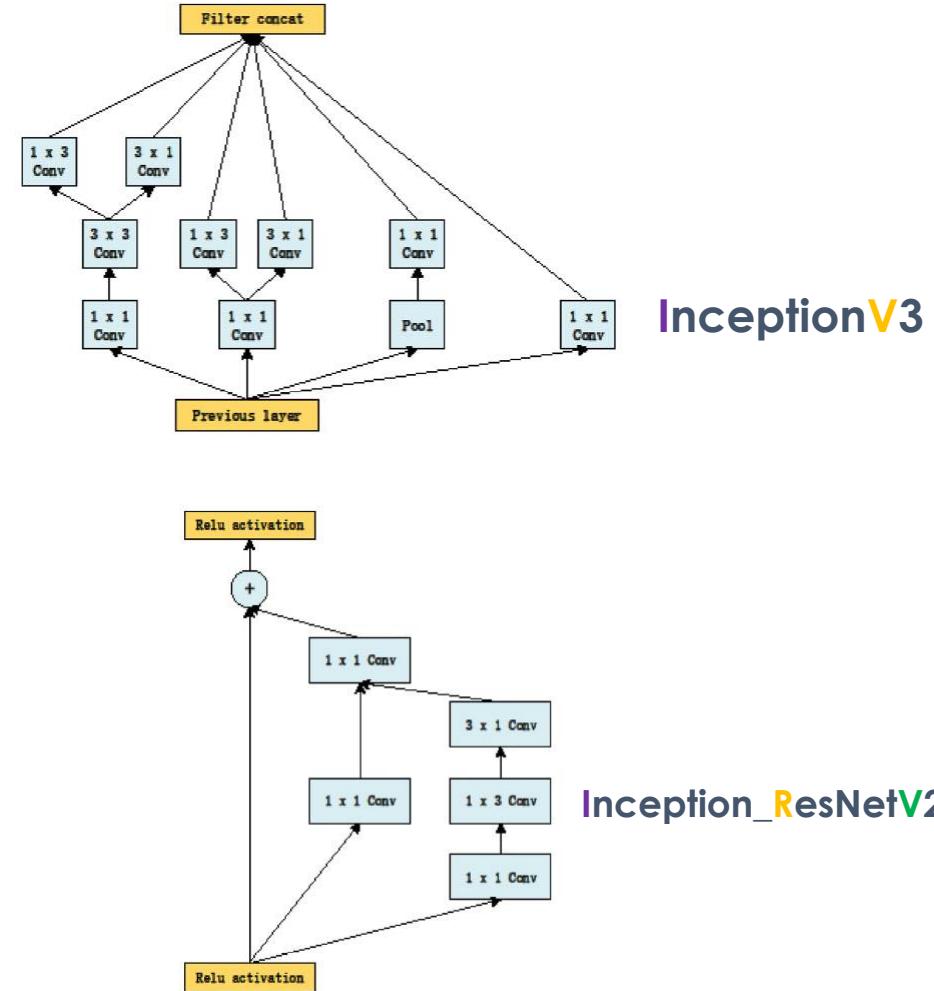


Breast Cancer Classification

Deep Learning Based Analysis of Histopathological Images of Breast Cancer (2019)



Classification	Network	Criteria	Magnification factors			
			40X	100X	200X	400X
Binary	INV3	Se	98.00	98.48	99.01	96.41
		Sp	94.31	<u>93.46</u>	91.40	90.99
		PPV	97.41	<u>96.67</u>	95.88	95.89
		DOR	81,233	92,303	106,700	27,105
		ACC_IL	96.84	96.76	96.49	94.71
		ACC_PL	97.74	94.19	<u>87.21</u>	<u>96.67</u>
		F1	97.70	97.56	97.42	96.15
		AUC	99.47	<u>99.03</u>	99.29	97.91
		Kappa	92.64	92.74	91.95	87.68
IRV2	IRV2	Se	<u>98.48</u>	<u>98.90</u>	99.13	98.06
		Sp	96.63	92.95	<u>92.80</u>	92.10
		PPV	98.46	96.45	<u>96.39</u>	96.51
		DOR	185,774	<u>118,782</u>	147,138	58,835
		ACC_IL	97.90	<u>96.88</u>	<u>96.98</u>	<u>96.98</u>
		ACC_PL	98.03	<u>97.07</u>	82.74	88.12
		F1	98.47	<u>97.66</u>	<u>97.74</u>	<u>97.28</u>
		AUC	99.57	98.84	99.61	98.81
		Kappa	95.12	<u>92.96</u>	93.18	91.05
Multi-class	INV3	ACC_IL	90.28	85.35	83.99	82.08
		ACC_PL	90.44	<u>89.05</u>	80.63	<u>81.08</u>
		Macro-F1	88.55	82.59	79.64	77.98
		Micro-F1	90.28	85.35	83.99	82.08
		Kappa	87.37	80.26	77.91	76.39
	IRV2	ACC_IL	92.07	<u>88.06</u>	<u>87.62</u>	84.50
		ACC_PL	89.11	88.45	<u>86.07</u>	71.42
		Macro-F1	90.89	<u>85.67</u>	<u>84.08</u>	<u>80.13</u>
		Micro-F1	92.07	<u>88.06</u>	<u>87.62</u>	84.50
		Kappa	89.74	<u>84.03</u>	<u>82.84</u>	79.70

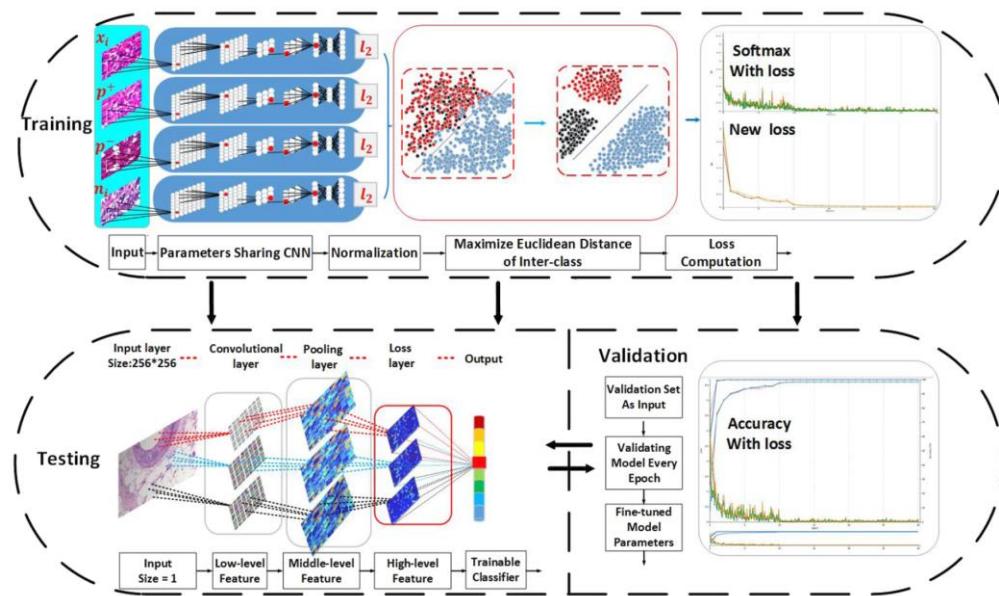


Breast Cancer Classification

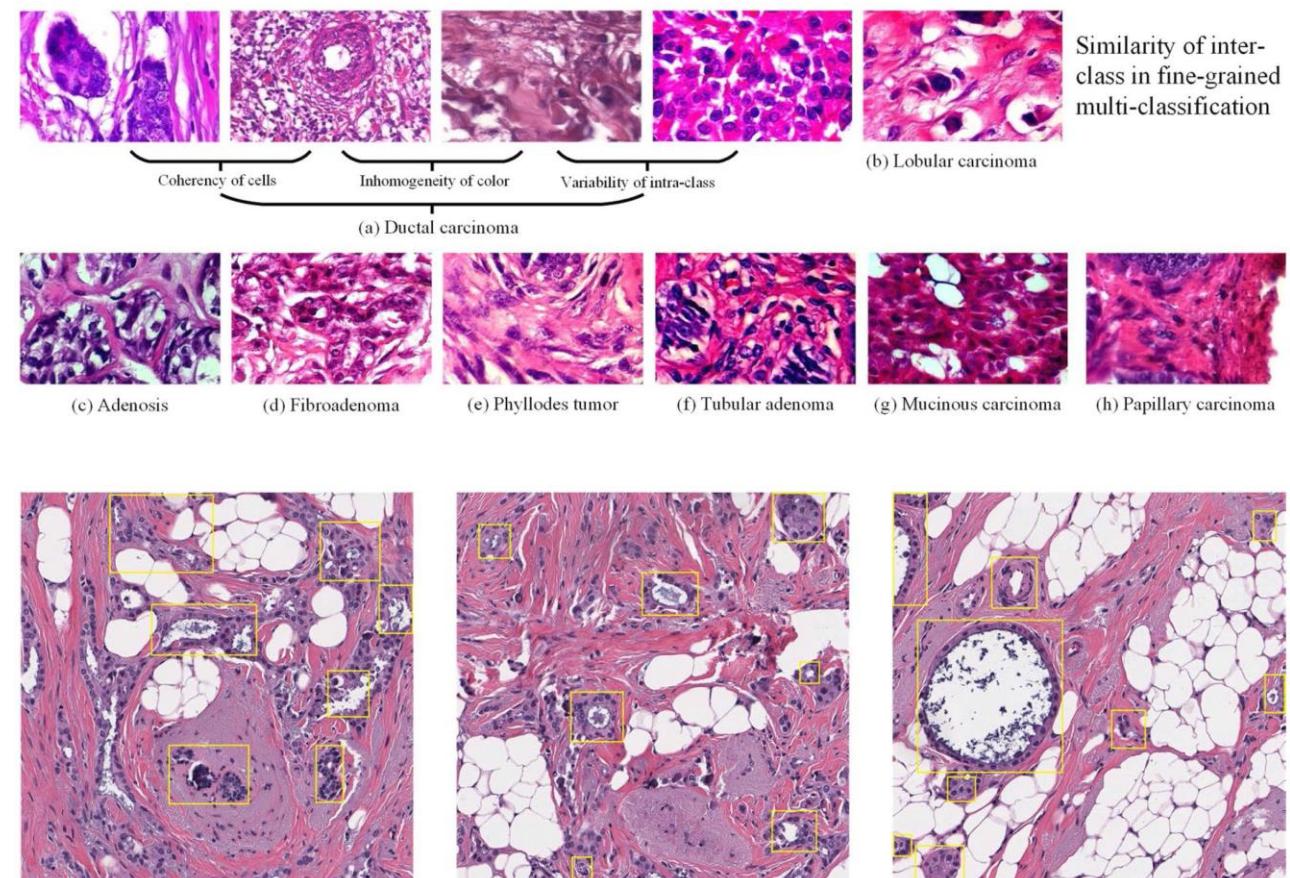
Breast Cancer Multi-classification from Histopathological Images
with Structured Deep Learning Model (2017)

nature

SCIENTIFIC
REPORTS



Accuracy at	Methods	Magnification factors			
		40X	100X	200X	400X
Image level	AlexNet ¹⁷	85.6 ± 4.8	83.5 ± 3.9	83.1 ± 1.9	80.8 ± 3.0
	CSDCNN	95.8 ± 3.1	96.9 ± 1.9	96.7 ± 2.0	94.9 ± 2.8
Patient level	PFTAS + QDA ¹²	83.8 ± 4.1	82.1 ± 4.9	84.2 ± 4.1	82.0 ± 5.9
	PFTAS + SVM ¹²	81.6 ± 3.0	79.9 ± 5.4	85.1 ± 3.1	82.3 ± 3.8
	GLCM + 1-NN ¹²	74.7 ± 1.0	76.8 ± 2.1	83.4 ± 3.3	81.7 ± 3.3
	PFTAS + RF ¹²	81.8 ± 2.0	81.3 ± 2.8	83.5 ± 2.3	81.0 ± 3.8
	AlexNet ¹⁷	90.0 ± 6.7	88.4 ± 4.8	84.6 ± 4.2	86.1 ± 6.2
	CSDCNN	97.1 ± 1.5	95.7 ± 2.8	96.5 ± 2.1	95.7 ± 2.2



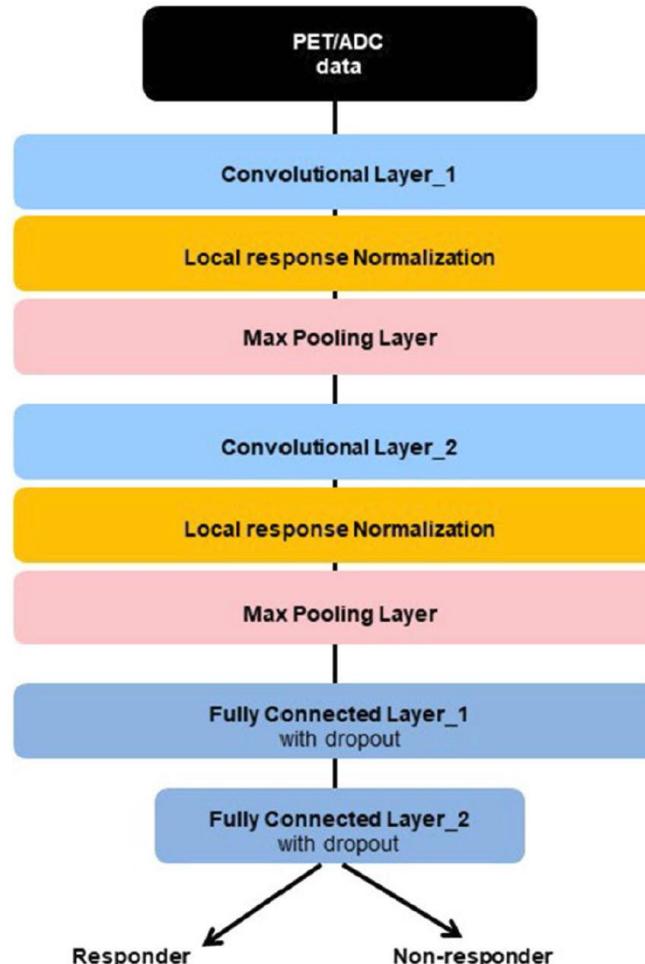
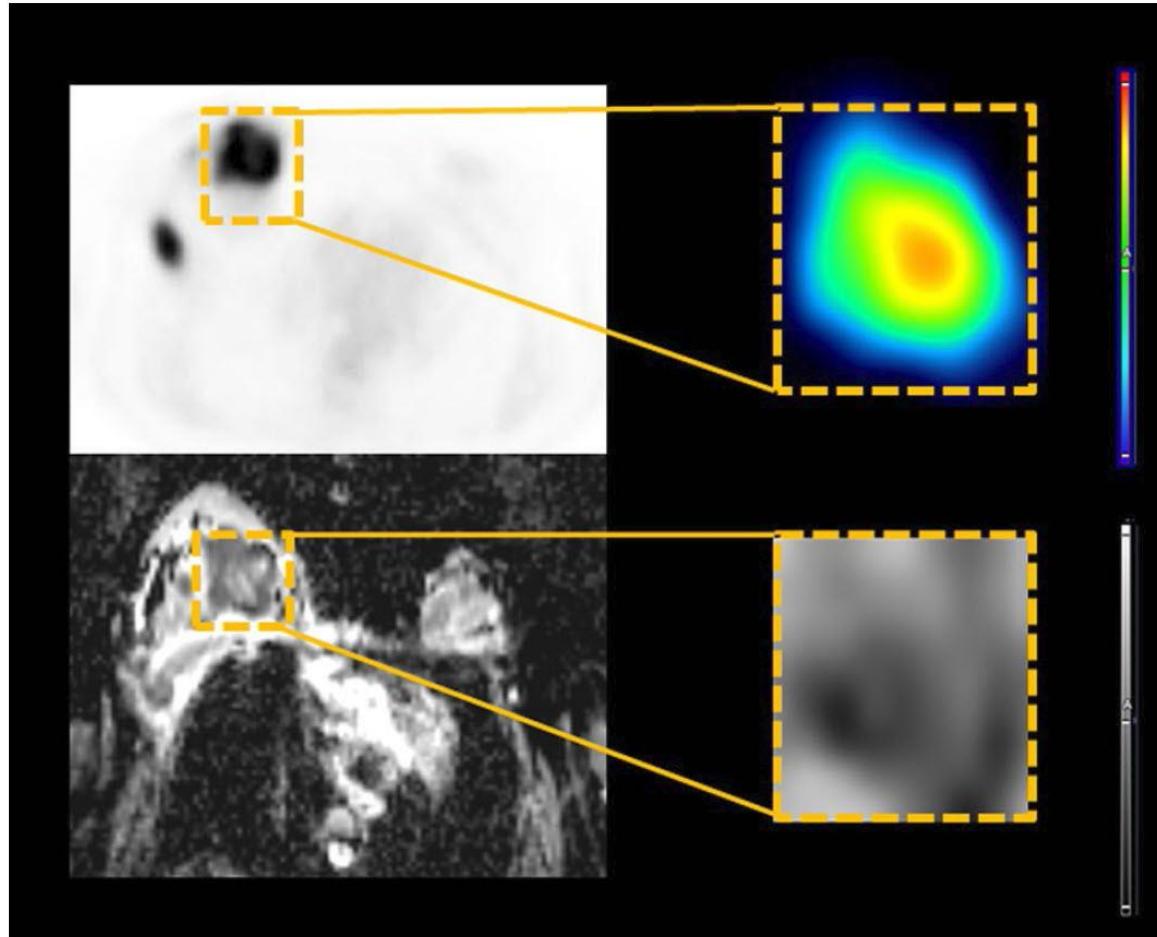
Breast Cancer Treatment Plan

Early prediction of neoadjuvant chemotherapy response for advanced breast cancer

Using PET/MRI image deep learning

nature

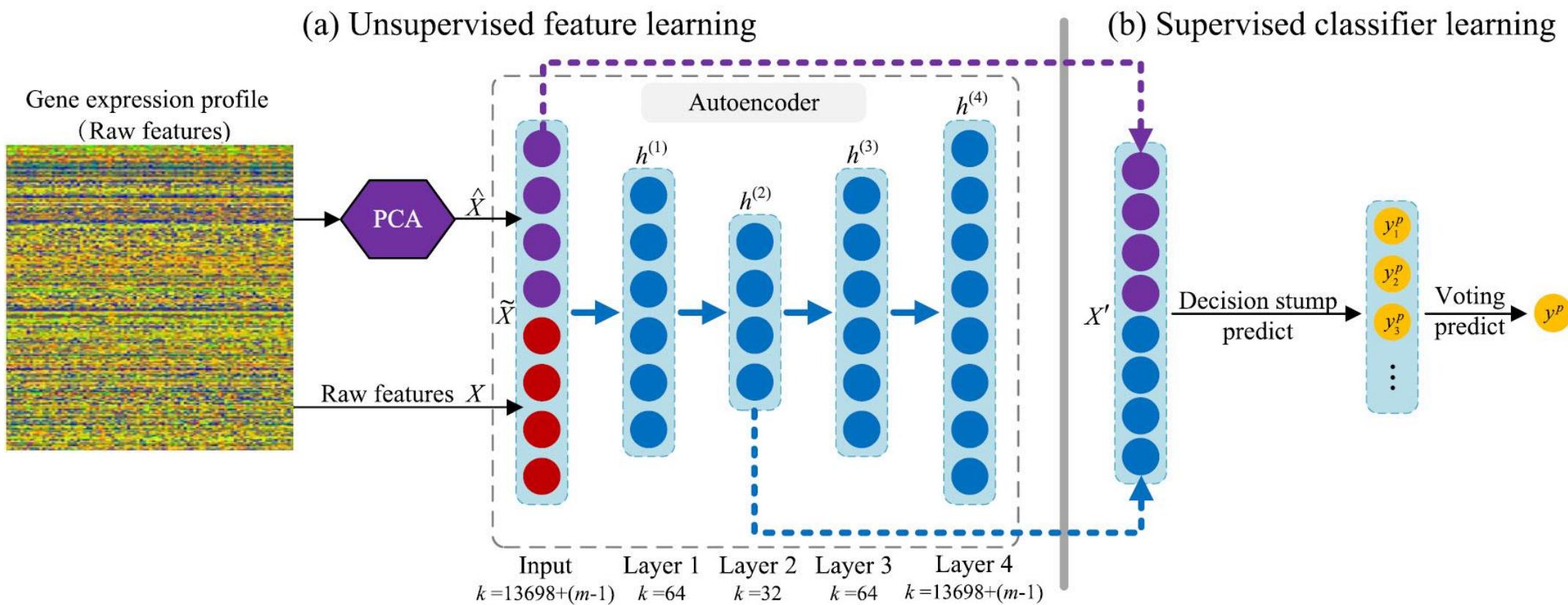
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Breast Cancer Prognosis

Integrating Feature Selection and Feature Extraction Methods With Deep Learning to Predict Clinical Outcome of Breast Cancer (2018)

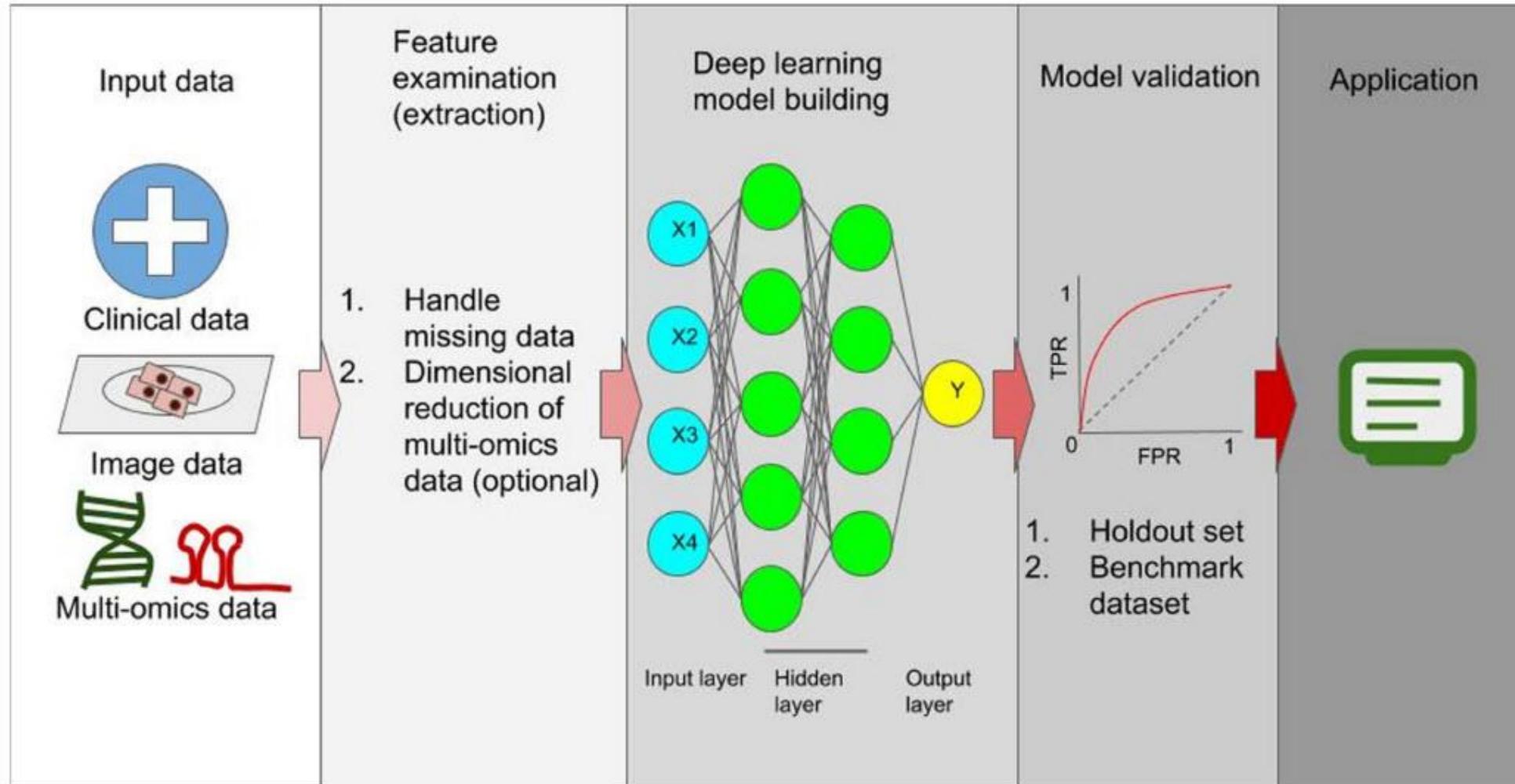
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Breast Cancer Prognosis

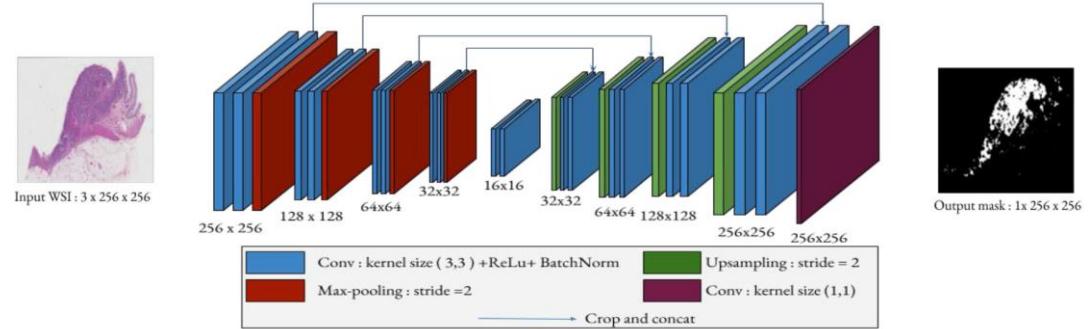


The Application of Deep Learning in Cancer Prognosis Prediction (2020)

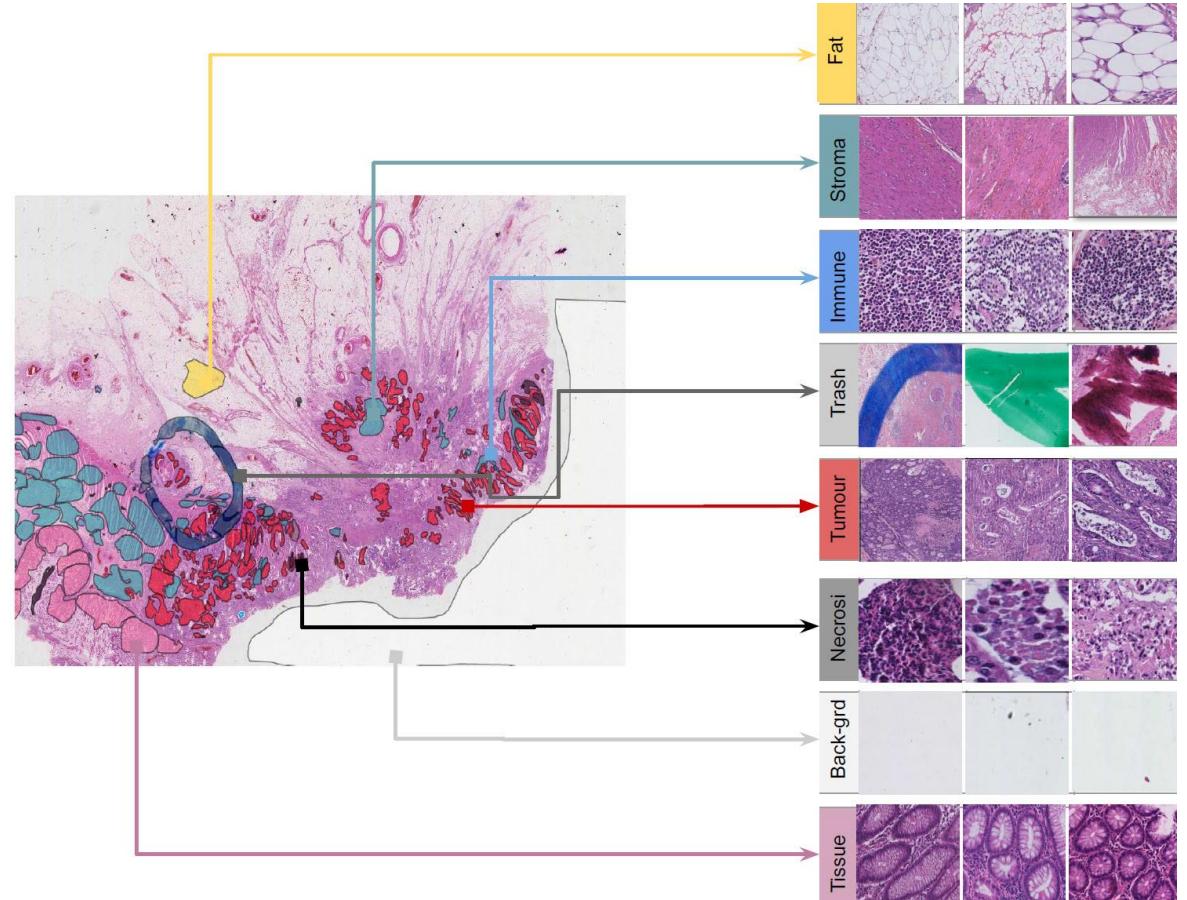


Colon Cancer Analysis

Deep learning for colon cancer histopathological images analysis (2021)

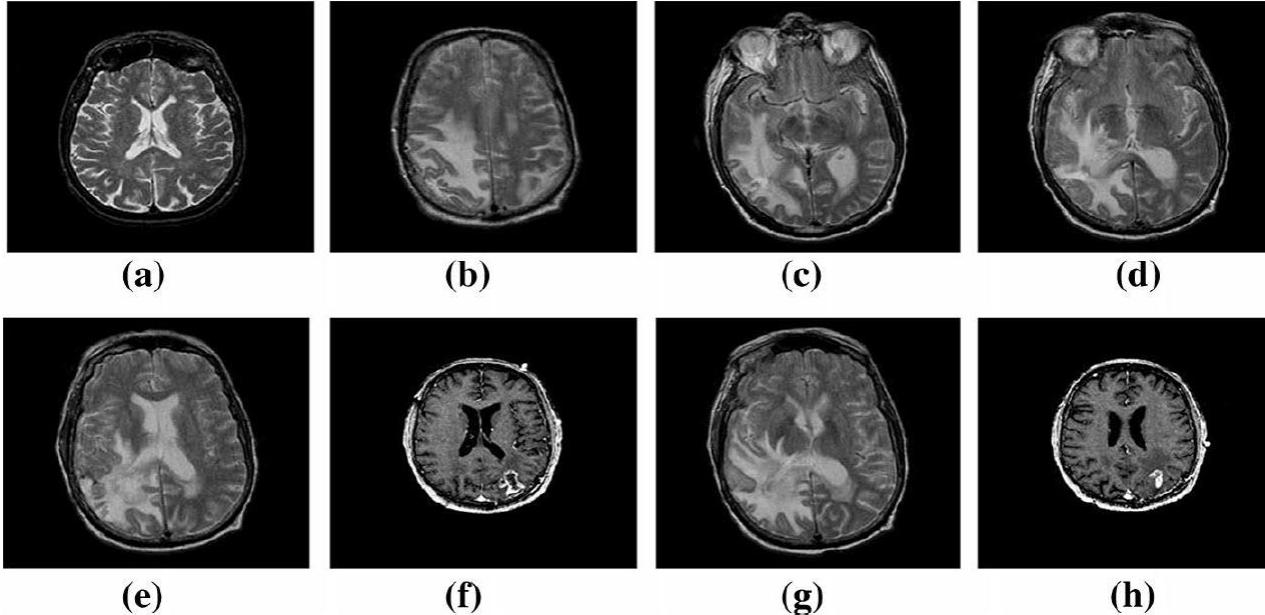


	Trash	Necrosis	Fat	Tumour	Stroma	Tissue	Immune	Backgrnd
Trash	0.9742	0	0	0	0	0	0	0
Necrosis	0	0.9151	0	0.0002	0	0	0.0015	0
Fat	0	0	1.0000	0	0	0	0	0
Tumour	0	0.0011	0	0.9900	0	0	0.0100	0
Stroma	0	0	0	0	0.9706	0.0020	0	0
Tissue	0	0	0	0	0	0.9606	0	0
Immune	0	0.0071	0	0.0038	0	0.0080	0.9320	0
Backgrnd	0.0023	0	0	0	0	0	0	0.9786

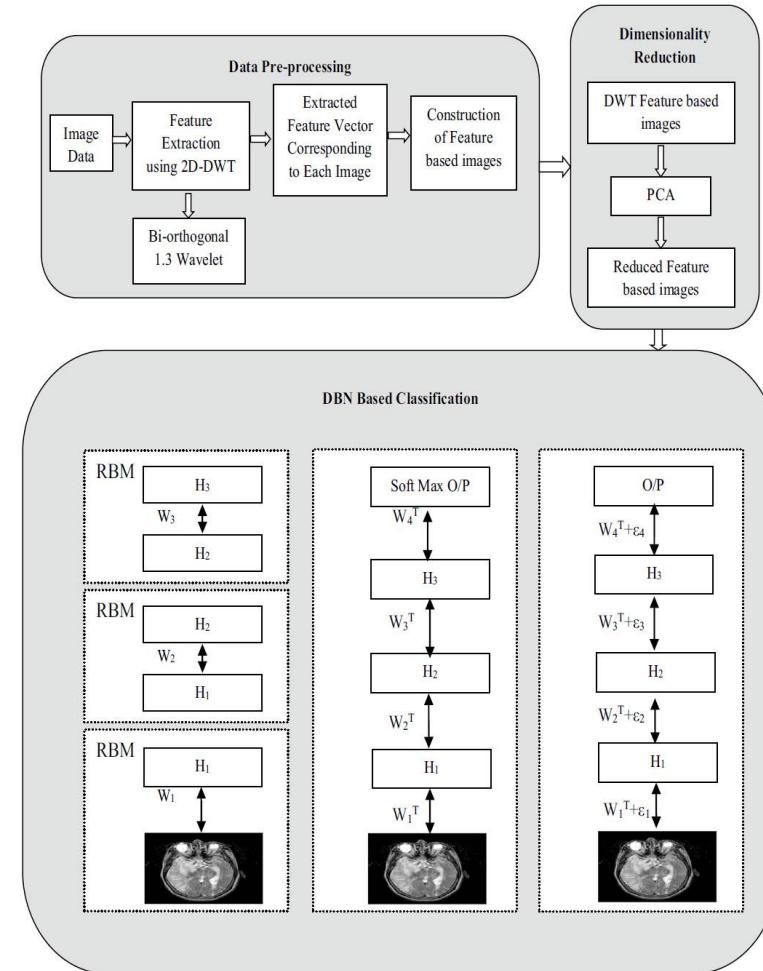


Glioblastoma Detection

Analyzing MRI scans to detect glioblastoma tumor using hybrid deep belief networks (2020)

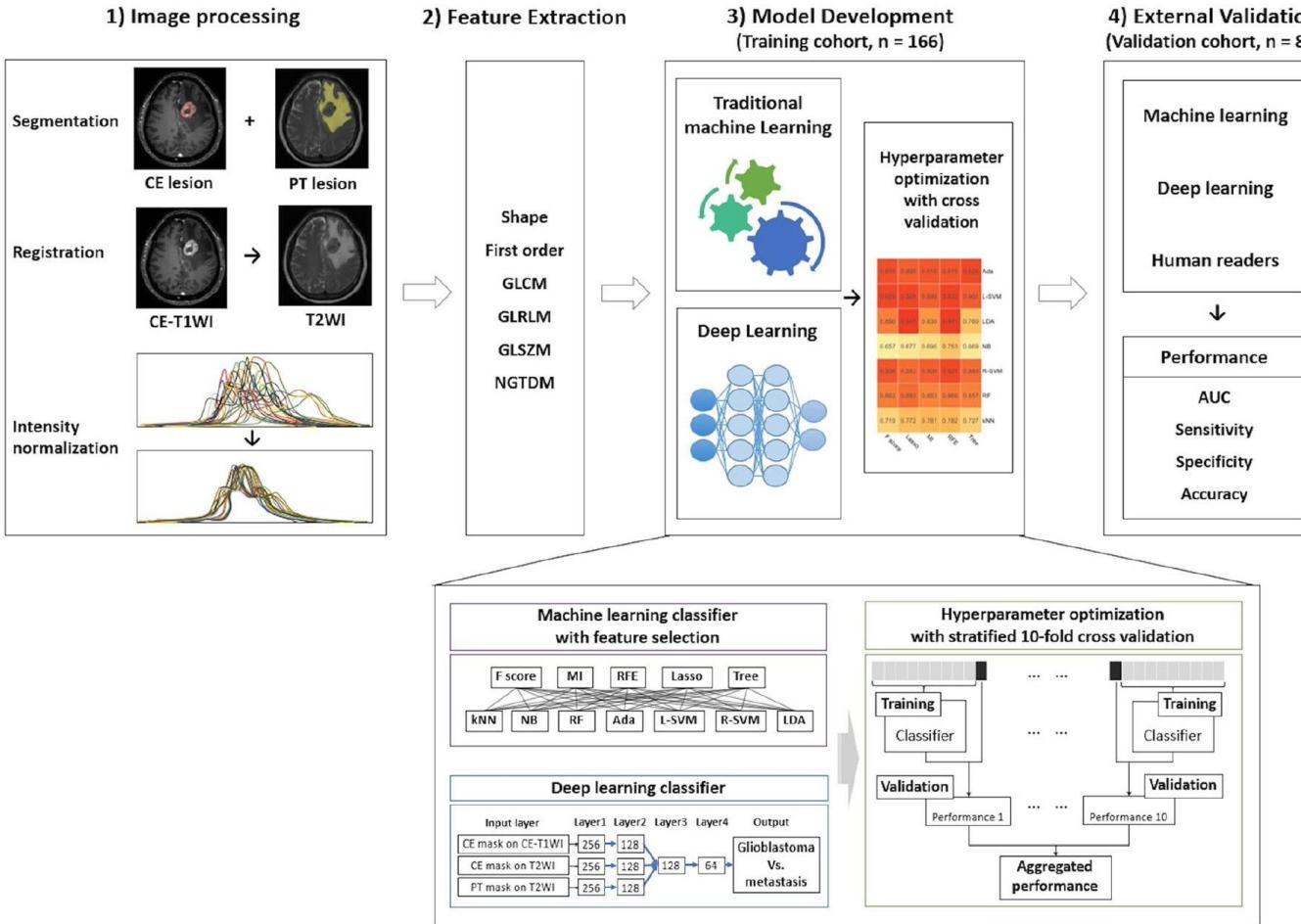


Classifiers	Overall accuracies (%)	Average accuracies (%)	Kappa statistics
DBN	91	89	0.4811
DWT-DBN	93	91	0.5732
DWT-PCA-DBN	97	95	0.6522



Glioblastoma Distinguishing

Robust performance of deep learning for distinguishing glioblastoma from single brain metastasis using radiomic features (2020)



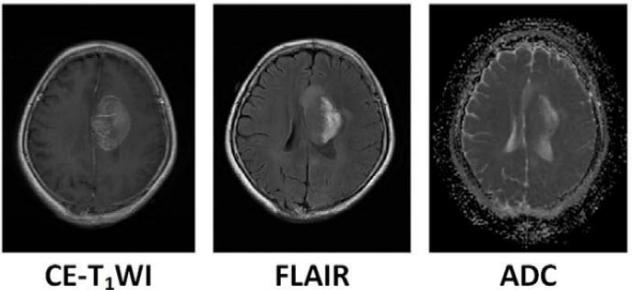
Data	Classifier	Cross validation	External validation			
		Mean AUC	AUC	Sensitivity (%)	Specificity (%)	Accuracy (%)
<i>Traditional machine learning</i>						
CE mask	AdaBoost with LASSO	0.870	0.858 (0.787–0.926)	68.0 (53.3–80.5)	93.8 (79.2–99.2)	78.0 (67.5–86.4)
	L-SVM with Tree-based selection	0.875	0.833 (0.755–0.904)	62.0 (47.2–75.4)	93.8 (79.2–99.2)	74.4 (63.6–83.4)
	LDA with LASSO	0.863	0.818 (0.737–0.891)	64.0 (49.2–77.1)	87.5 (71.0–96.5)	73.2 (62.2–82.4)
PT mask	AdaBoost with Tree-based selection	0.816	0.773 (0.668–0.870)	94.0 (83.5–98.8)	34.4 (18.6–53.2)	70.7 (59.7–80.3)
	L-SVM with RFE	0.830	0.803 (0.718–0.879)	86.0 (75.5–96.5)	65.6 (46.8–81.4)	78.0 (67.2–88.8)
	LDA with MI	0.818	0.787 (0.695–0.870)	94.0 (83.5–98.8)	50.0 (31.9–68.1)	76.8 (66.2–85.4)
Combined mask	AdaBoost with Tree-based selection	0.926	0.890 (0.823–0.947)	80.0 (62.3–90.0)	87.5 (71.0–94.5)	82.9 (73.0–90.3)
	L-SVM with RFE	0.932	0.886 (0.798–0.927)	80.0 (62.3–90.0)	84.4 (67.2–94.7)	81.7 (71.6–89.4)
	LDA with LASSO	0.945	0.899 (0.839–0.951)	84.0 (70.9–92.8)	78.1 (70.9–90.7)	81.7 (71.6–89.4)
<i>Deep learning</i>						
CE mask	DNN	0.887	0.887 (0.812–0.951)	62.5 (45.7–79.3)	96.0 (90.6–100)	82.9 (74.8–91.1)
PT mask	DNN	0.865	0.825 (0.722–0.887)	75.0 (60.2–90.1)	82.0 (71.4–92.6)	79.3 (70.5–88.0)
Combined mask	DNN	0.986	0.956 (0.918–0.990)	90.6 (80.5–100)	88.0 (79.0–97.0)	89.0 (82.3–95.8)
<i>Human reading</i>						
Images	Reader 1		0.774 (0.685–0.852)	97.0 (91.1–100)	50.0 (36.1–63.9)	68.7 (58.7–78.7)
Images	Reader 2		0.904 (0.852–0.951)	81.8 (68.7–95.0)	78.0 (66.5–89.5)	79.5 (70.8–88.2)

Glioblastoma Distinguishing

Deep Learning for Automatic Differential Diagnosis of Primary Central Nervous System Lymphoma

And Glioblastoma: Multi-Parametric Magnetic Resonance Imaging (2021)

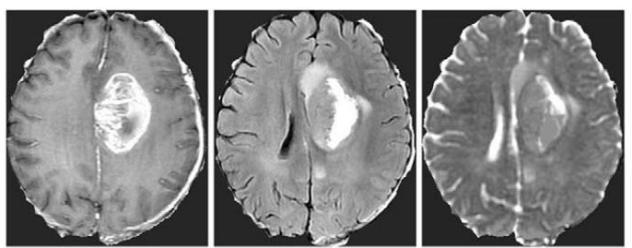
I. Automatic image preprocessing



1. Registration



2. Brain extraction



3. Standardization

II. Single-parametric CNN model development

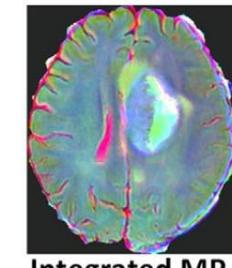


Pre-trained on ImageNet and fine-tuned using MRI images

DenseNet based CNN model training through transfer learning

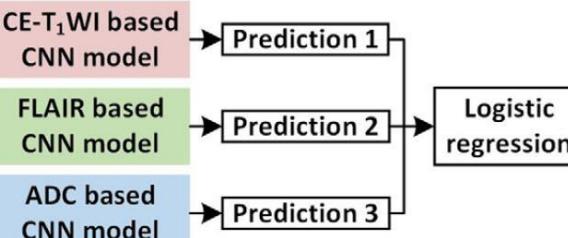
CE-T₁WI based CNN model
FLAIR based CNN model
ADC based CNN model

III. Multi-parametric CNN model development



Integrated MP-MR image

Image-level fusion based multi-parametric CNN model (IF-CNN model)



Decision-level fusion based multi-parametric CNN model (DF-CNN model)

Glioblastoma Distinguishing

Deep Learning for Automatic Differential Diagnosis of Primary Central Nervous System Lymphoma

And Glioblastoma: Multi-Parametric Magnetic Resonance Imaging (2021)

Models and Radiologists	Model Type	ACC	SEN	SPE	AUC
CNN models	CE-T1WI	0.884	0.934	0.841	0.956
	FLAIR	0.782	0.772	0.790	0.852
	ADC	0.700	0.860	0.558	0.828
	IF-CNN	0.830	0.772	0.882	0.919
	DF-CNN	0.899	0.934	0.867	0.964
Radiomics models	CE-T1WI	0.865	0.838	0.890	0.925
	FLAIR	0.771	0.757	0.784	0.867
	ADC	0.796	0.758	0.830	0.868
	IF-RADS	0.856	0.831	0.878	0.929
	DF-RADS	0.820	0.838	0.804	0.911
Junior radiologist	–	0.875	0.823	0.921	–
Intermediate-level radiologist	–	0.878	0.801	0.947	–
Senior radiologist	–	0.906	0.853	0.954	–

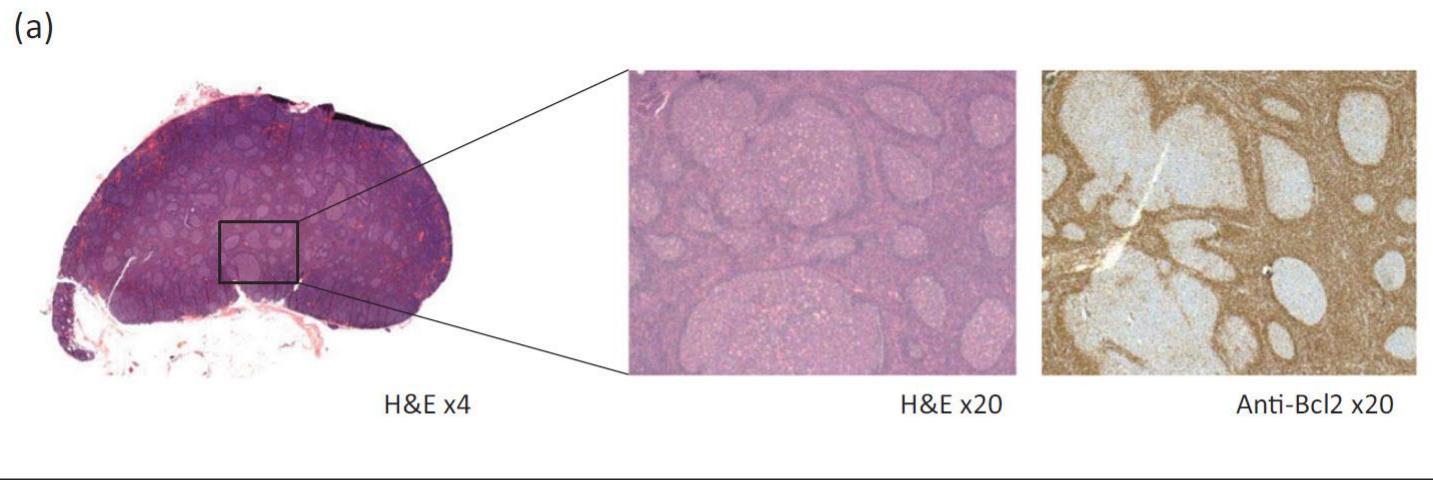
CNN = convolutional neural network; ACC = accuracy; SEN = sensitivity; SPE = specificity; AUC = area under the ROC curve;
IF = image-level fusion; DF = decision-level fusion.

Lymphoma Accurate Diagnosis

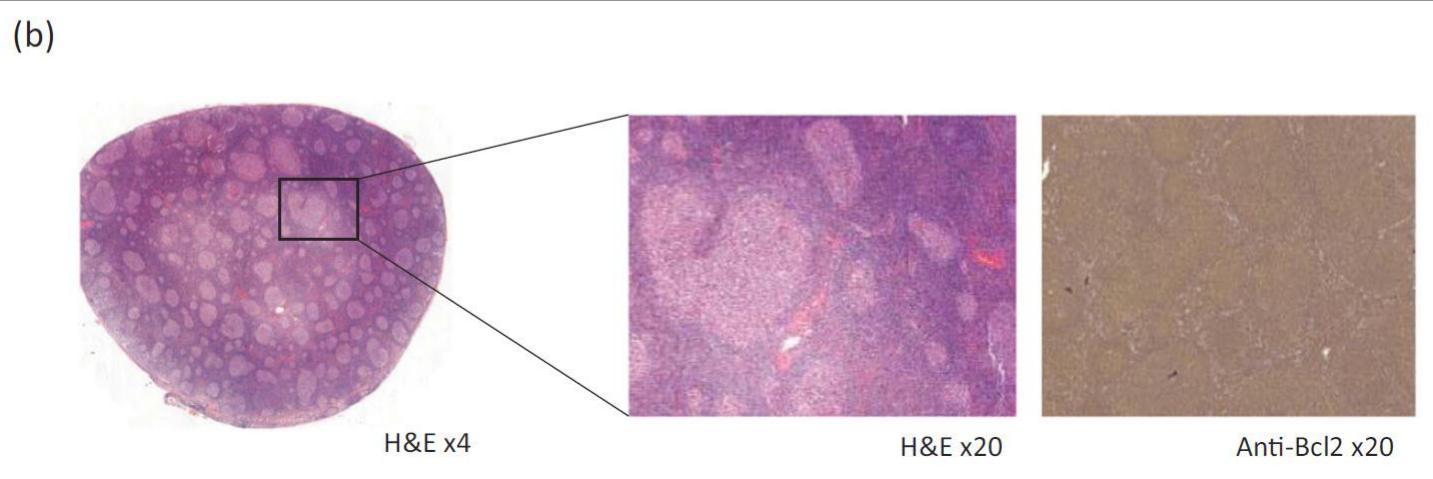
Accurate diagnosis of lymphoma on whole-slide histopathology images
using deep learning (2020)

nature

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Follicular Hyperplasia (FH)



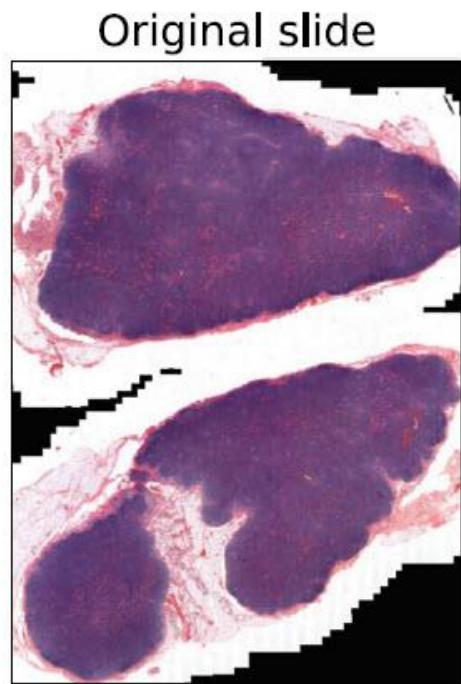
Follicular Lymphoma (FL)

Lymphoma Accurate Diagnosis

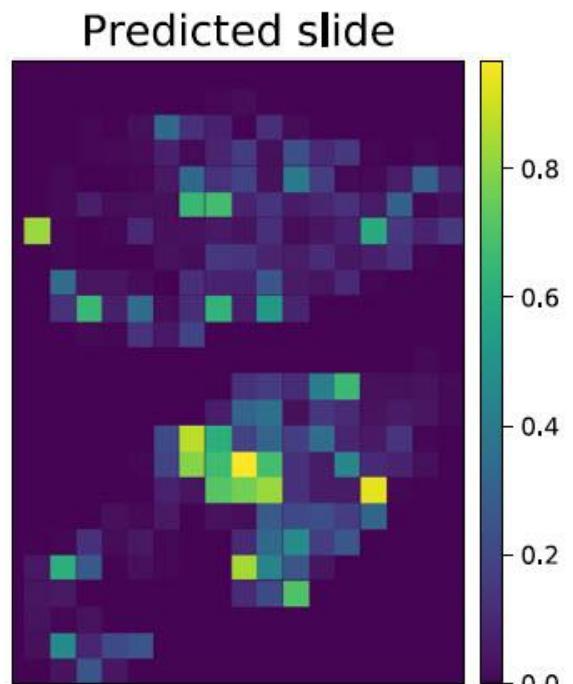
Accurate diagnosis of lymphoma on whole-slide histopathology images
using deep learning (2020)

nature

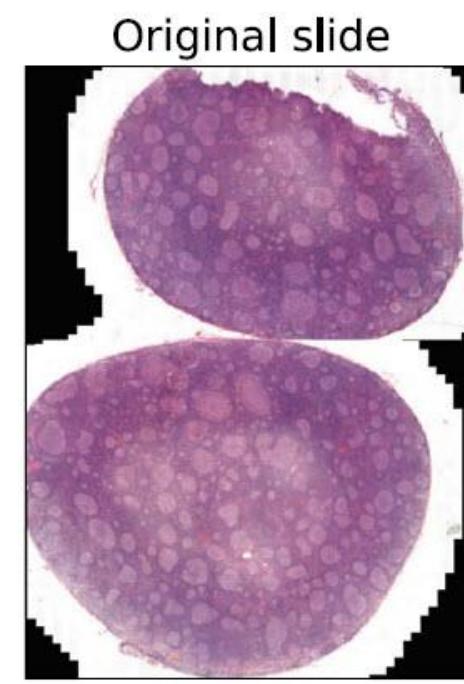
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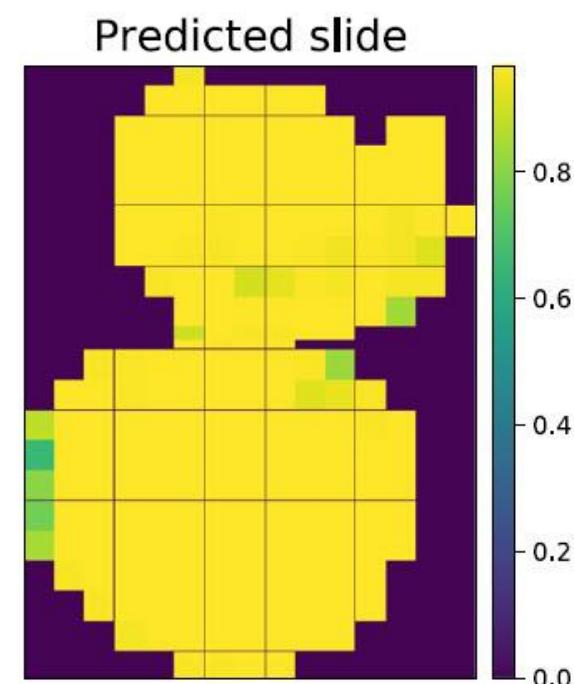
(a)



(b)



(c)

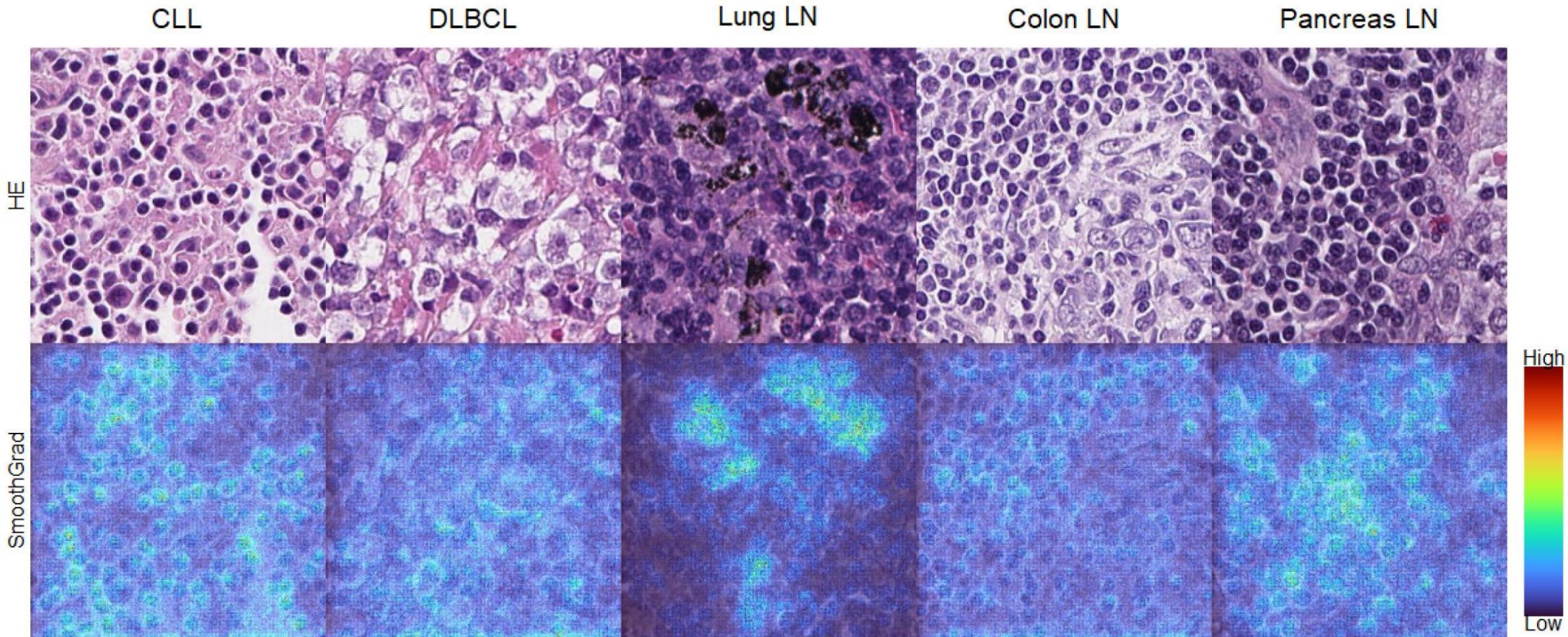


(d)

Lymphoma Non-Hodgkin Classification



Deep Learning for the Classification of Non-Hodgkin Lymphoma on Histopathological Images (2021)



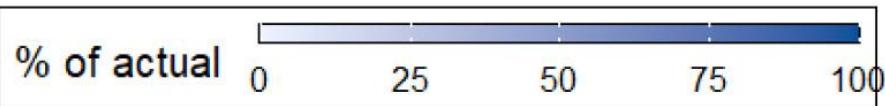
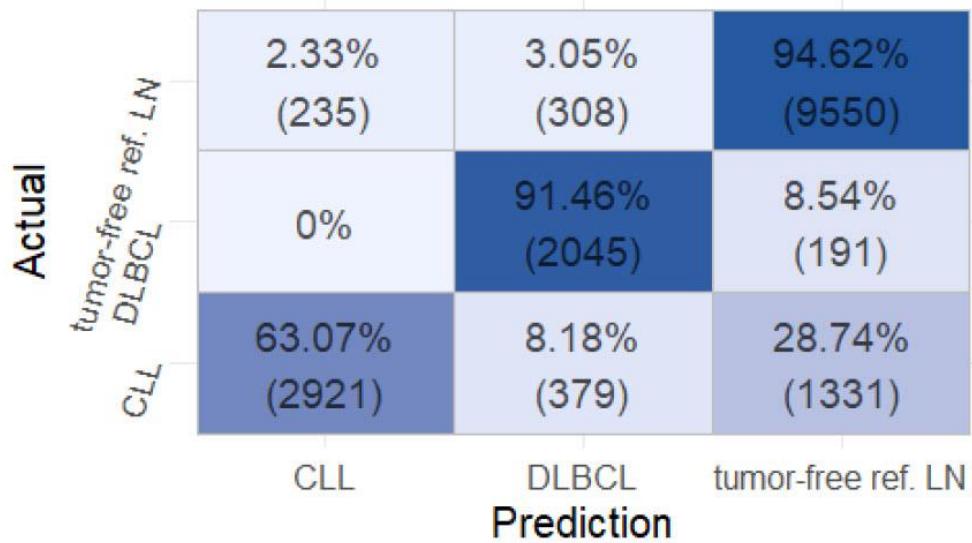
Lymphoma Non-Hodgkin Classification



Deep Learning for the Classification of Non-Hodgkin Lymphoma on Histopathological Images (2021)

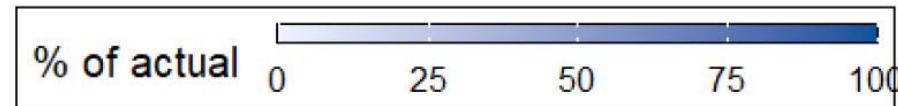
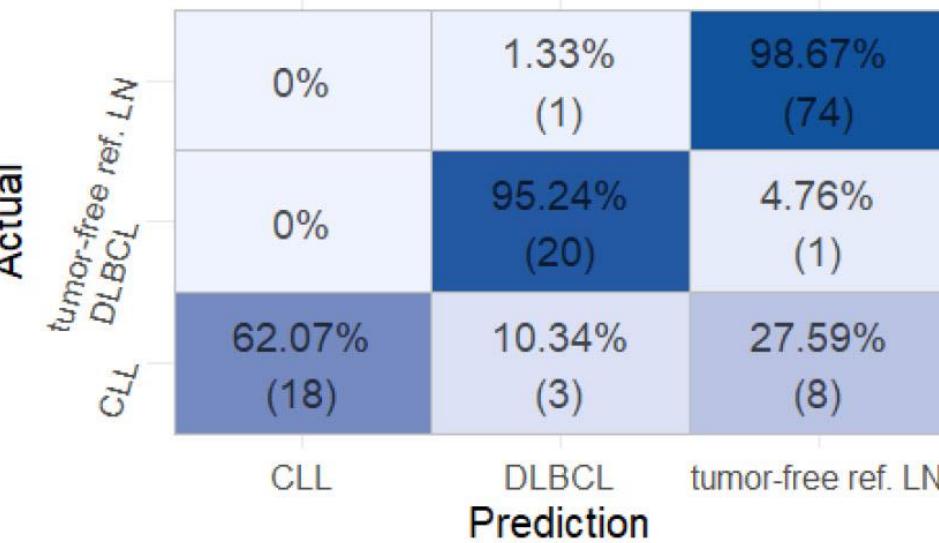
EfficientNetB3 Test Data

Patch based confusion matrix (BACC=0.83)



EfficientNetB3 Test Data

Case based confusion matrix (BACC=0.85)





Motivation & Hope...

Dream Big, Work Hard and Stay focused.



2003 - 2006

6 times Surgery

+30 times Chemotherapy and...



Thanks!

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Nov 2021

