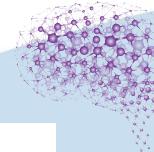




Institut et hôpital neurologiques de Montréal  
Montreal Neurological Institute and Hospital



Jewish General Hospital  
Lady Davis Institute for Medical Research



CENTRE  
**LUDMER**  
NEUROINFORMATIQUE & SANTÉ MENTALE | NEUROINFORMATICS & MENTAL HEALTH

# Intro to Brain Segmentation with Keras

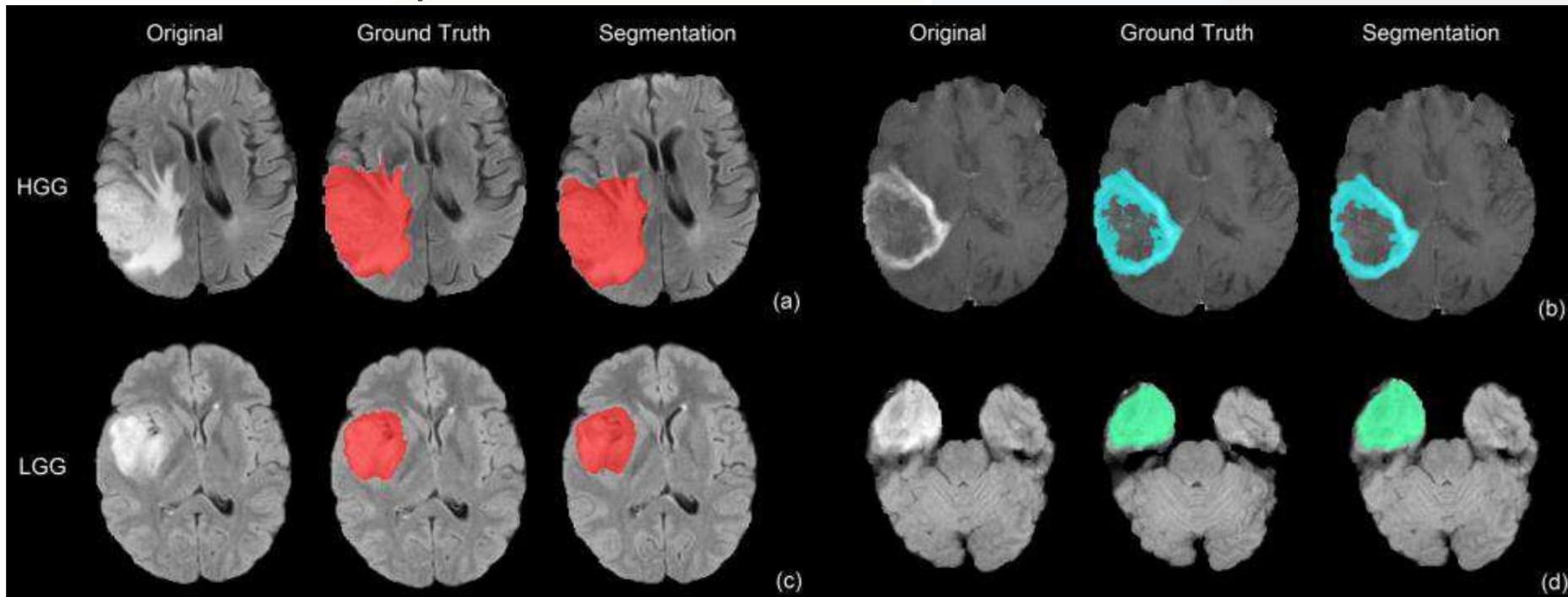
MAIN 2018 Educational Course

Thomas Funck, McGill University

Email: [tffunck@gmail.com](mailto:tffunck@gmail.com); Twitter: [@tffunck](https://twitter.com/tffunck)

# Why CNNs for segmentation?

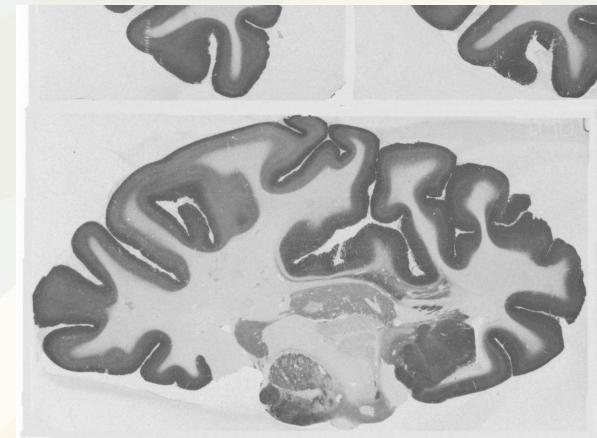
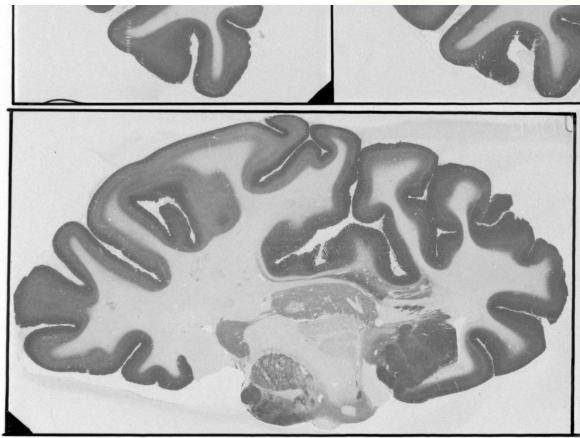
- Can solve complicated problems
  - ...if you have the data + labels



Dong, et al. (2017) Automatic Brain Tumor Detection and Segmentation Using U-Net Based Fully Convolutional Networks. <https://arxiv.org/abs/1705.03820>

# Why CNNs for segmentation?

- Can solve complicated problems
- Can also solve simple, but annoying problems



# Why CNNs for segmentation?

- Can solve complicated problems
- Can also solve simple, but annoying problems
- Goal: Dive into nitty-gritty to building a CNN
  - So that you can go back to the lab and create your own



# Outline

1. Basic Concepts
2. Simple Networks
3. U-Net



# Outline

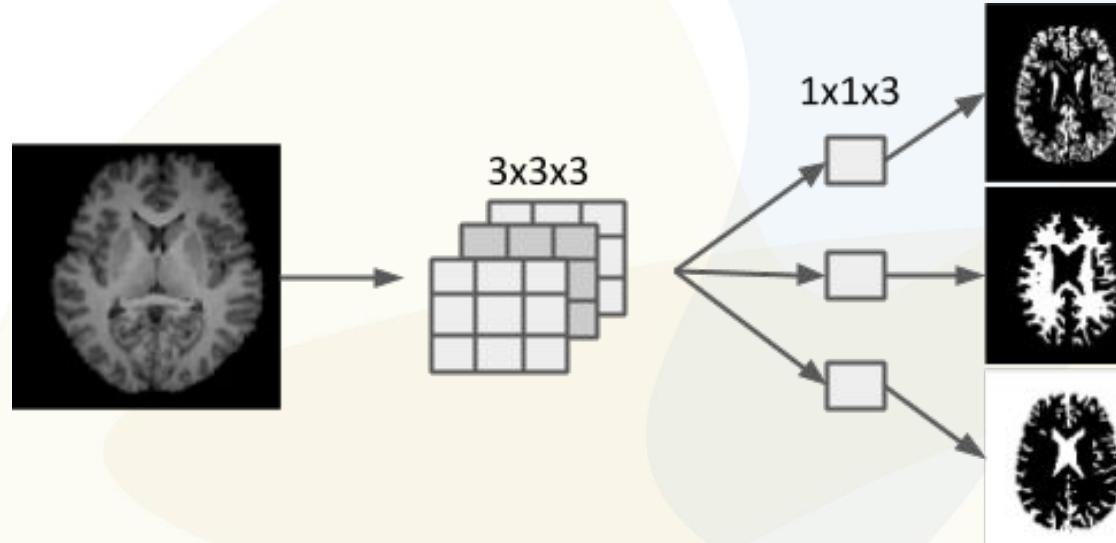
## 1. Basic Concepts

- a. Kernels
- b. Receptive field & dilations
- c. Upsampling & downsampling
- d. Final activation functions
- e. Loss functions
- f. Metrics
- g. Cross-validation
- h. Feature extraction



# Example of a simple convolutional network

- 1 Layer, 3 Kernels of 3x3 size



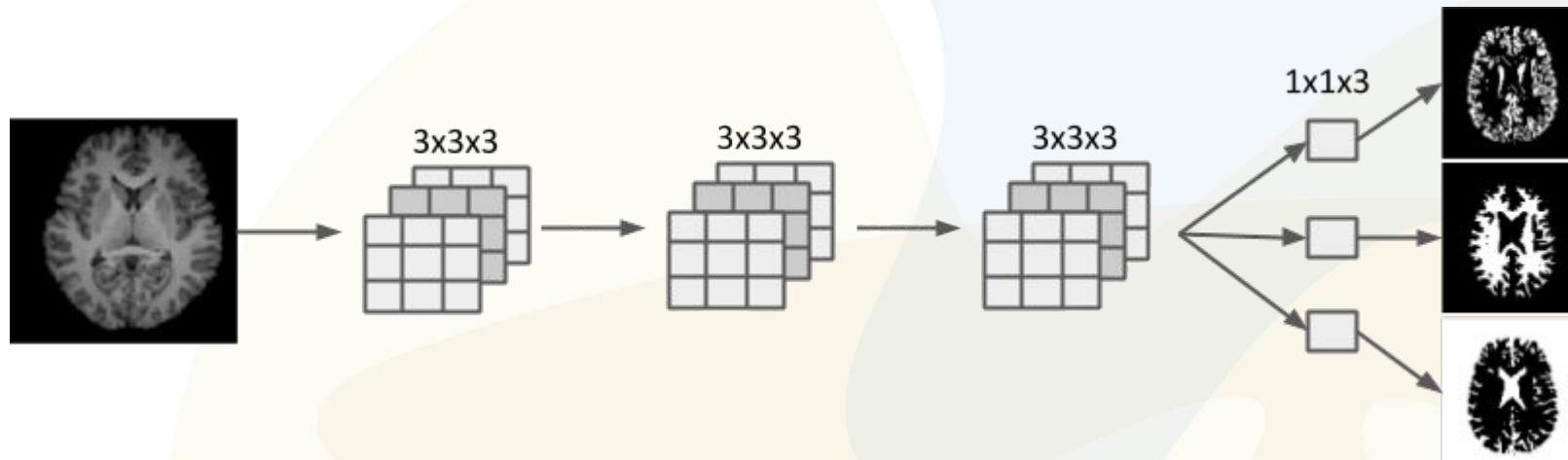
# Example of a simple convolutional network

Layer (type)	Output Shape	Param #
=====		
=====		
input_1 (InputLayer)	(None, 110, 92, 1)	0
=====		
conv2d_1 (Conv2D)	(None, 110, 92, 3)	30
=====		
dropout_1 (Dropout)	(None, 110, 92, 3)	0
=====		
conv2d_2 (Conv2D)	(None, 110, 92, 3)	12
=====		
=====		
Total params: 42		
Trainable params: 42		
Non-trainable params: 0		



# Example of a simple convolutional network

- 3 Layer, 3 Kernels of 3x3 size



# Example of a simple convolutional network

- 3 Layer, 3 Kernels of 3x3 size

Layer (type)	Output Shape	Param #
input_2 (InputLayer)	(None, 110, 92, 1)	0
conv2d_3 (Conv2D)	(None, 110, 92, 3)	30
dropout_2 (Dropout)	(None, 110, 92, 3)	0
conv2d_4 (Conv2D)	(None, 110, 92, 3)	84
dropout_3 (Dropout)	(None, 110, 92, 3)	0
conv2d_5 (Conv2D)	(None, 110, 92, 3)	84
dropout_4 (Dropout)	(None, 110, 92, 3)	0
conv2d_6 (Conv2D)	(None, 110, 92, 3)	12

Total params: 210

Trainable params: 210

Non-trainable params: 0



# *Different number of parameters. What gives?*

- 3 Layer, 3 Kernels of 3x3 size

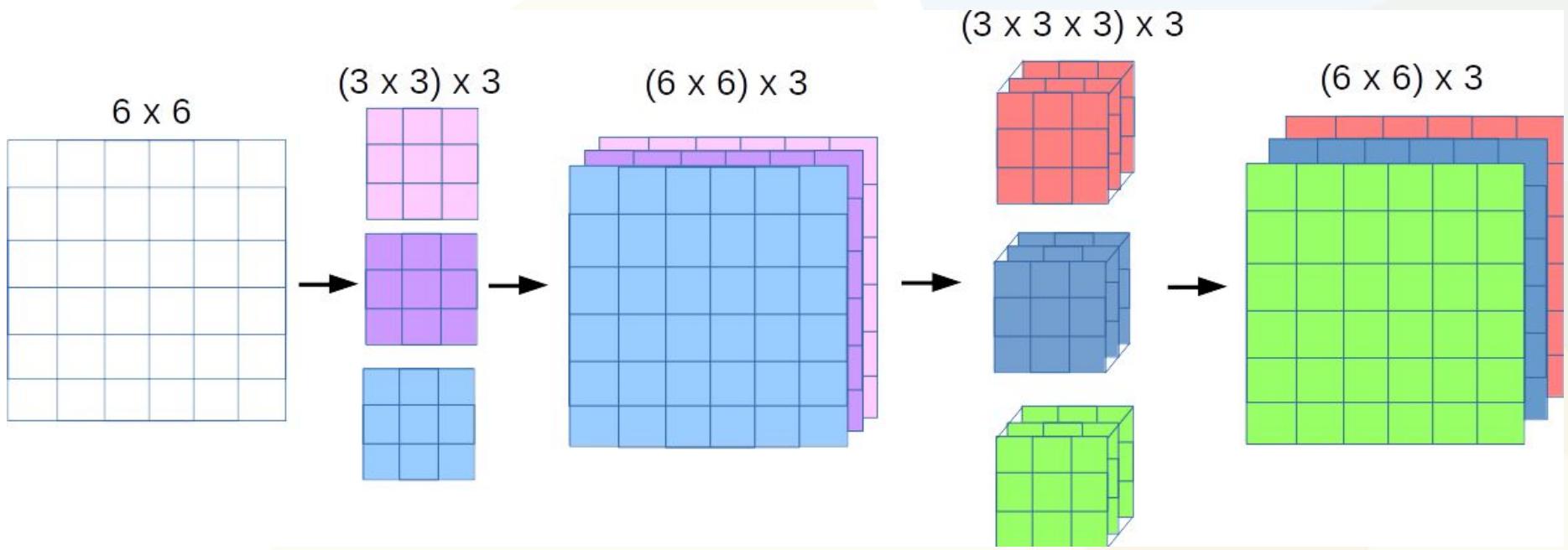
Layer (type)	Output Shape	Param #
input_2 (InputLayer)	(None, 110, 92, 1)	0
<b>conv2d_3 (Conv2D)</b>	<b>(None, 110, 92, 3)</b>	<b>30</b>
dropout_2 (Dropout)	(None, 110, 92, 3)	0
<b>conv2d_4 (Conv2D)</b>	<b>(None, 110, 92, 3)</b>	<b>84</b>
dropout_3 (Dropout)	(None, 110, 92, 3)	0
conv2d_5 (Conv2D)	(None, 110, 92, 3)	84
dropout_4 (Dropout)	(None, 110, 92, 3)	0
conv2d_6 (Conv2D)	(None, 110, 92, 3)	12

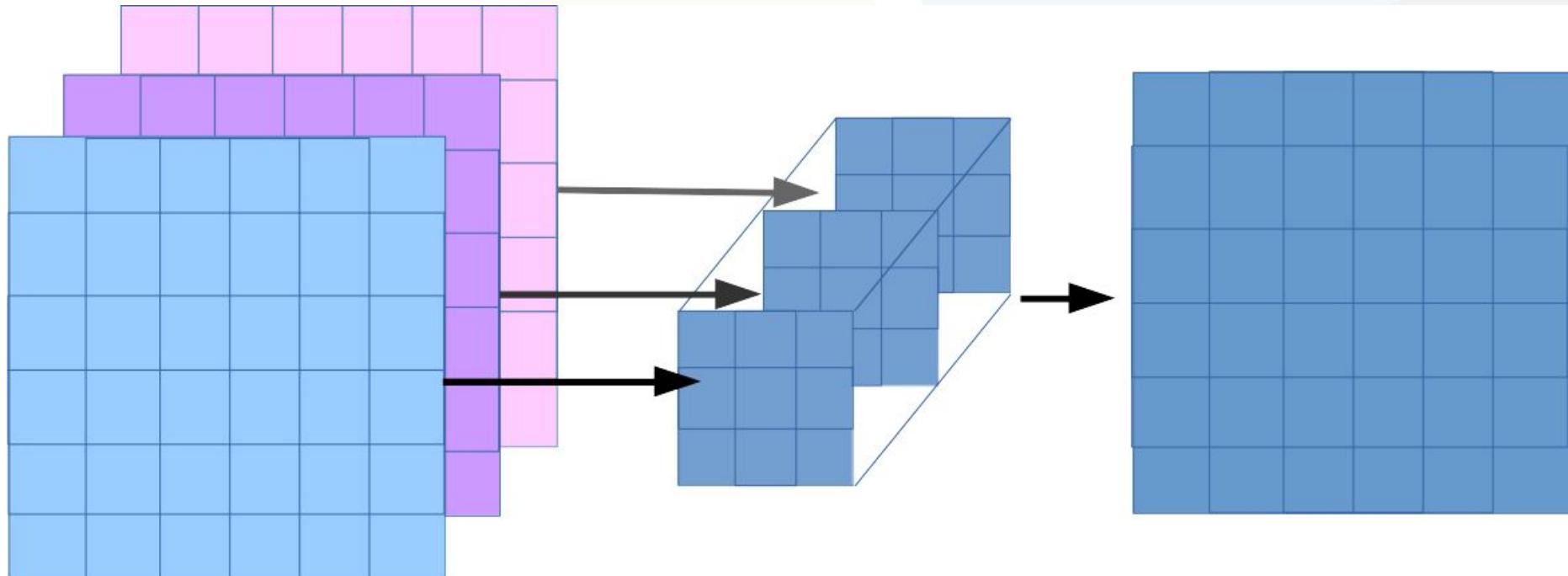
Total params: 210

Trainable params: 210

Non-trainable params: 0







$$N_{parameters} = N_{kernels} \times Kernel_{Dimension1} \times Kernel_{Dimension2} \times Kernel_{Channels} + N_{kernels}$$

$$84 = 3 \times 3 \times 3 \times 3 + 3$$

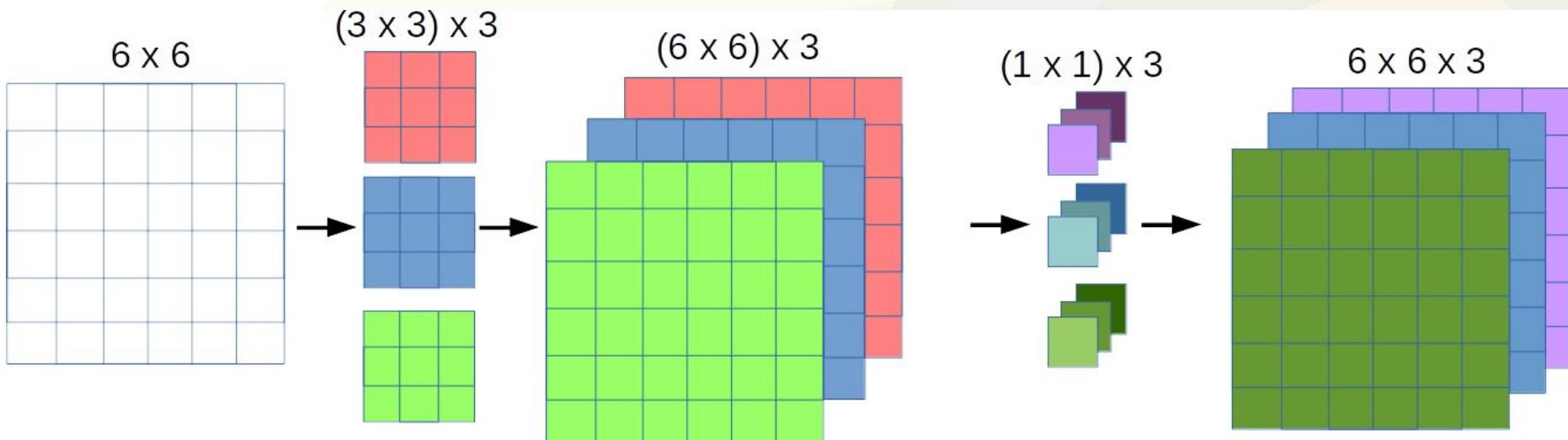
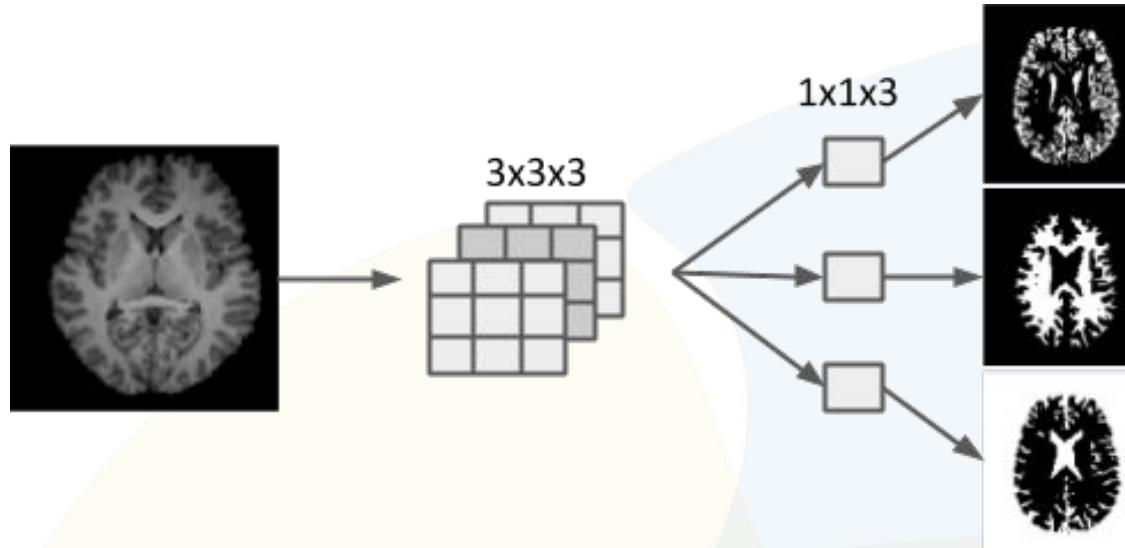
Layer (type)	Output Shape	Param #
input_2 (InputLayer)	(None, 110, 92, 1)	0
<b>conv2d_3 (Conv2D)</b>	<b>(None, 110, 92, 3)</b>	<b>30</b>
dropout_2 (Dropout)	(None, 110, 92, 3)	0
<b>conv2d_4 (Conv2D)</b>	<b>(None, 110, 92, 3)</b>	<b>84</b>
dropout_3 (Dropout)	(None, 110, 92, 3)	0
conv2d_5 (Conv2D)	(None, 110, 92, 3)	84
dropout_4 (Dropout)	(None, 110, 92, 3)	0
<b>conv2d_6 (Conv2D)</b>	<b>(None, 110, 92, 3)</b>	<b>12</b>

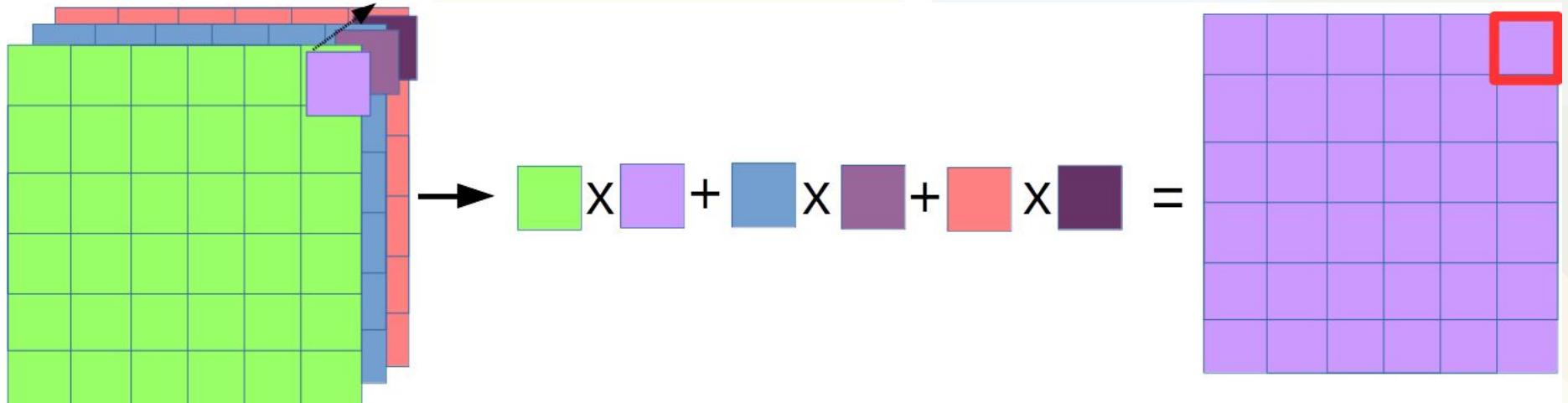
Total params: 210

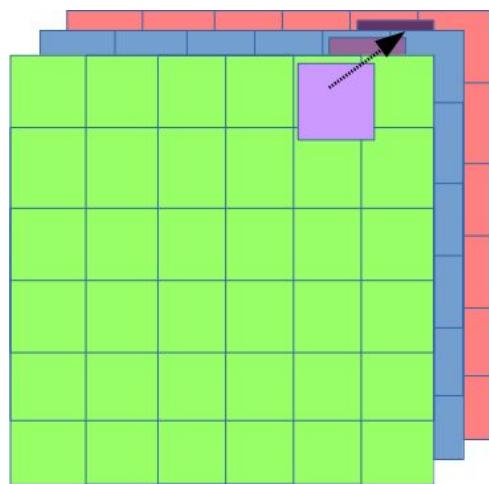
Trainable params: 210

Non-trainable params: 0

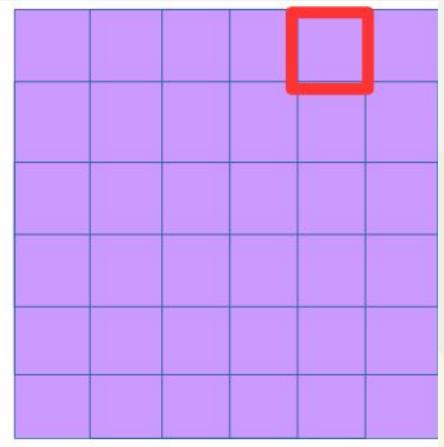


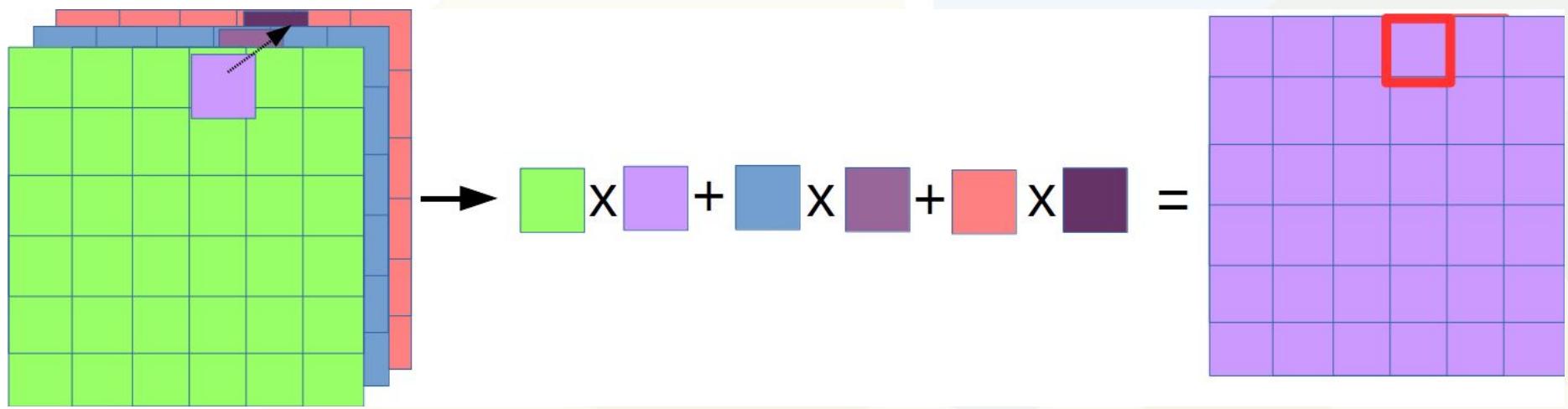






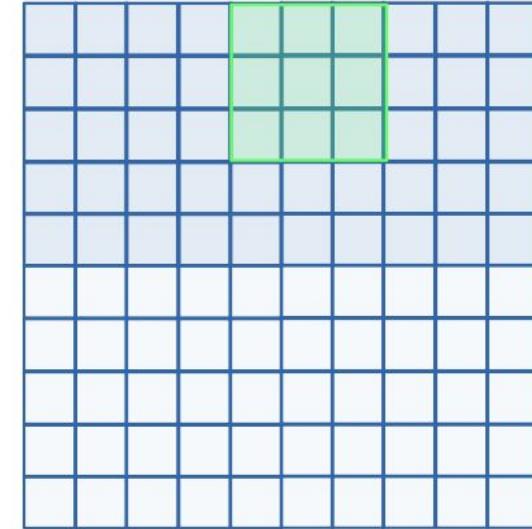
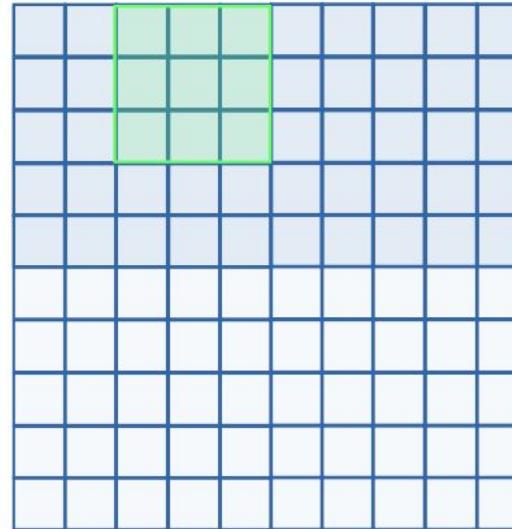
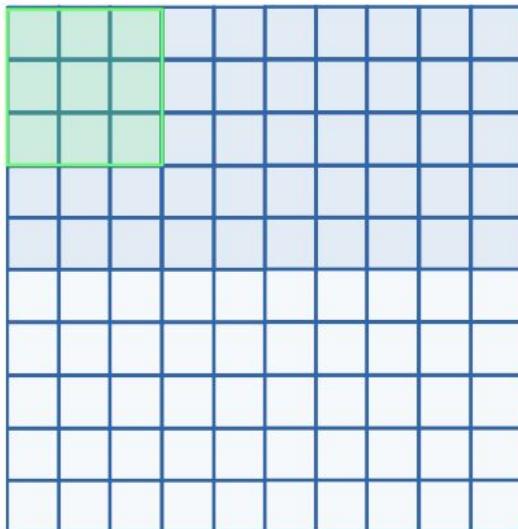
$$\text{[Green]} \times \text{[Purple]} + \text{[Blue]} \times \text{[Purple]} + \text{[Red]} \times \text{[Purple]} =$$





# Kernel Stride

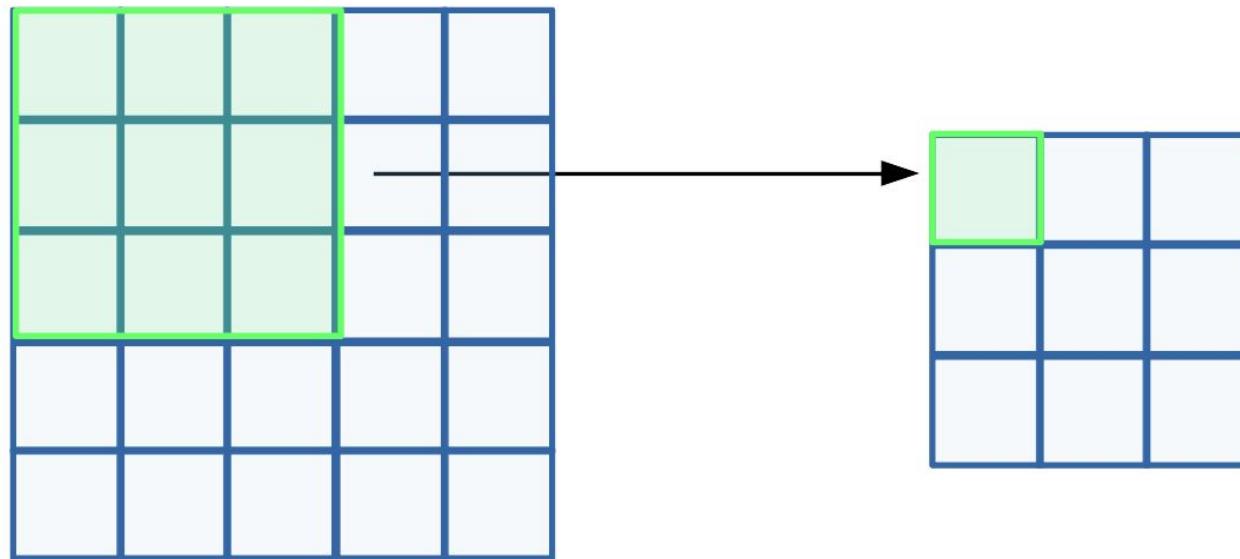
- must divide the image evenly
  - you can pad the image if necessary



kernel stride = 2

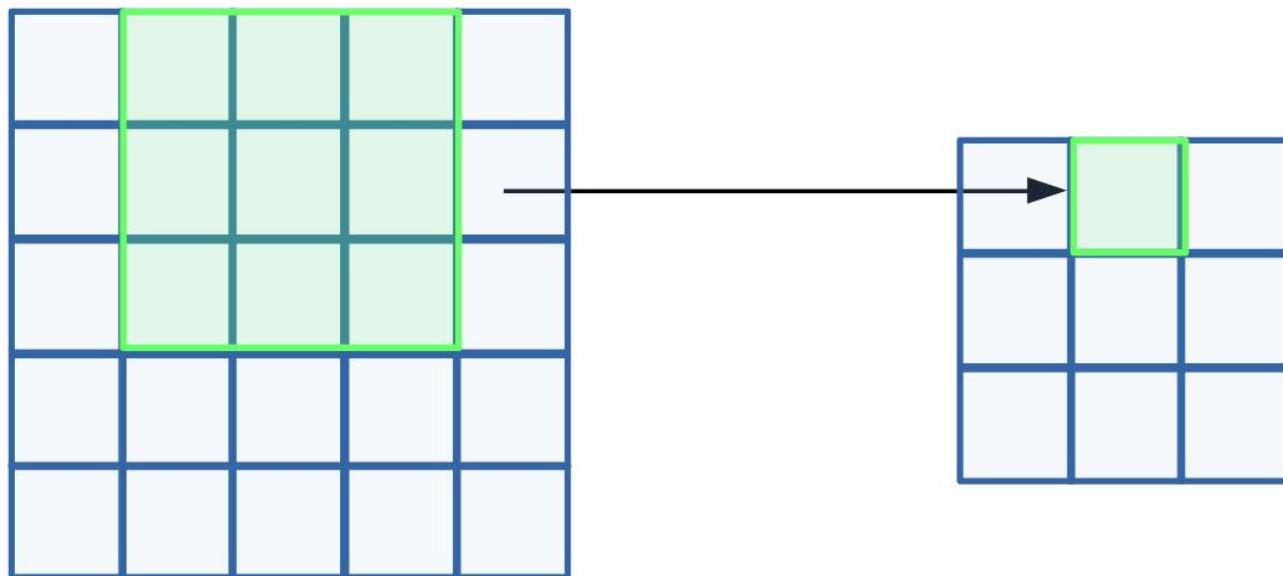
# Kernel Padding

*padding = “valid”*



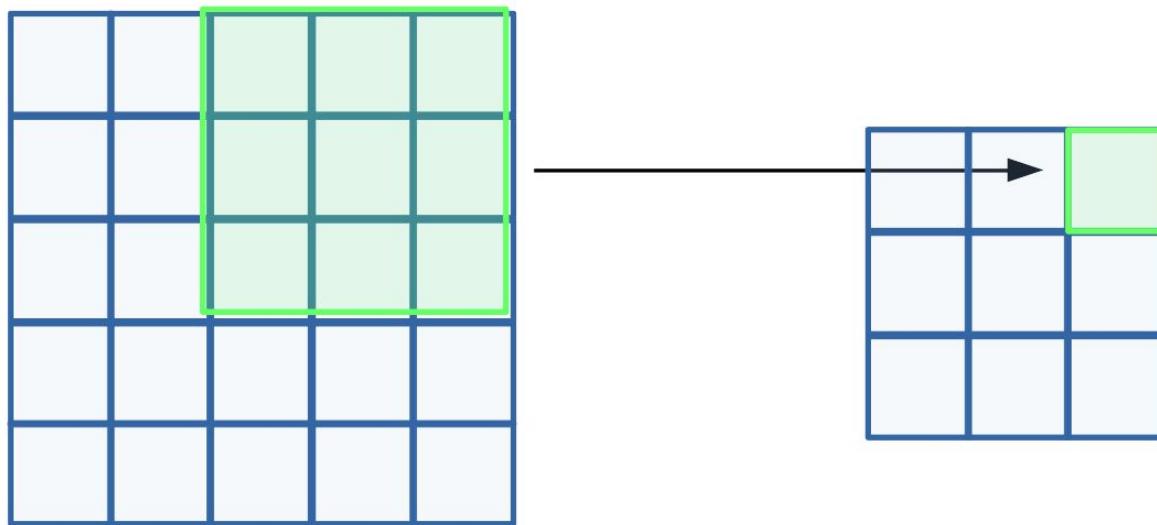
# Kernel Padding

*padding = “valid”*



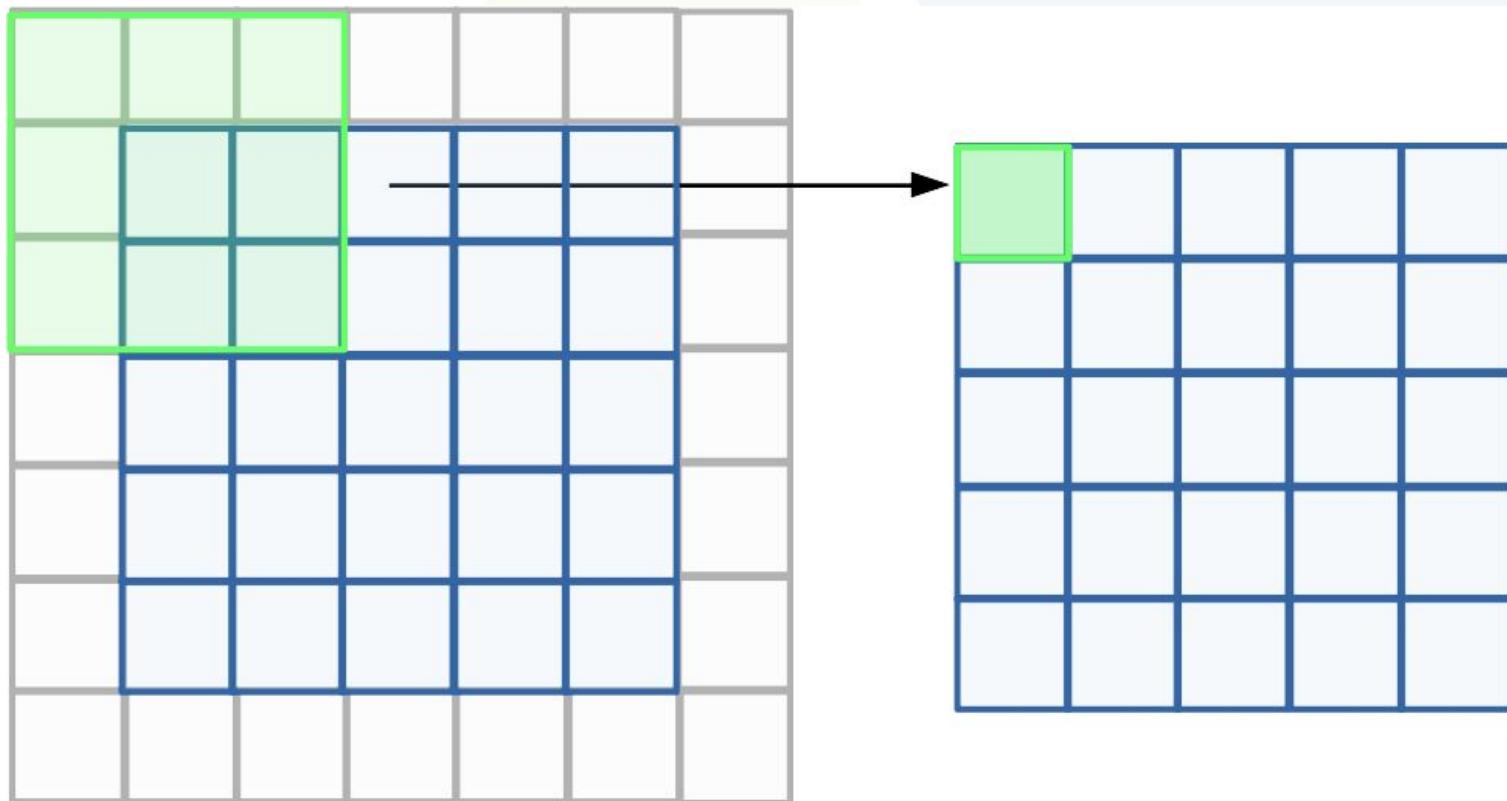
# Kernel Padding

*padding = “valid”*



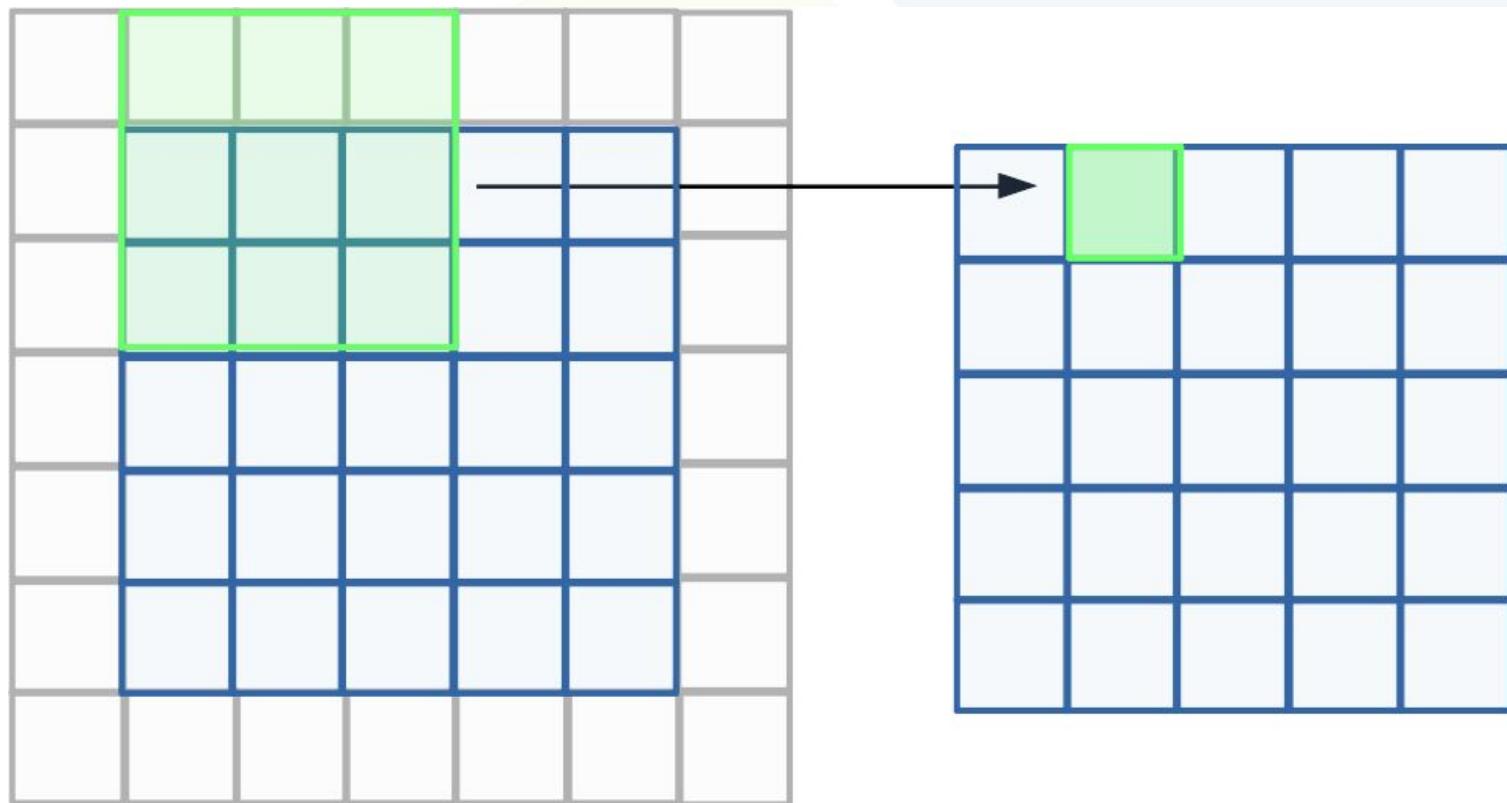
# Kernel Padding

*padding = "same"*



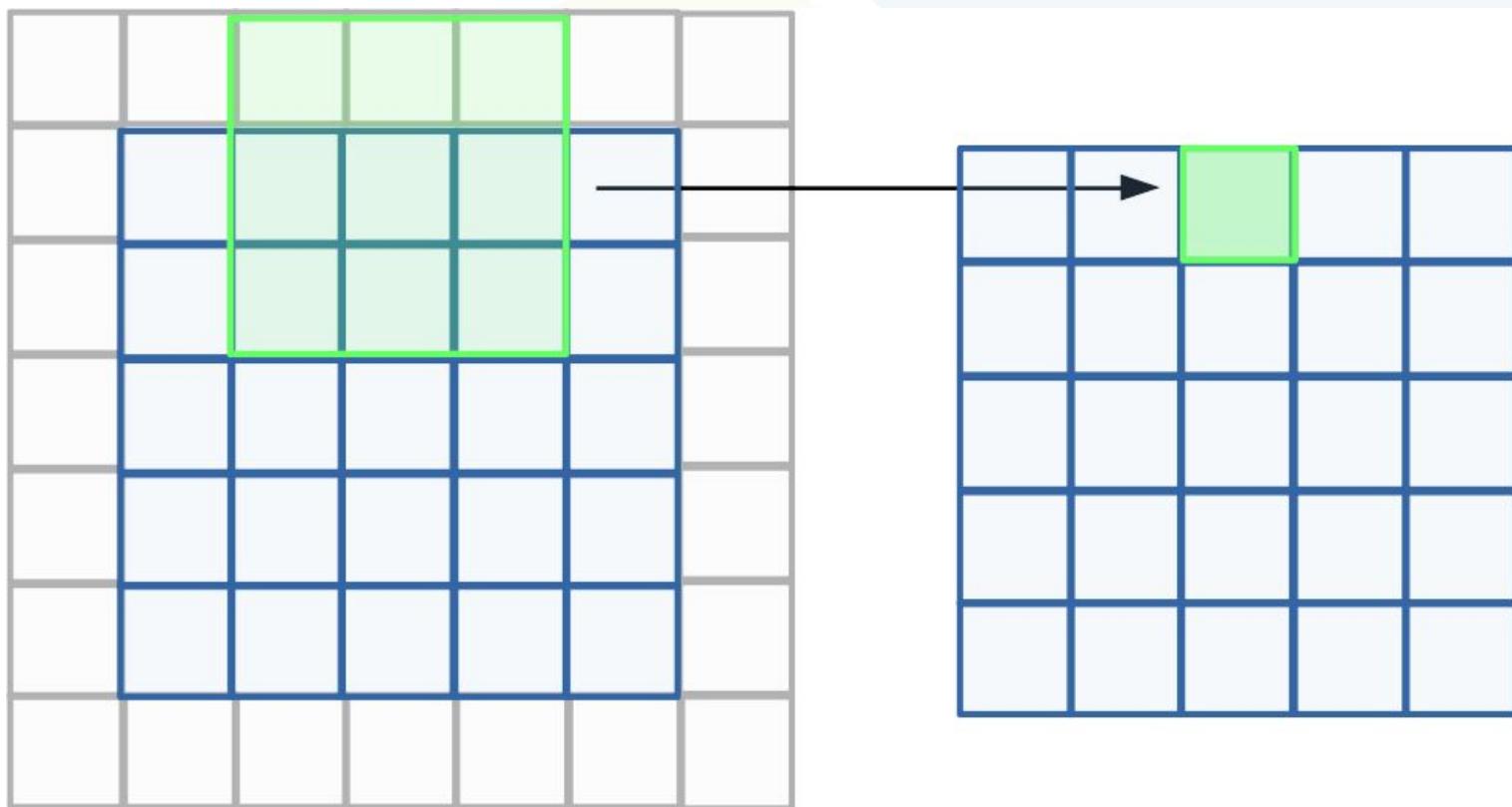
# Kernel Padding

*padding = "same"*



# Kernel Padding

*padding = "same"*

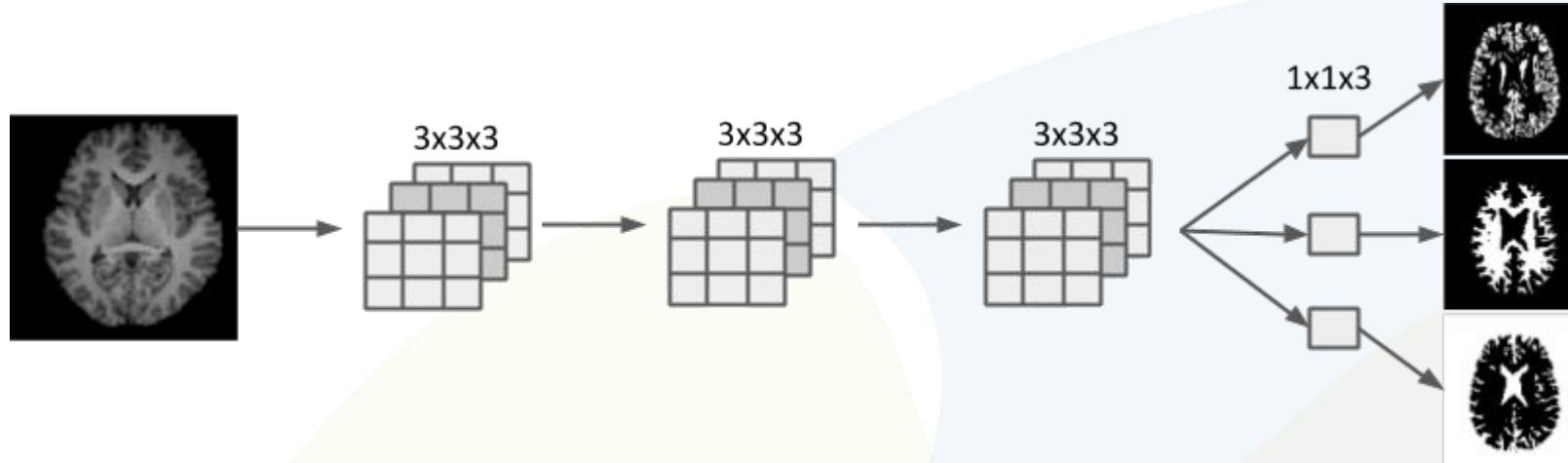


# Outline

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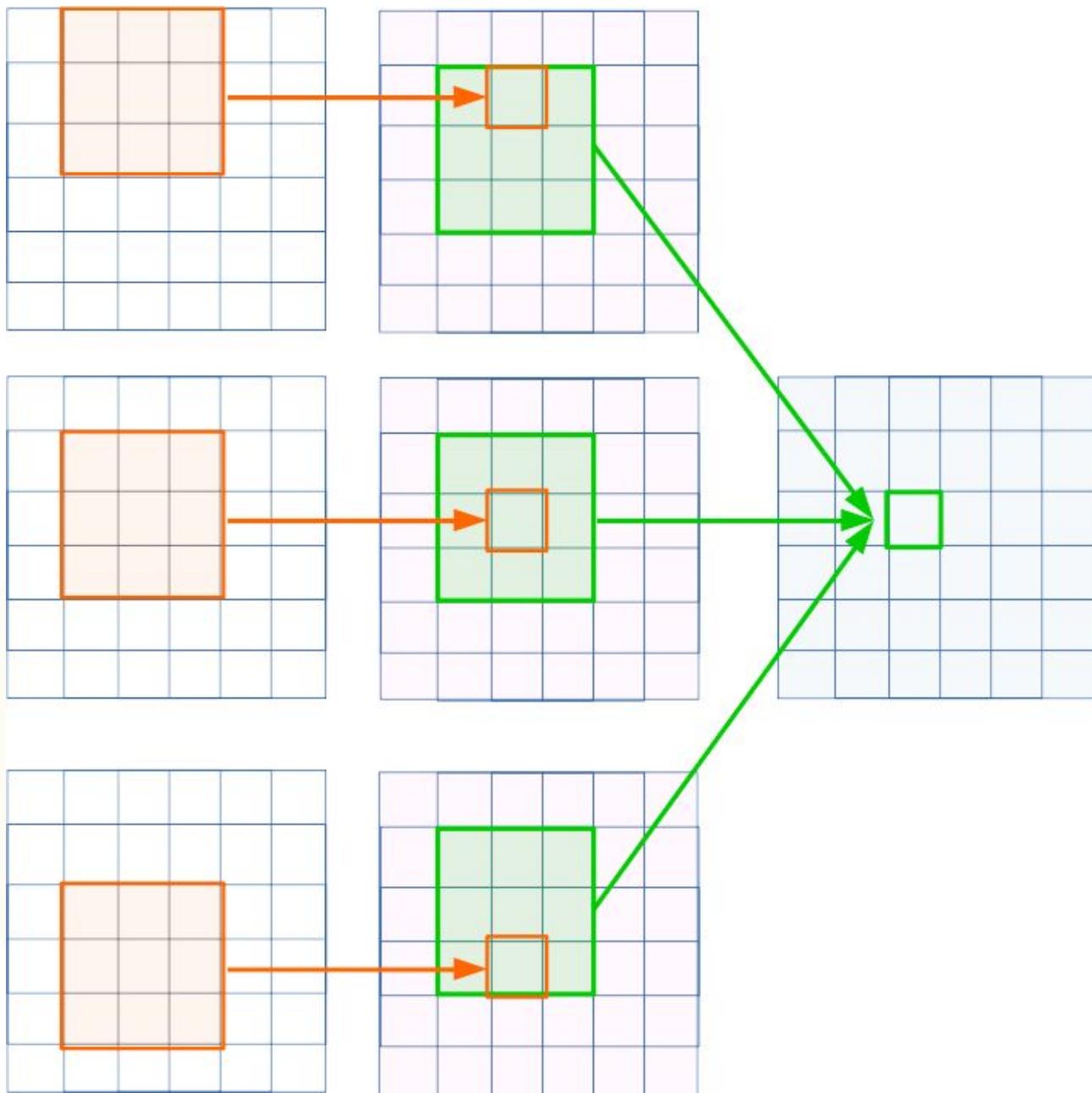


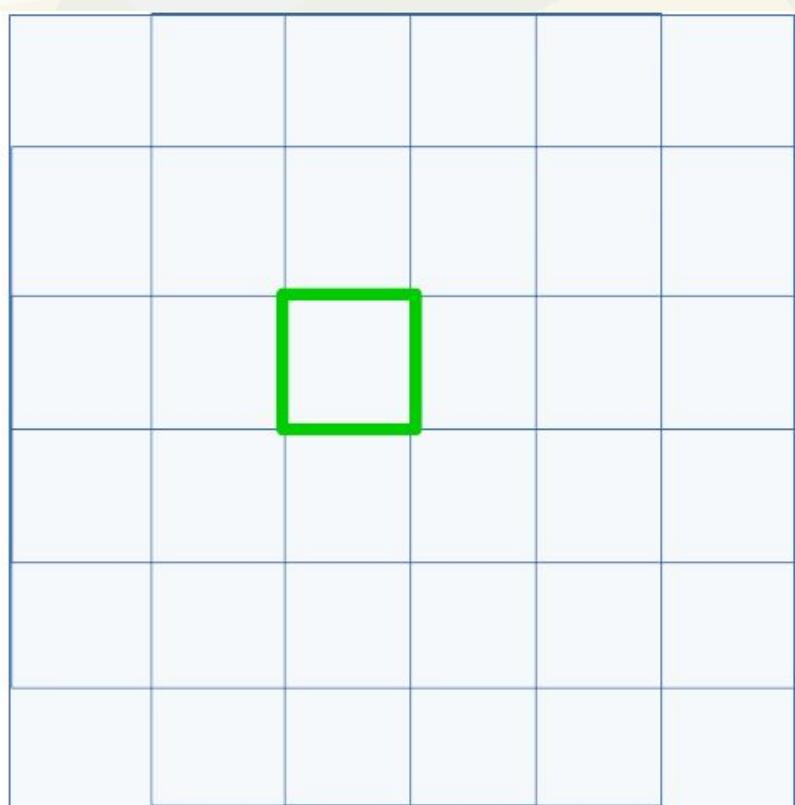
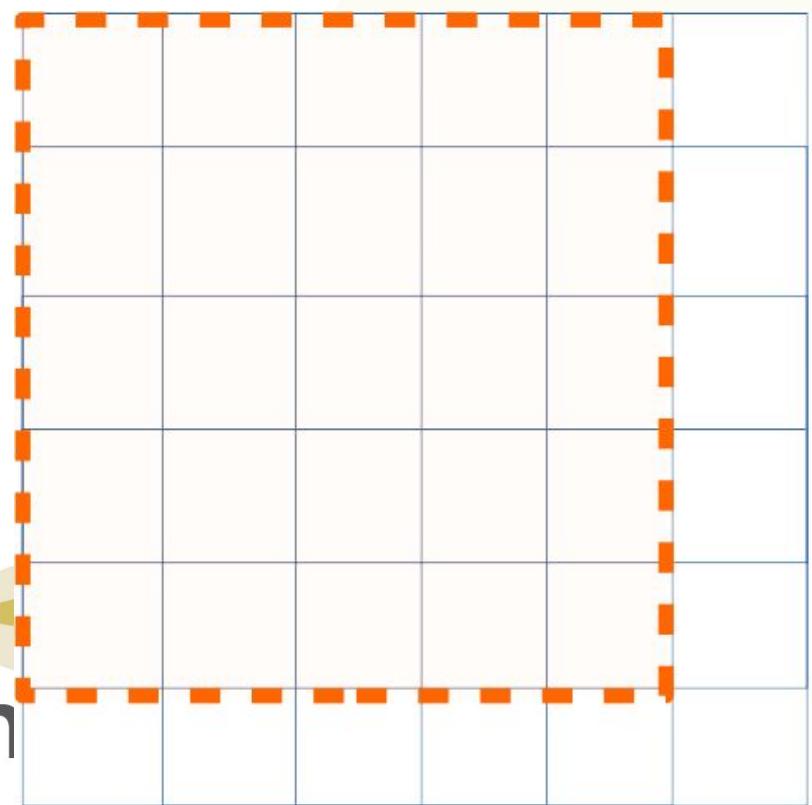
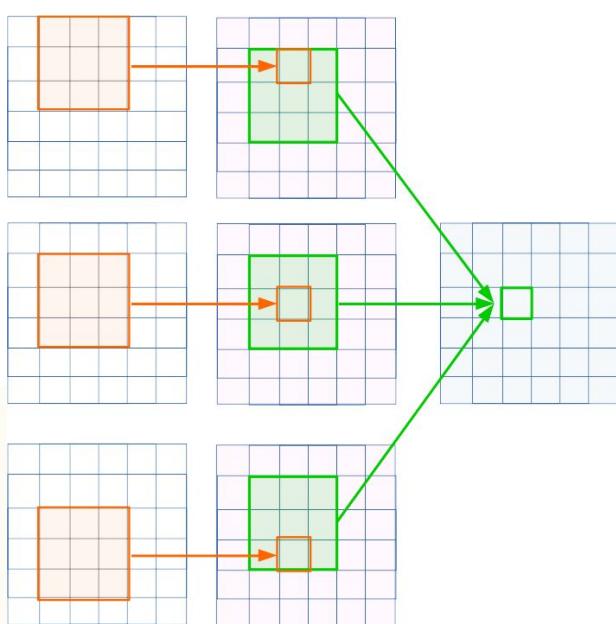


*“The **receptive field** of an individual sensory neuron is the particular region of the sensory space in which a stimulus will modify the firing of that neuron.”*

[https://en.wikipedia.org/wiki/Receptive\\_field](https://en.wikipedia.org/wiki/Receptive_field)

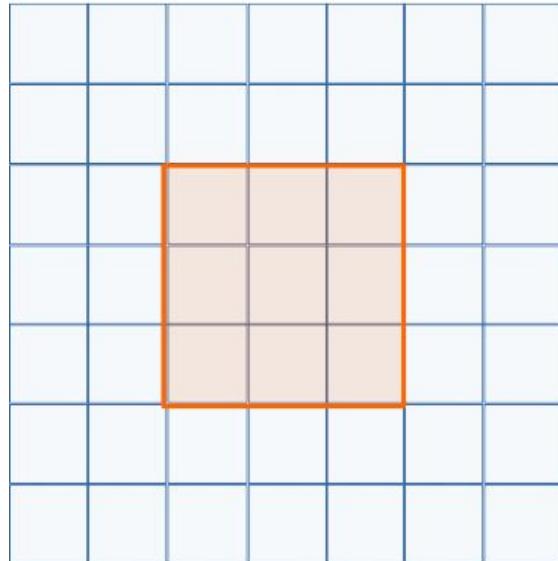




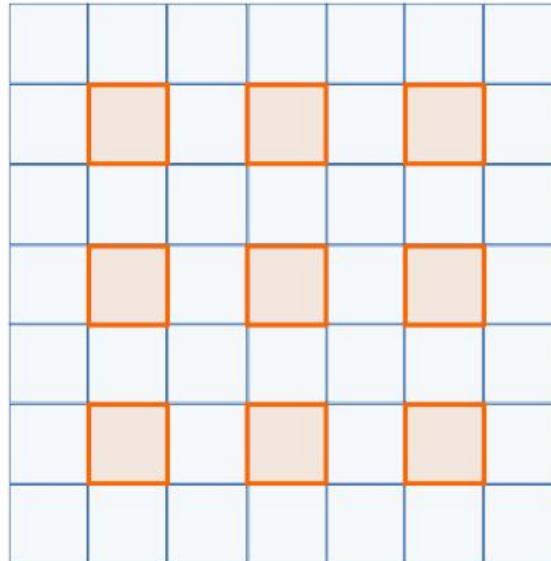


# Dilations

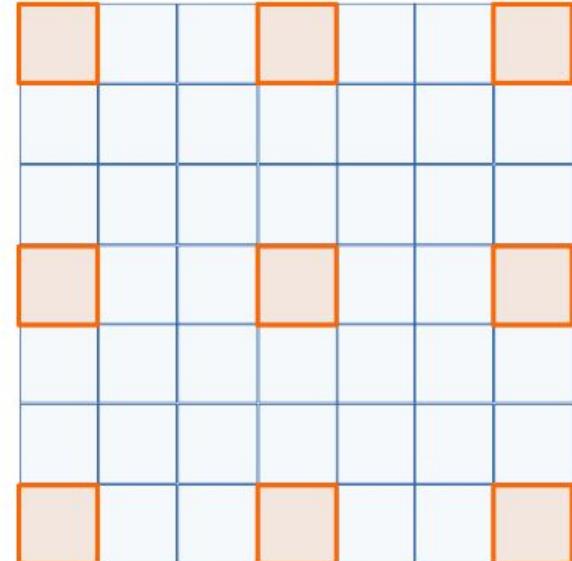
Dilation=1, (3 x 3)



Dilation=2, (3 x 3)



Dilation=3, (3 x 3)



- Increase receptive field without increasing parameters



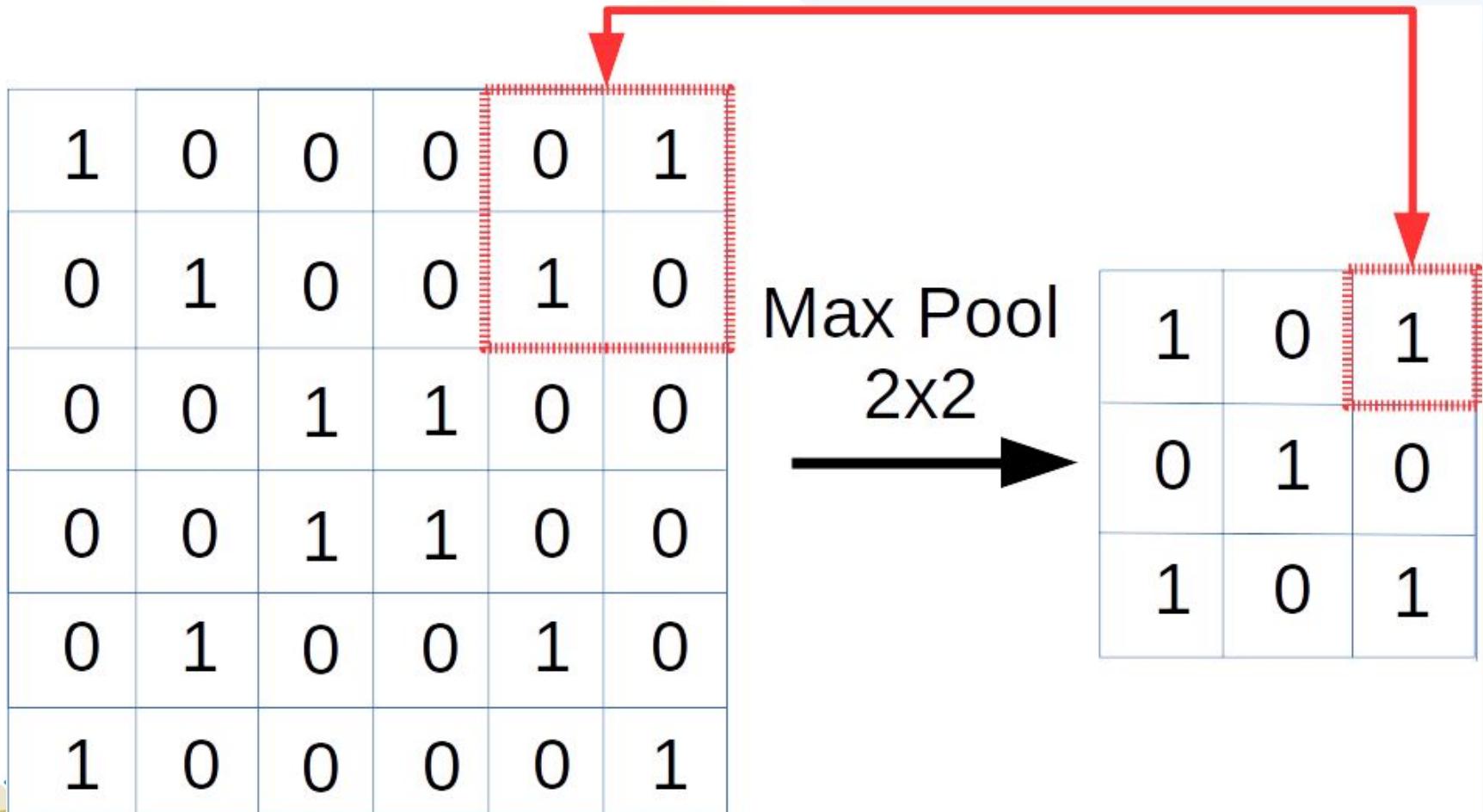
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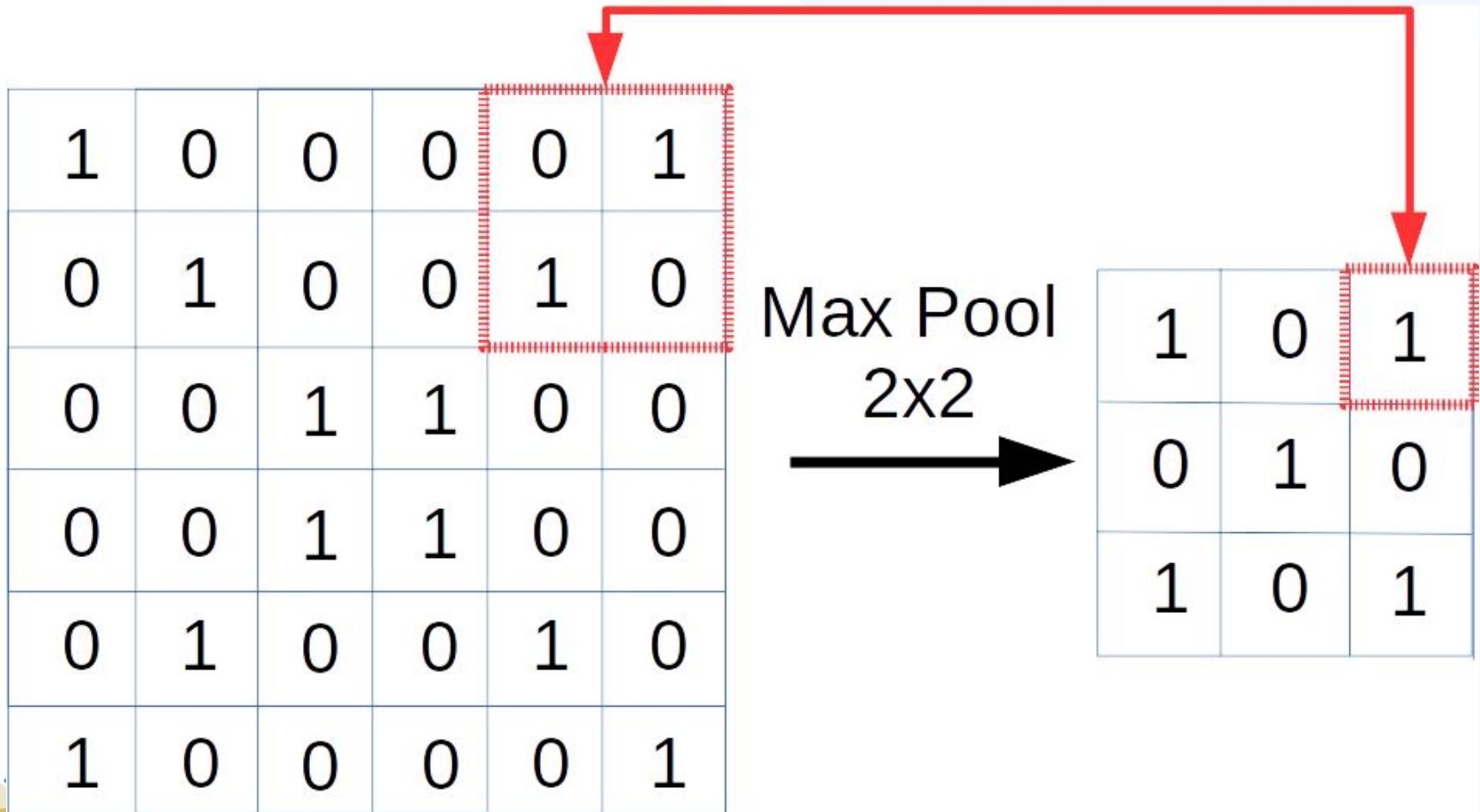


# Downsampling



*Summarizes essential features of image at lower spatial resolution*

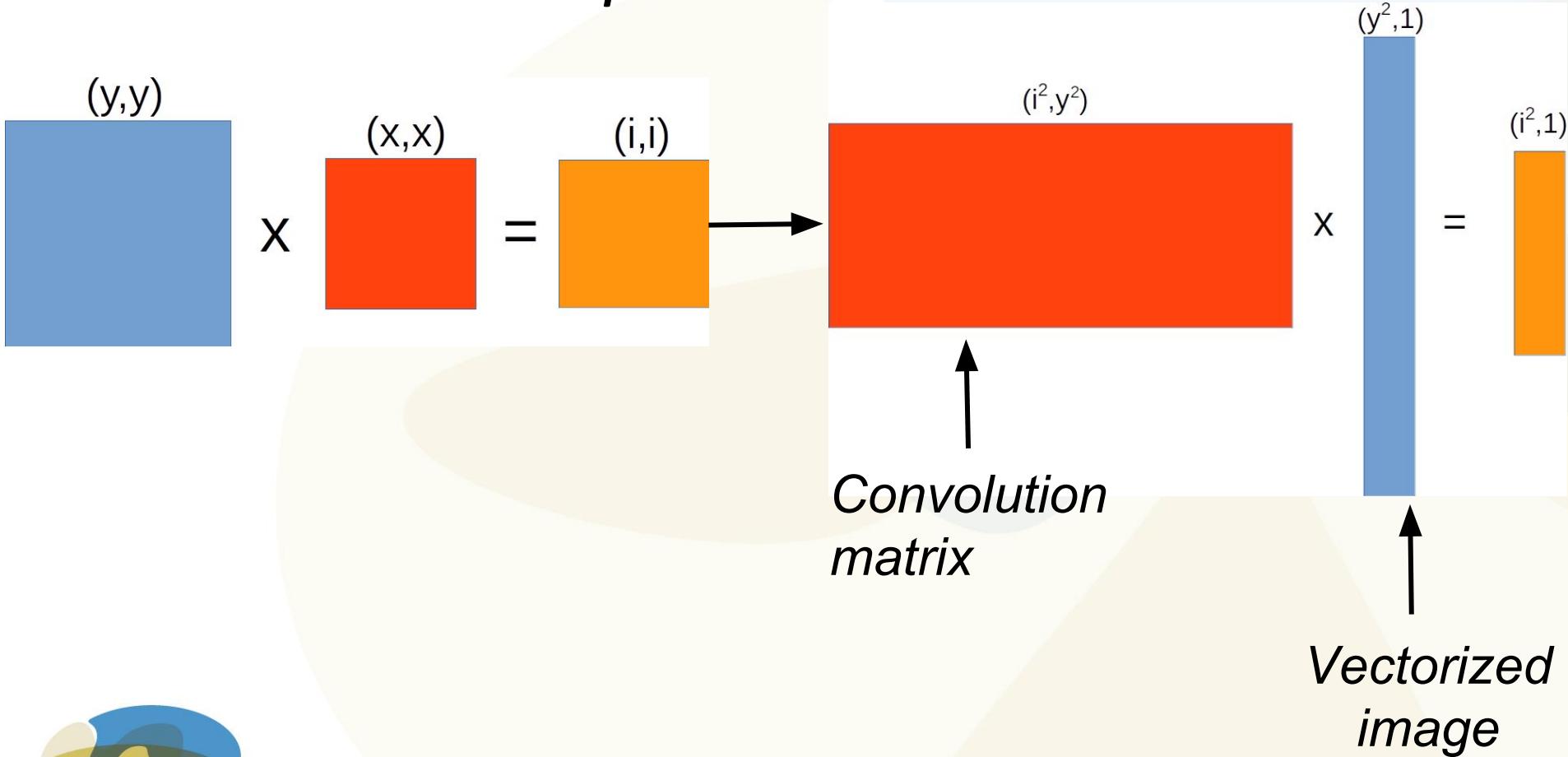
# Downsampling



*Decreases the image size by half!*

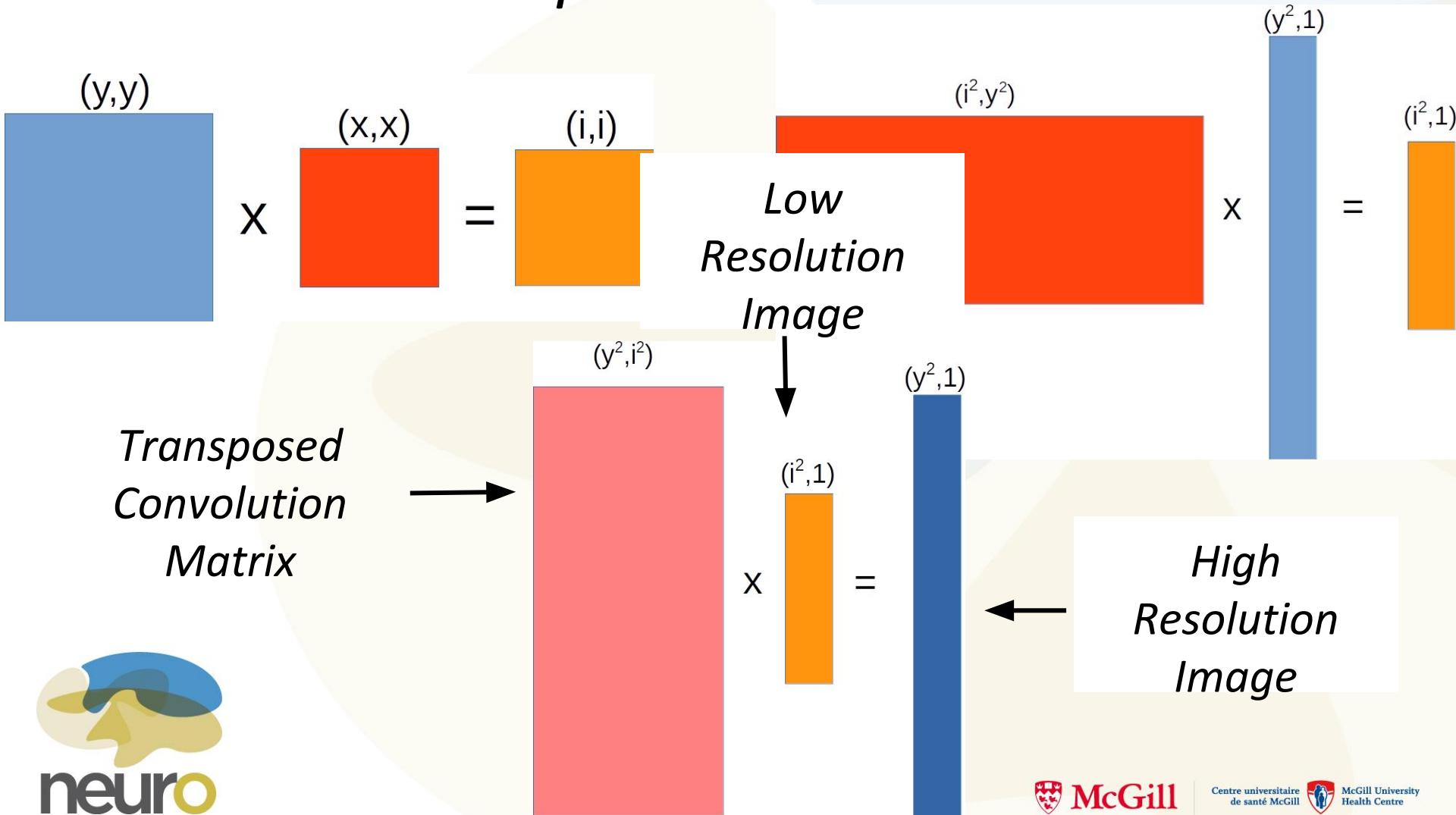
# Upsampling

## *Transpose Convolution*



# Upsampling

## *Transpose Convolution*

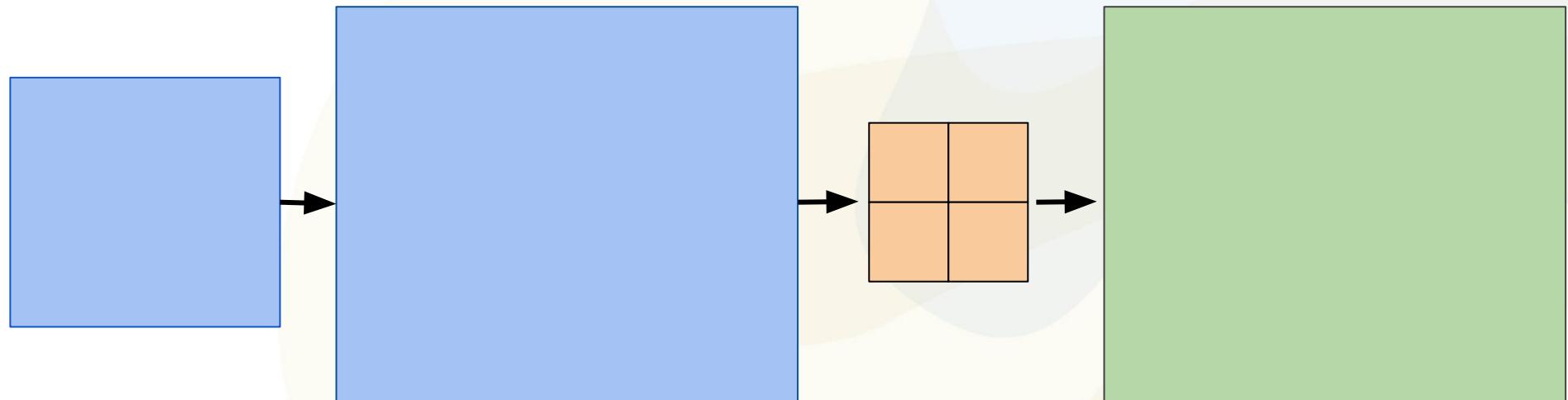


# Upsampling

*Interpolation + 2x2 Convolution*

Interpolation  
(nearest neighbours)

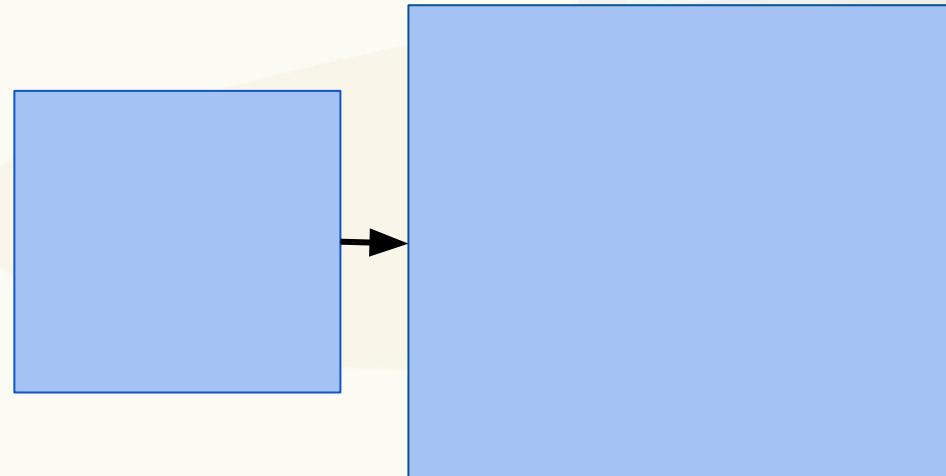
2x2 Convolution



# Upsampling

## *Interpolation*

Interpolation  
(nearest neighbours)

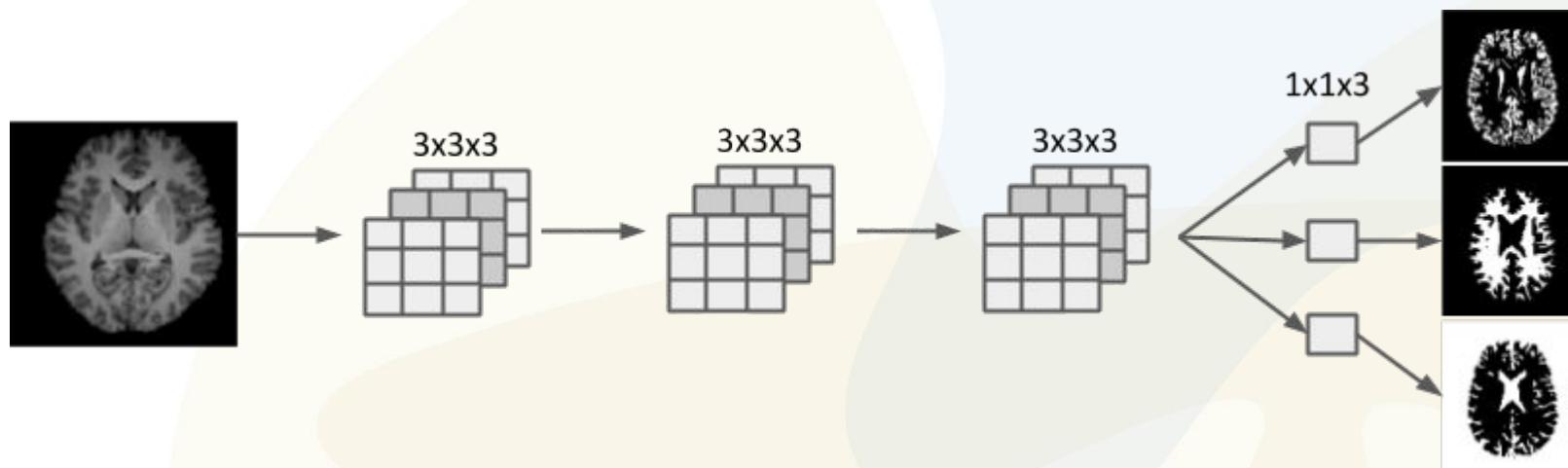


# Outline

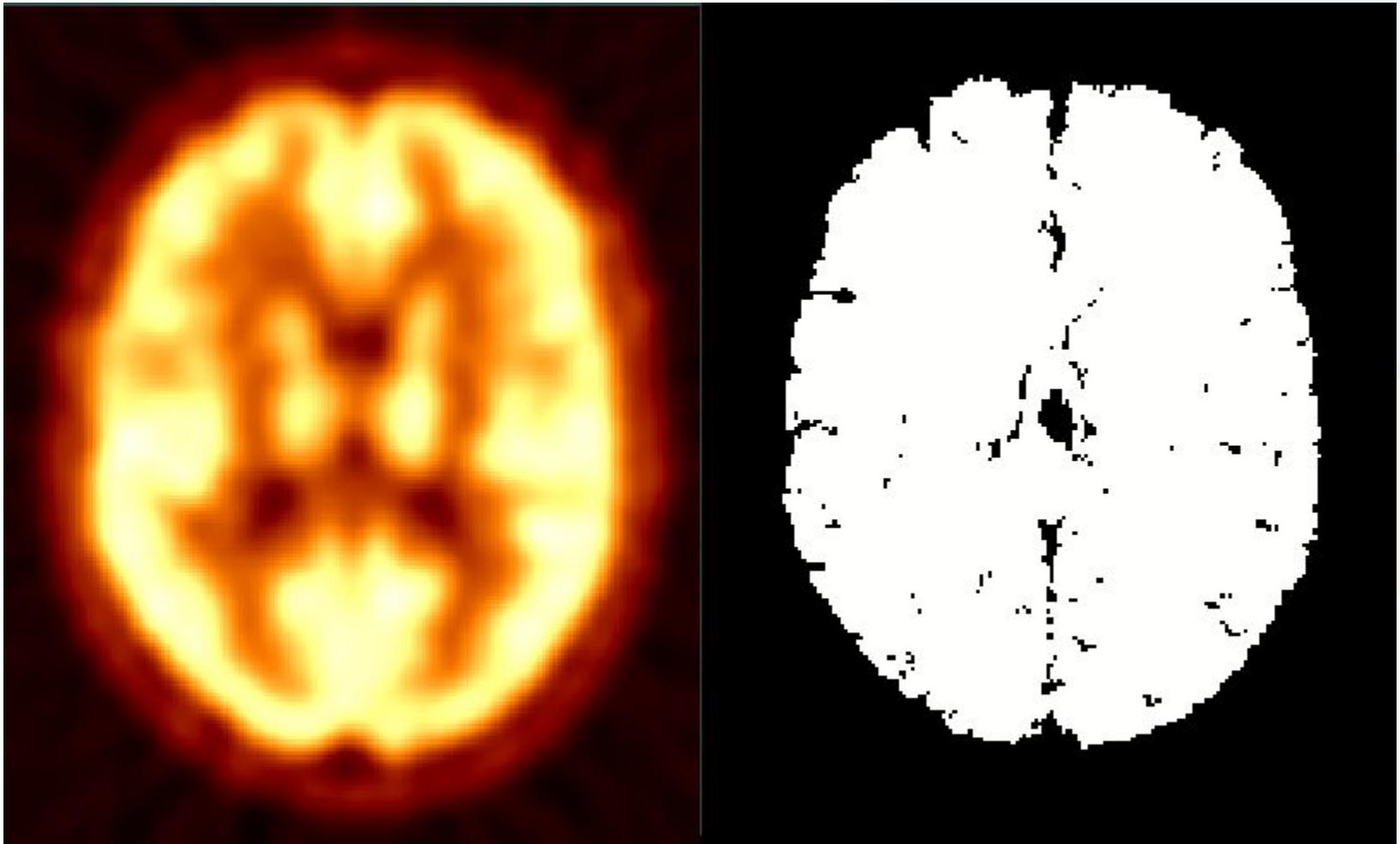
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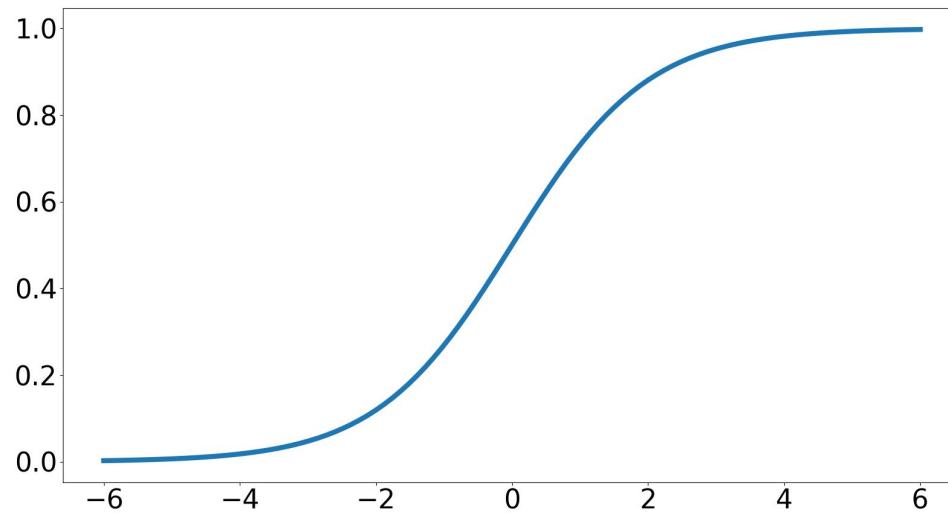
# Binary Classification



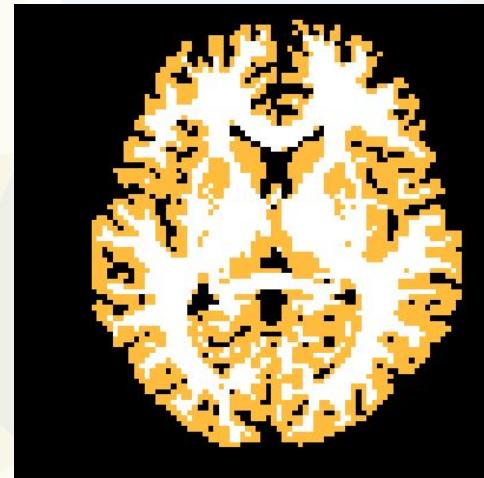
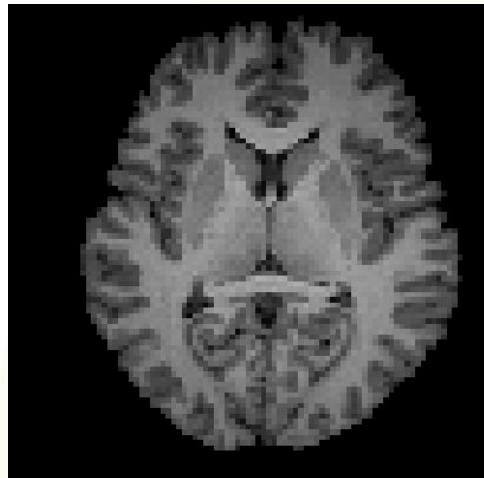
# Binary Classification

Sigmoid :

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



# Multi-category Classification



# Multi-category Classification

## Softmax :

- Generalization of binary sigmoid
- Creates pseudo-probability distribution

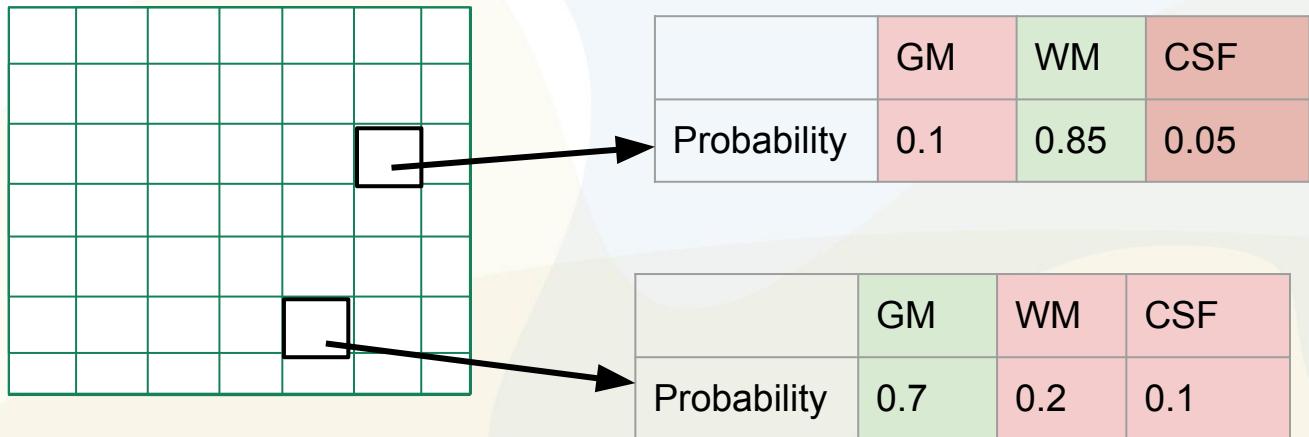
$$\sigma(x_j) = \frac{e^{x_j}}{\sum_i e^{x_i}}$$



# Softmax Output of Network

## Softmax

- Each pixel = [P(GM) P(WM), P(BG)]
- 3D output array = (Width, Height, 3)



- Transform to 2D image by finding class with max probability at each pixel



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# Choosing loss function

- **Cross Entropy :**

- From information theory
- Quantifies difference between two probability distributions

$$CE = \frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$



# Choosing loss function

## Cross Entropy :

$$CE = \frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$



Ground truth  
probability  
from labels



Bits of  
information  
for predicted  
label

# Cross Entropy

*True Distribution*

	GM	WM	CSF
<i>data_0</i>	1	0	0
<i>data_1</i>	0	1	0
<i>data_2</i>	0	1	0

*Predicted Distribution*

	GM	WM	CSF
<i>data_0</i>	.5	.2	.3
<i>data_1</i>	.2	.4	.4
<i>data_2</i>	.1	.5	.4

*Predicted distribution produced with softmax / sigmoid final activation*



# Cross Entropy

<i>True</i>			
	GM	WM	CSF
<i>data_0</i>	1	0	0
<i>data_1</i>	0	1	0
<i>data_2</i>	0	1	0

<i>Predicted</i>			
	GM	WM	CSF
<i>data_0</i>	.5	.2	.3
<i>data_1</i>	.2	.4	.4
<i>data_2</i>	.1	.5	.4

$$CE = \frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$

$$1 \times -\log_2(0.5) + 0 \times \log_2(.2) + 0 \times \log_2(.3) +$$

# Cross Entropy

		<i>True</i>				
		GM	WM	CSF		
data_0	1	0	0			
	0	1	0			
	0	1	0			
		<i>Predicted</i>				
		GM	WM	CSF		
data_0	.5	.2	.3			
	.2	.4	.4			
	.1	.5	.4			

$$CE = \frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$

$$1 \times -\log_2(0.5) + 0 \times \log_2(.2) + 0 \times \log_2(.3) + \\ 0 \times -\log_2(0.2) + 1 \times \log_2(.4) + 0 \times \log_2(.4) + \\ 0 \times -\log_2(0.1)$$

# Cross Entropy

<i>True</i>			
	GM	WM	CSF
<i>data_0</i>	1	0	0
<i>data_1</i>	0	1	0
<i>data_2</i>	0	1	0

<i>Predicted</i>			
	GM	WM	CSF
<i>data_0</i>	.5	.2	.3
<i>data_1</i>	.2	.4	.4
<i>data_2</i>	.1	.5	.4

$$CE = \frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$

$$\begin{aligned} & 1 \times -\log_2(0.5) + 0 \times \log_2(.2) + 0 \times \log_2(.3) + \\ & 0 \times -\log_2(0.2) + 1 \times \log_2(.4) + 0 \times \log_2(.4) + \\ & 0 \times -\log_2(0.1) + 1 \times \log_2(.5) + 0 \times \log_2(.4) \end{aligned}$$

# Cross Entropy

	<i>True</i>			<i>Predicted</i>			
	GM	WM	CSF		GM	WM	CSF
<i>data_0</i>	1	0	0	<i>data_0</i>	.5	.2	.3
<i>data_1</i>	0	1	0	<i>data_1</i>	.2	.4	.4
<i>data_2</i>	0	1	0	<i>data_2</i>	.1	.5	.4

$$CE = -\frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$

$$\begin{aligned} & 1 \times -\log_2(0.5) + 0 \times \log_2(.2) + 0 \times \log_2(.3) + \\ & 0 \times -\log_2(0.2) + 1 \times \log_2(.4) + 0 \times \log_2(.4) + \\ & 0 \times -\log_2(0.1) + 1 \times \log_2(.5) + 0 \times \log_2(.4) \\ & = 1 \times -\log_2(0.5) + 1 \times \log_2(.4) + 1 \times \log_2(.4) \\ & = 3.64 \end{aligned}$$



# Cross Entropy

<i>True</i>			
	GM	WM	CSF
<i>data_0</i>	1	0	0
<i>data_1</i>	0	1	0
<i>data_2</i>	0	1	0

<i>Predicted</i>			
	GM	WM	CSF
<i>data_0</i>	.8	.1	.1
<i>data_1</i>	.1	.8	.1
<i>data_2</i>	.1	.8	.1

# Cross Entropy

	True			Predicted		
	GM	WM	CSF	GM	WM	CSF
<i>data_0</i>	1	0	0	.8	.1	.1
<i>data_1</i>	0	1	0	.1	.8	.1
<i>data_2</i>	0	1	0	.1	.8	.1

$$CE = -\frac{1}{N} \sum_i^N \sum_c^C True_i^c \times \log_2(Predicted_i^c)$$

$$\begin{aligned} &= 1 \times -\log_2(0.8) + 1 \times \log_2(.8) + 1 \times \log_2(.8) \\ &= 0.97 \end{aligned}$$

*By improving our prediction, we've decreased the value of the cross entropy function*



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# Metrics

1. Classification labels are integers
2. Dice Metric
  - a. Binary classification
  - b. Custom metric, not included by default in Keras
  - c. Perfect overlap =  $(2 \times |X \cap Y|) / (|X| + |Y|) = 1$

$$\frac{2|X \cap Y|}{|X| + |Y|}$$



# Metrics

1. Classification labels are integers
  - a. → Can't use MSE or similar metrics
2. Dice Metric
3. Categorical Accuracy
  - a. Binary and multi-class labels
  - b. Use 'acc' in Keras

$$\frac{1}{N} \sum_i I(Predicted_i = Label_i)$$



# Metrics

1. Classification labels are integers
2. Dice Metric
3. Categorical Accuracy
4. Baseline for metrics is not usually not 0
  - a. Ex. Guess all 0 → metric = 0.6



# Outline

## 1. Basic Concepts

- a. Kernels
- b. Receptive field & dilations
- c. Upsampling & downsampling
- d. Final activation functions
- e. Loss functions
- f. Metrics
- g. Cross-validation
- h. Feature extraction



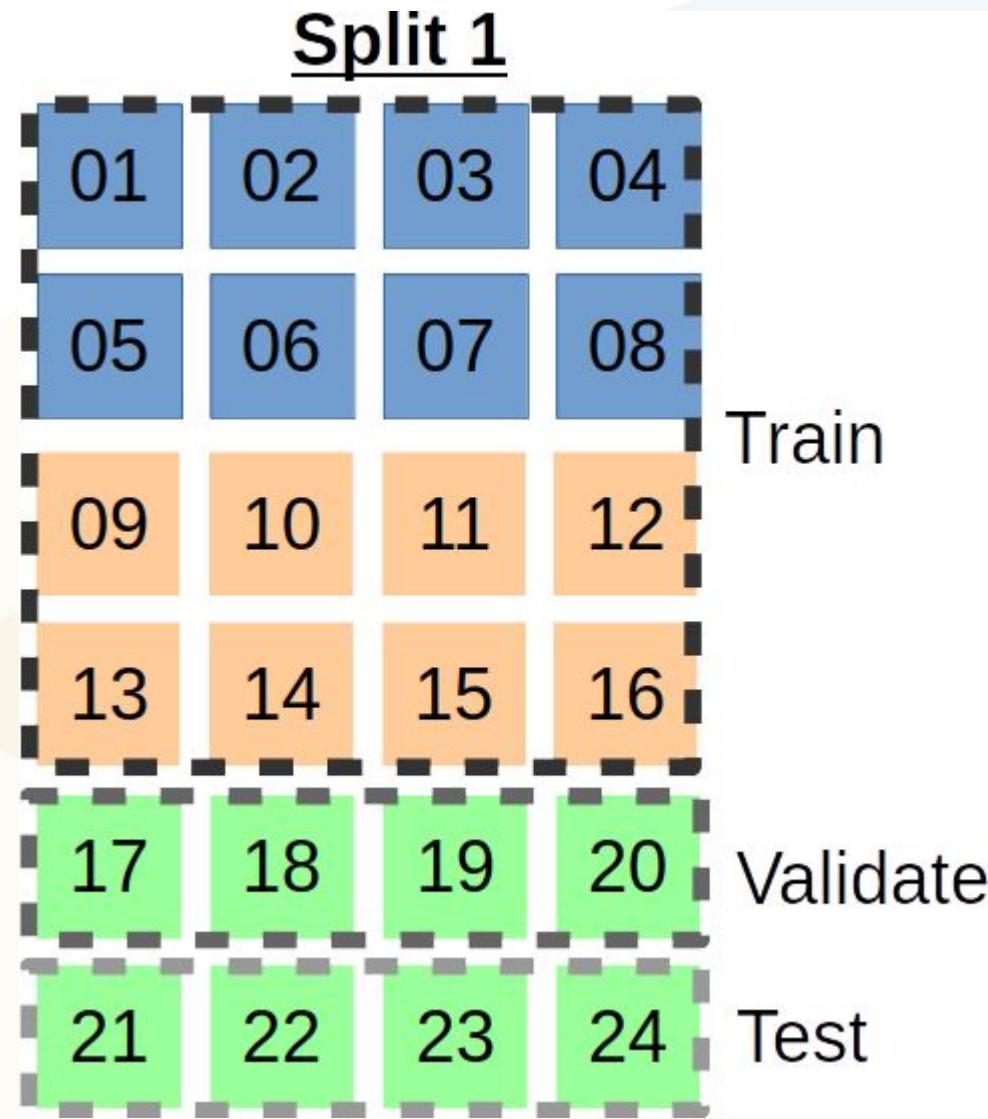
## K-folds cross validation

01	02	03	04	05	06	07	08	09	10	11	12	13	14	15	16
01	02	03	04	05	06	07	08	09	10	11	12	13	14	15	16
01	02	03	04	05	06	07	08	09	10	11	12	13	14	15	16
01	02	03	04	05	06	07	08	09	10	11	12	13	14	15	16

# Data

01	02	03	04
05	06	07	08
09	10	11	12
13	14	15	16
17	18	19	20
21	22	23	24

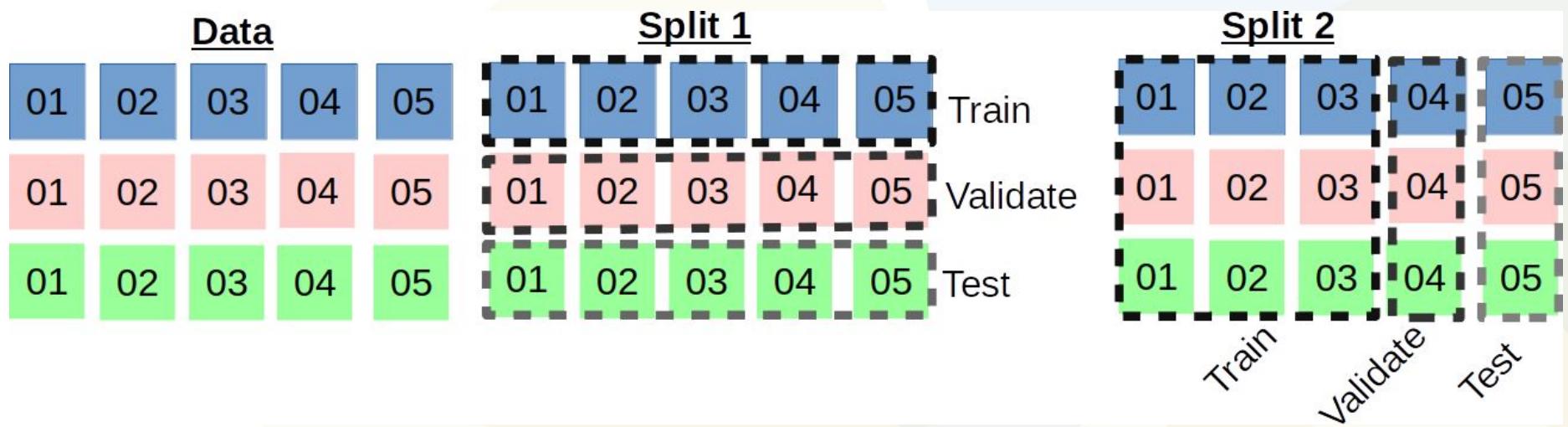




## Split 2

01	02	03		
05	06	07		
09	10	11		
13	14	15	12	16
17	18	19	04	08
21	22	23	24	20





# Outline

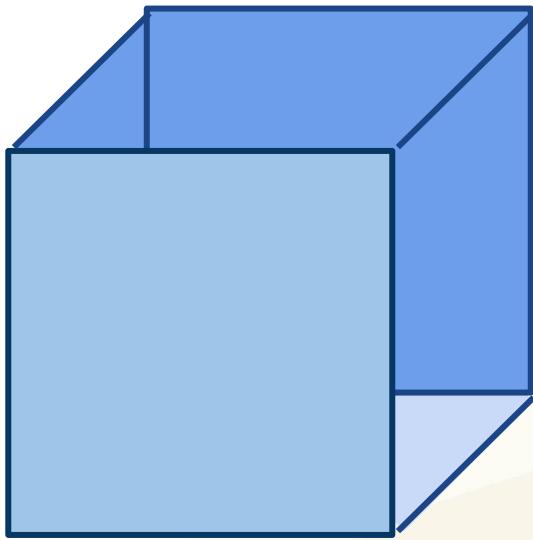
## 1. Basic Concepts

- a. Kernels
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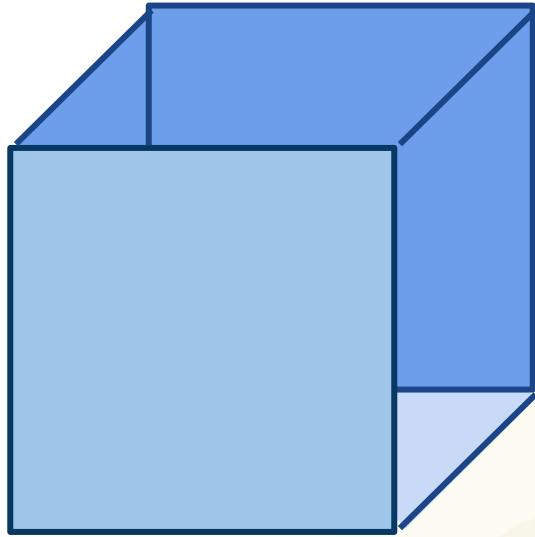
# Many ways to analyze volumes

Full 3D Volume

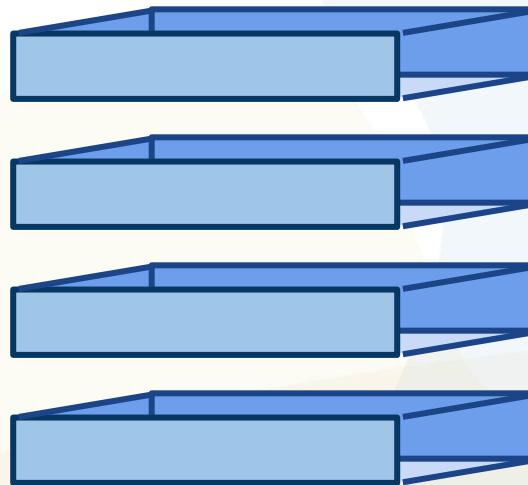


# Many ways to analyze volumes

Full 3D Volume

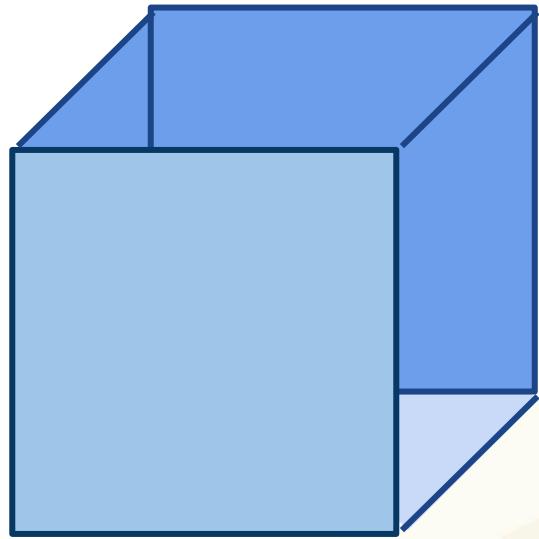


2D Slices

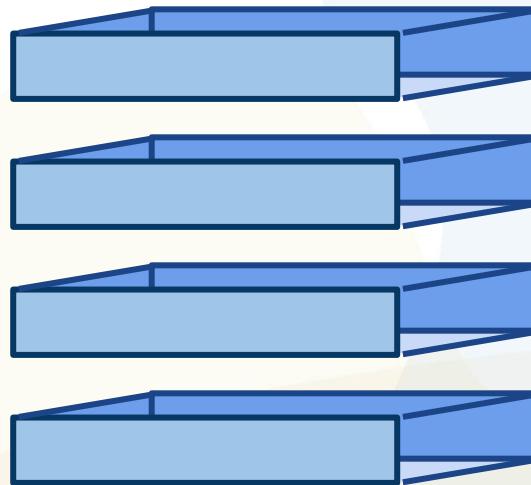


# Many ways to analyze volumes

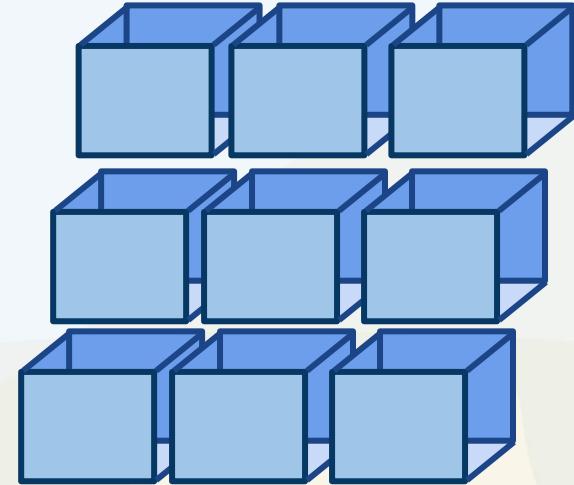
Full 3D Volume



2D Slices

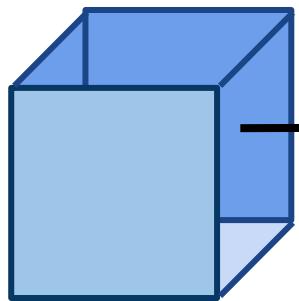


3D Patches

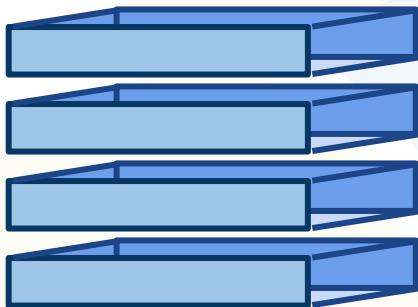


# Formatting data to feed into Keras

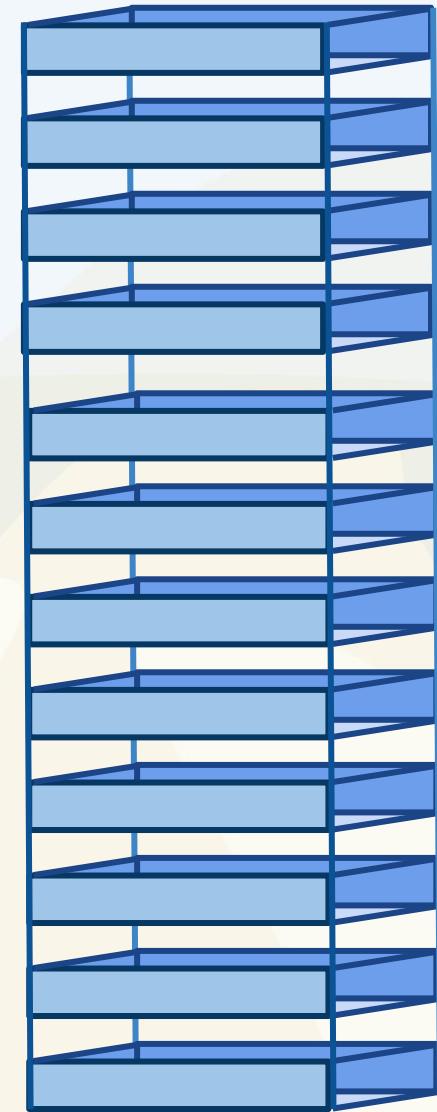
3D Nifti / MINC



2D Slices



HDF5 / NPY



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  - b. “Deep Learning” <http://neuralnetworksanddeeplearning.com/chap6.html> , Michael Nielsen
2. Kernels
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3. Softmax
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4. Cross Entropy
  - a. “A Short Introduction to Entropy, Cross-Entropy and KL-Divergence”, Aurélien Géron.  
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  - b. A Friendly Introduction to Cross-Entropy Loss, Rob DiPietro.  
<https://rdipietro.github.io/friendly-intro-to-cross-entropy-loss/>.

