

Lecture 6: Electronic Health Records Introduction

Announcements

- Upcoming deadlines:
 - A1 due Tue 10/6
 - Project proposal due Fri 10/9

What are electronic health records?

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Patient Timeline

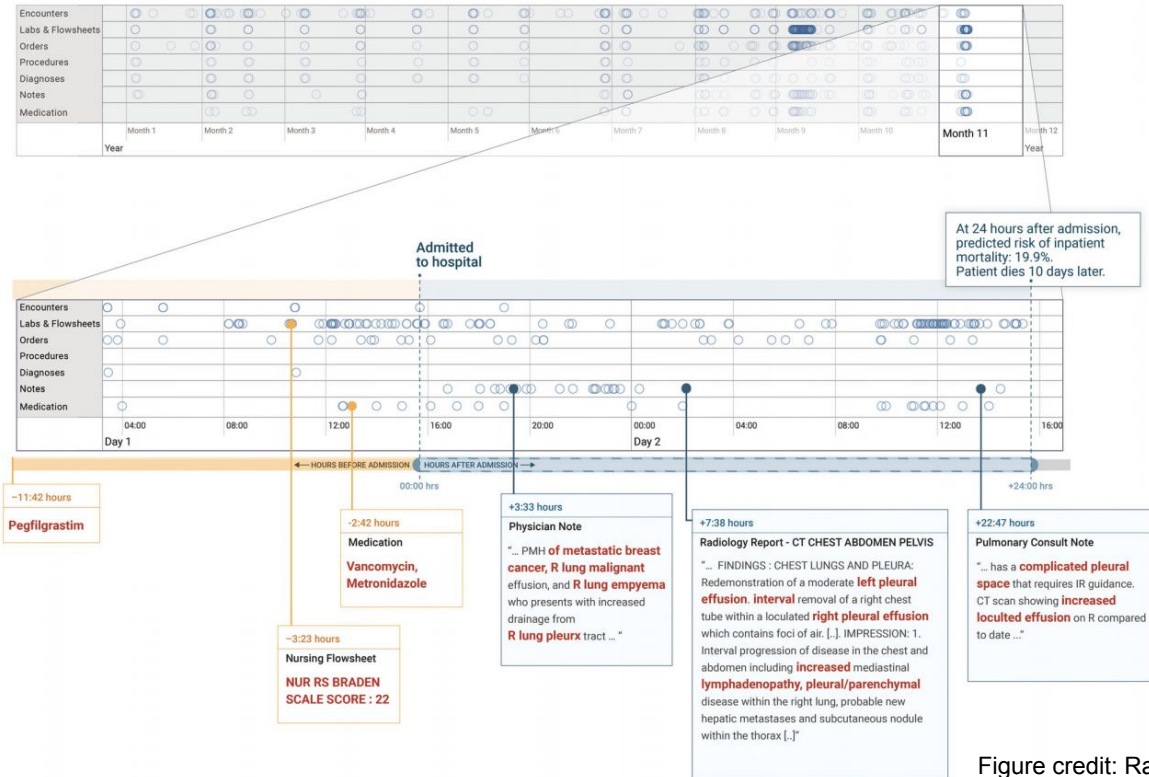


Figure credit: Rajkomar et al. 2018

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Stores patient information over time

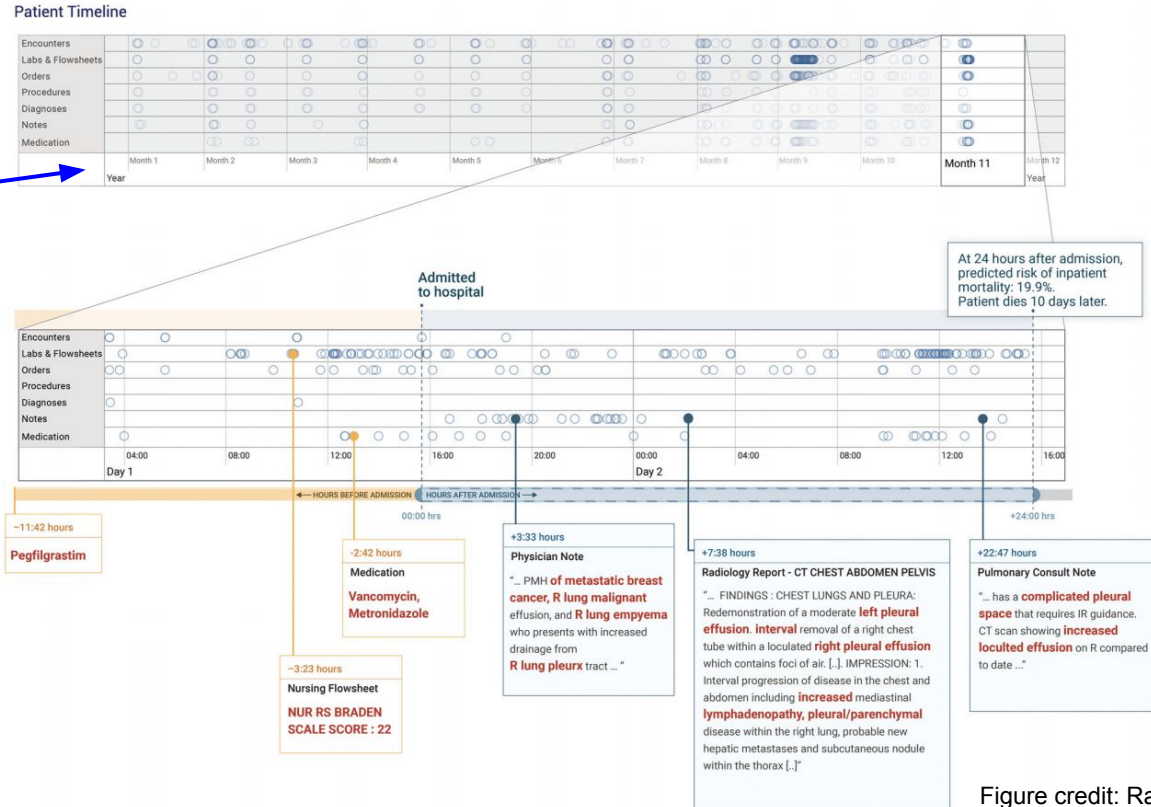
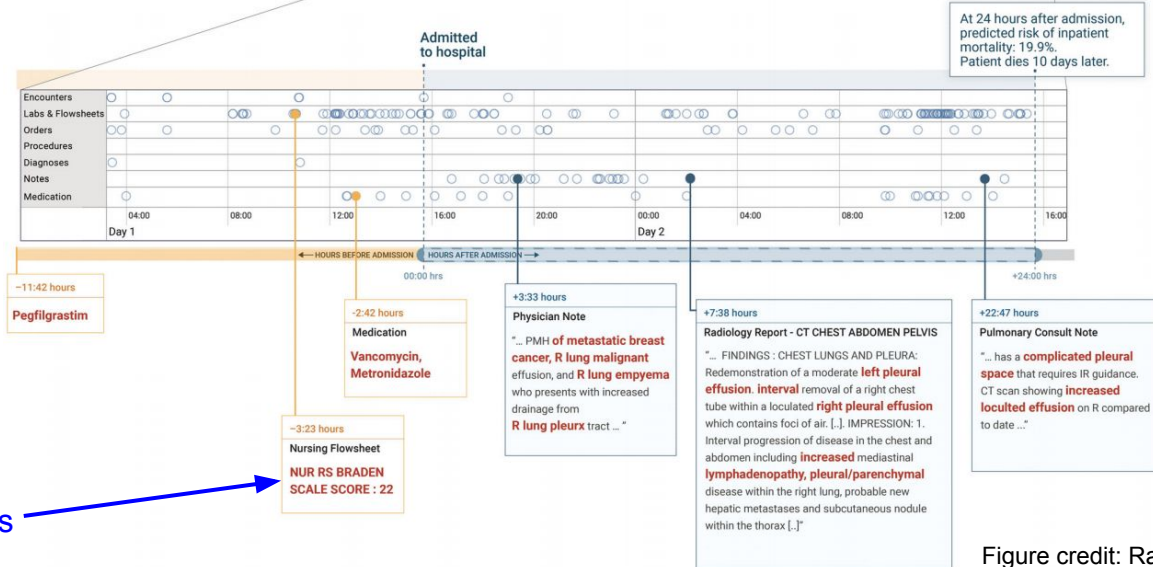
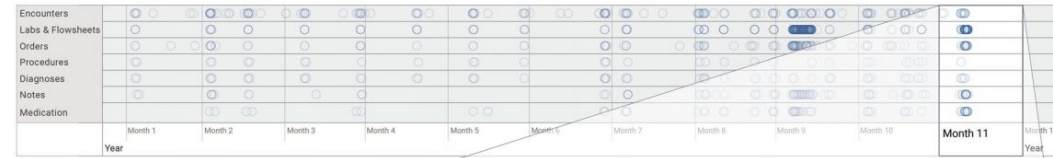


Figure credit: Rajkomar et al. 2018

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Patient Timeline



Nursing notes

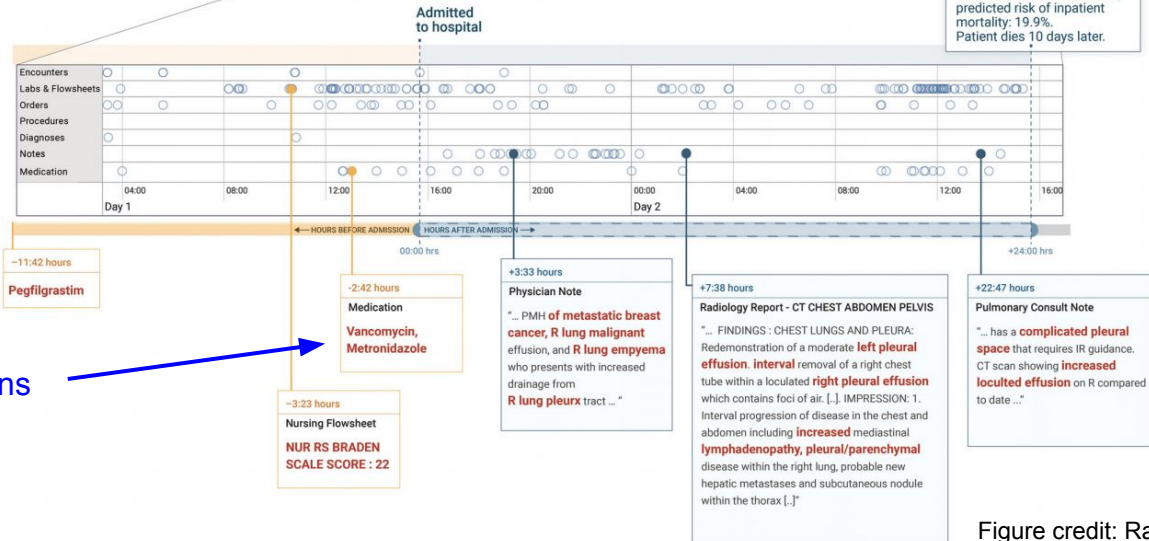
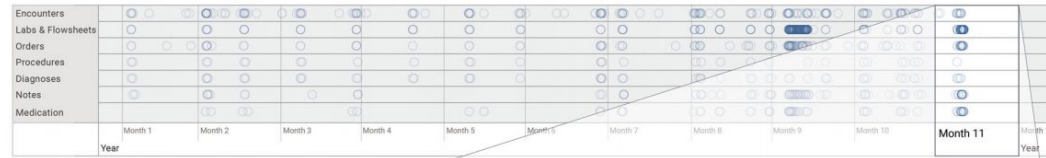


Figure credit: Rajkomar et al. 2018

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Patient Timeline



Medications

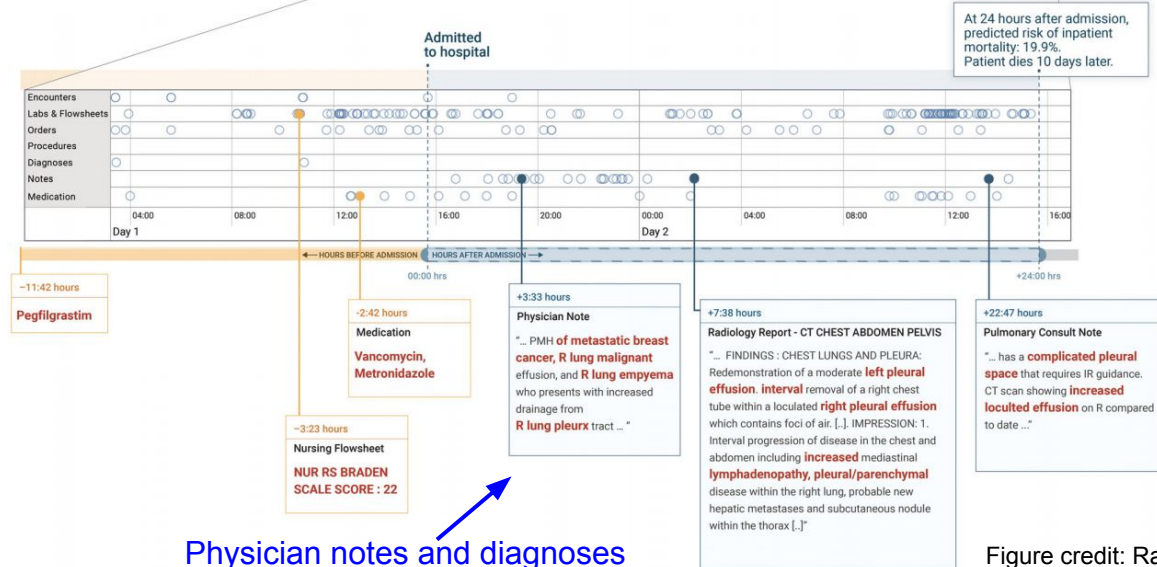
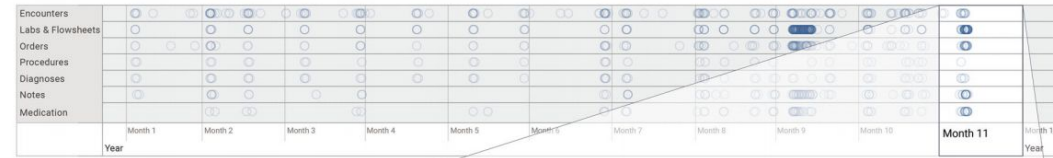


Figure credit: Rajkomar et al. 2018

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Patient Timeline



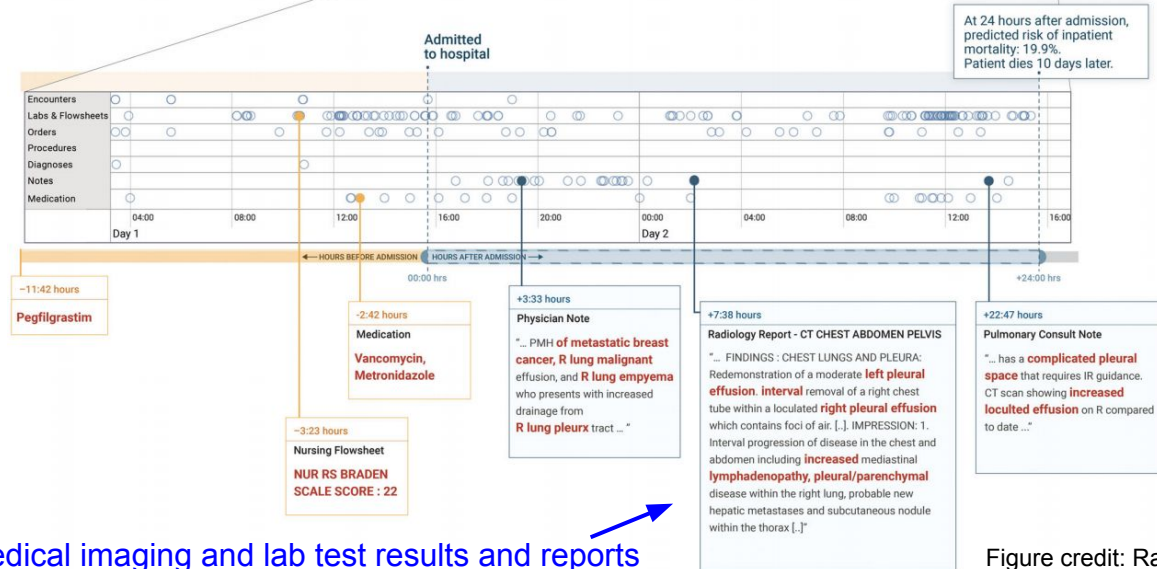
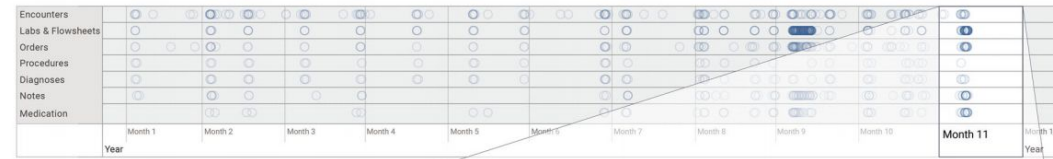
Physician notes and diagnoses

Figure credit: Rajkomar et al. 2018

What are electronic health records?

Patient chart in digital form, containing medical and treatment history

Patient Timeline

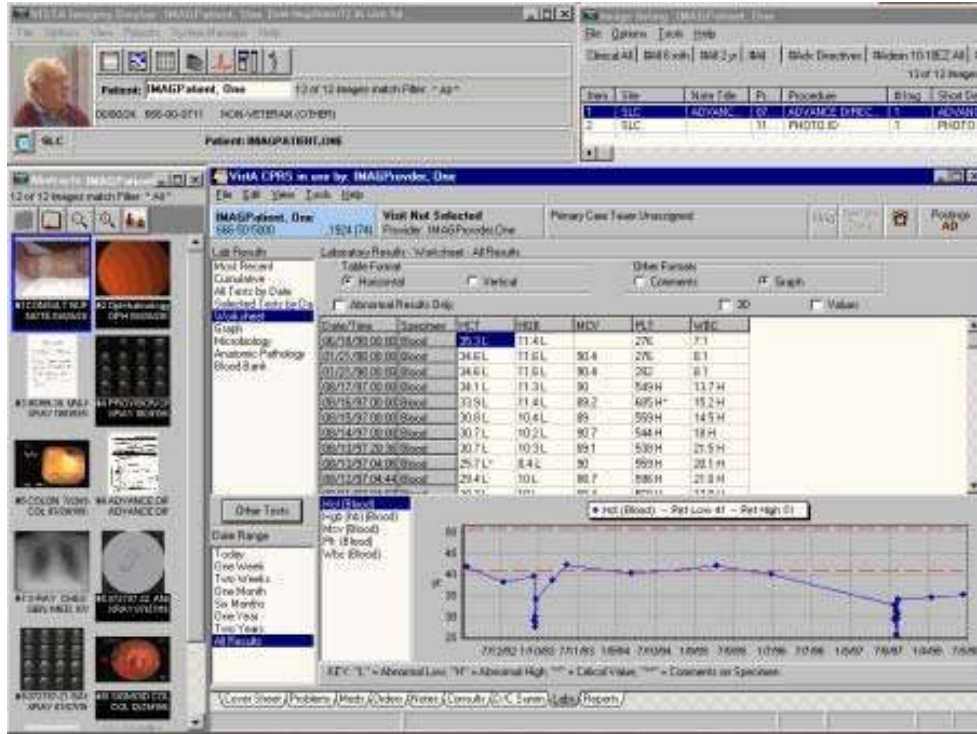


At 24 hours after admission, predicted risk of inpatient mortality: 19.9%. Patient dies 10 days later.

Medical imaging and lab test results and reports

Figure credit: Rajkomar et al. 2018

What are electronic health records?



1960s: invention
1980s: increased effort
2009: HITECH Act (Health Information Technology for Economic and Clinical Health Act) -- financial incentives for health care providers to adopt EHR

EHR adoption in the US (hospitals)

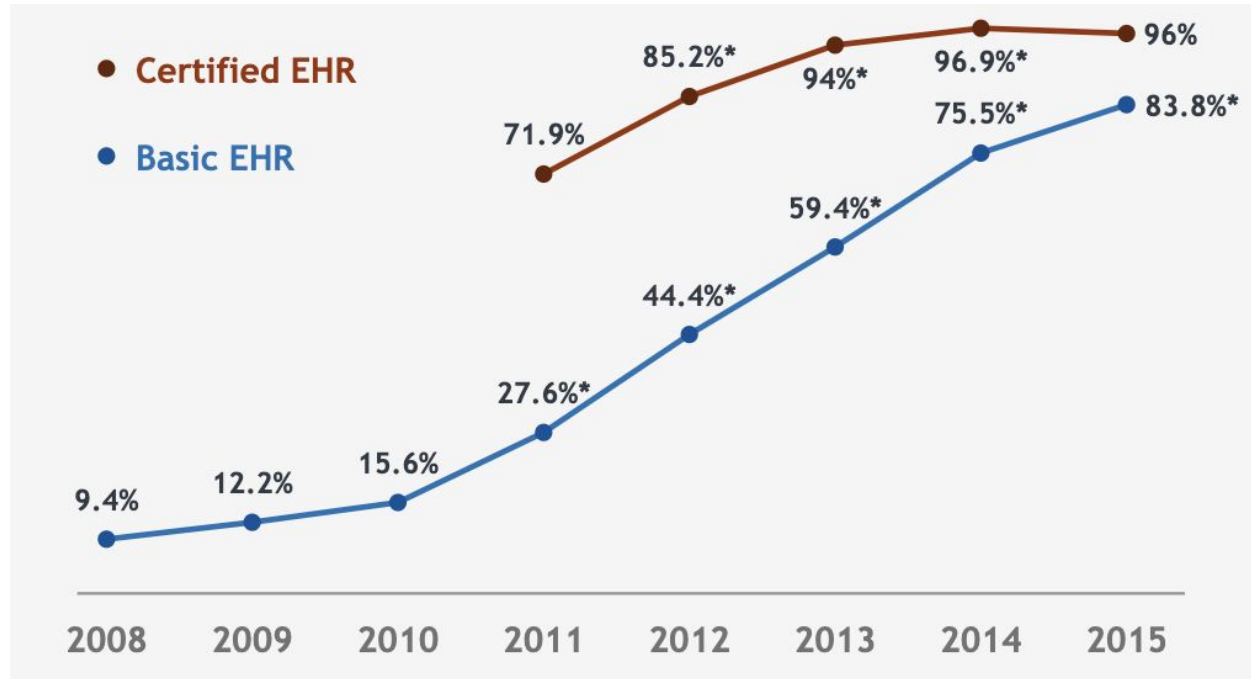


Figure credit: <https://dashboard.healthit.gov/evaluations/images/db-35-figure1.svg>

EHR adoption in the US (hospitals)

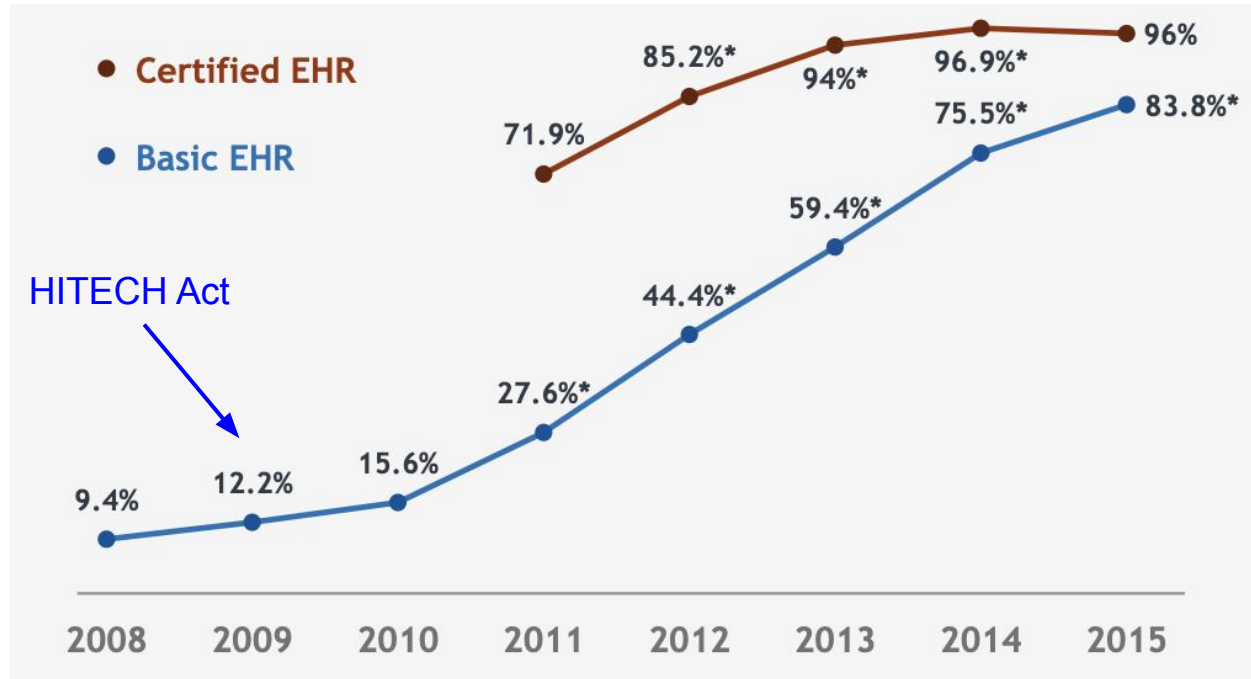


Figure credit: <https://dashboard.healthit.gov/evaluations/images/db-35-figure1.svg>

EHR adoption in the US (office-based physicians)

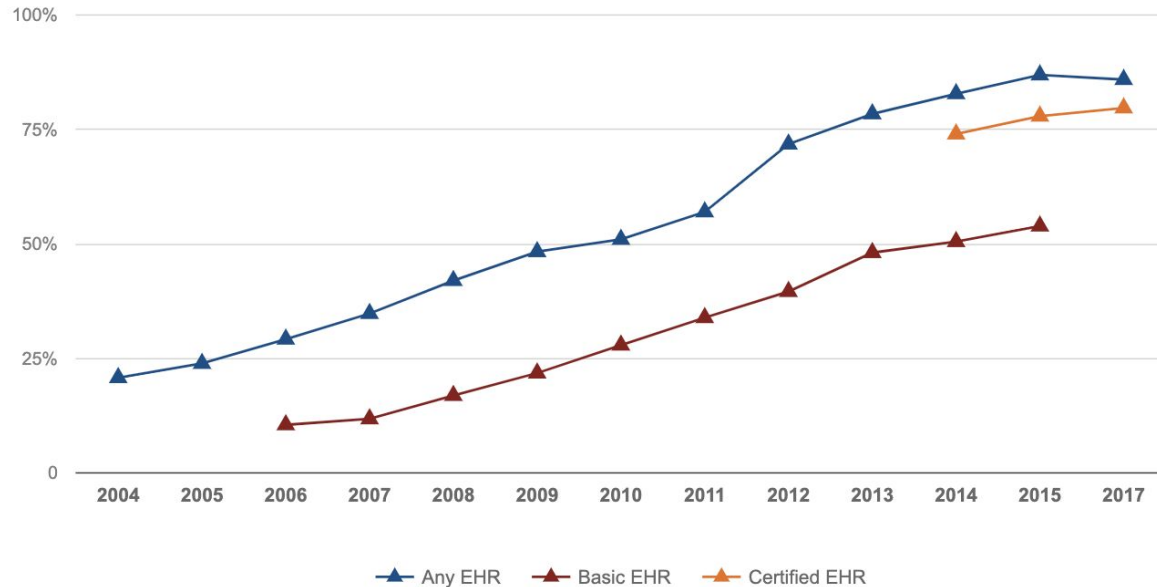
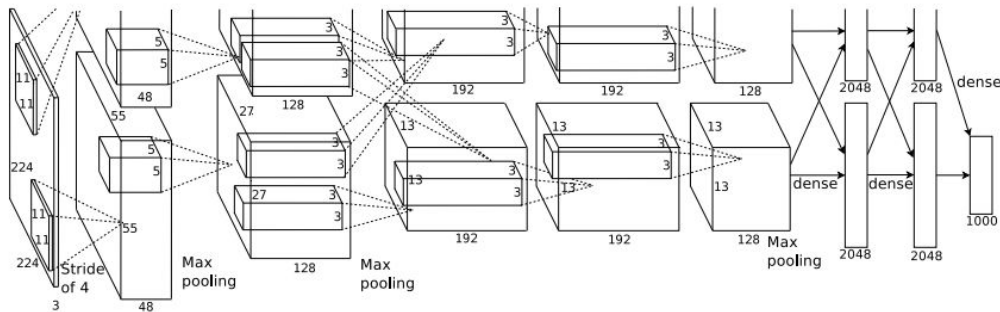


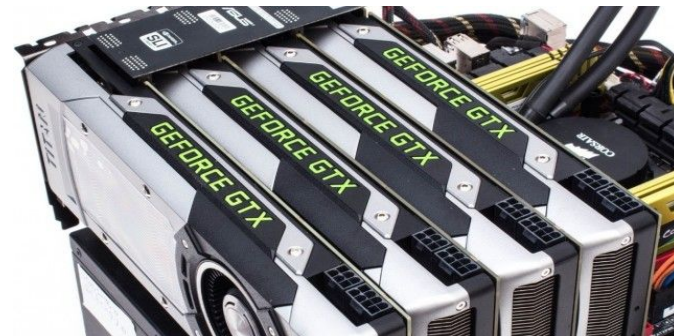
Figure credit: <https://dashboard.healthit.gov/quickstats/pages/physician-ehr-adoption-trends.php>

Convergence of key ingredients of deep learning

Algorithms



Compute

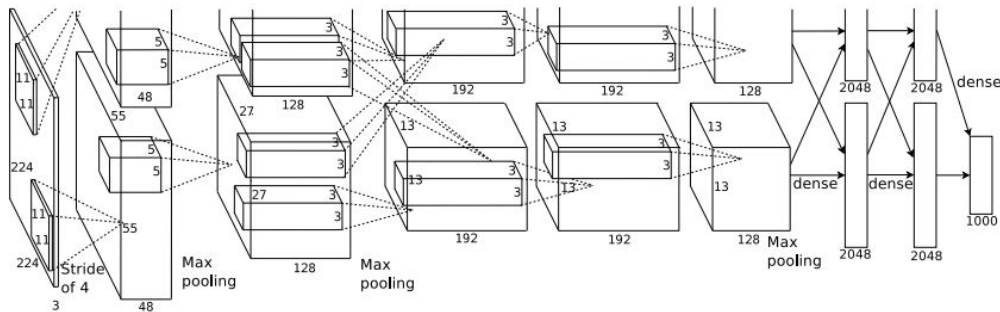


Data



Convergence of key ingredients of deep learning

Algorithms



Compute



Data



A real example of EHR data: MIMIC-III dataset

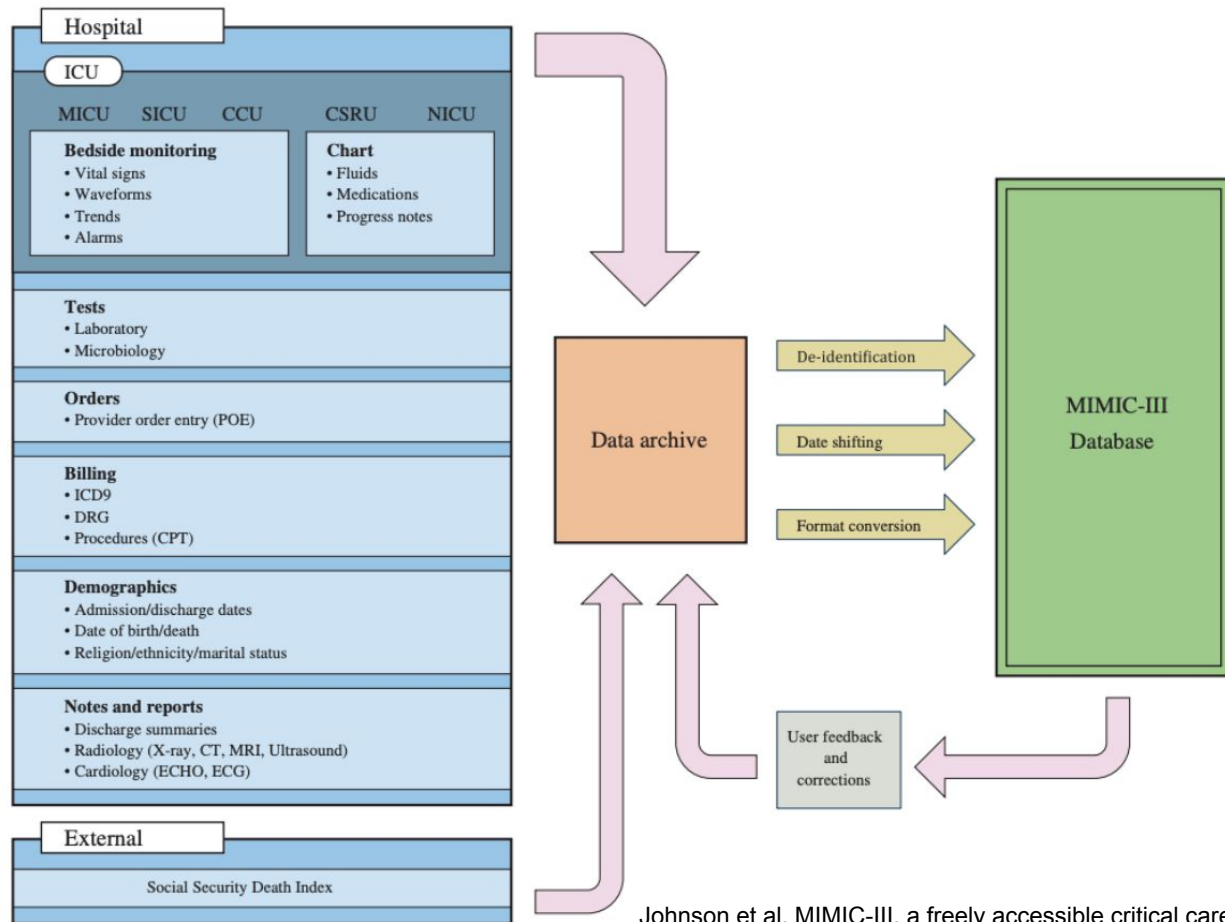
- Open source database of de-identified data for 38,597 adult patients, corresponding to 49,785 hospital admissions
- All patients admitted to critical care units at Beth Israel Deaconess Medical Center (Boston, MA) between 2001 - 2012
- Also 7870 neonates admitted between 2001-2008
- Median hospital stay length: 6.9 days
- Median ICU stay length: 2.1 days
- In-hospital mortality: 11.5%
- Mean of 4579 charted observations and 380 laboratory measurements for each admission

Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.

A real example of EHR data: MIMIC-III dataset

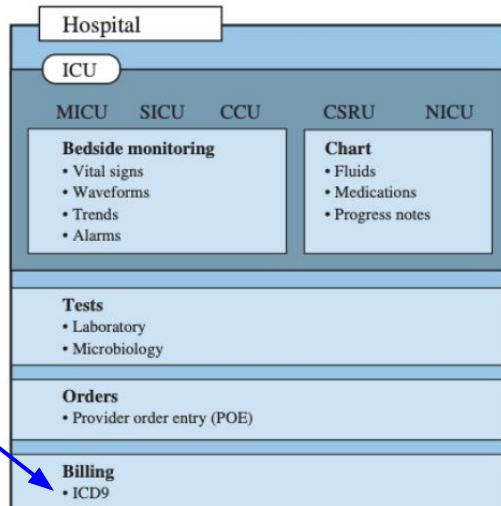
Critical care unit	CCU	CSRU	MICU	SICU	TSICU	Total
Distinct patients, no. (% of total admissions)	5,674 (14.7%)	8,091 (20.9%)	13,649 (35.4%)	6,372 (16.5%)	4,811 (12.5%)	38,597 (100%)
Hospital admissions, no. (% of total admissions)	7,258 (14.6%)	9,156 (18.4%)	19,770 (39.7%)	8,110 (16.3%)	5,491 (11.0%)	49,785 (100%)
Distinct ICU stays, no. (% of total admissions)	7,726 (14.5%)	9,854 (18.4%)	21,087 (39.5%)	8,891 (16.6%)	5,865 (11.0%)	53,423 (100%)
Age, years, median (Q1-Q3)	70.1 (58.4–80.5)	67.6 (57.6–76.7)	64.9 (51.7–78.2)	63.6 (51.4–76.5)	59.9 (42.9–75.7)	65.8 (52.8–77.8)
Gender, male, % of unit stays	4,203 (57.9%)	6,000 (65.5%)	10,193 (51.6%)	4,251 (52.4%)	3,336 (60.7%)	27,983 (55.9%)
ICU length of stay, median days (Q1-Q3)	2.2 (1.2–4.1)	2.2 (1.2–4.0)	2.1 (1.2–4.1)	2.3 (1.3–4.9)	2.1 (1.2–4.6)	2.1 (1.2–4.6)
Hospital length of stay, median days (Q1-Q3)	5.8 (3.1–10.0)	7.4 (5.2–11.4)	6.4 (3.7–11.7)	7.9 (4.4–14.2)	7.4 (4.1–13.6)	6.9 (4.1–11.9)
ICU mortality, percent of unit stays	685 (8.9%)	353 (3.6%)	2,222 (10.5%)	813 (9.1%)	492 (8.4%)	4,565 (8.5%)
Hospital mortality, percent of unit stays	817 (11.3%)	424 (4.6%)	2,859 (14.5%)	1,020 (12.6%)	628 (11.4%)	5,748 (11.5%)

Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.

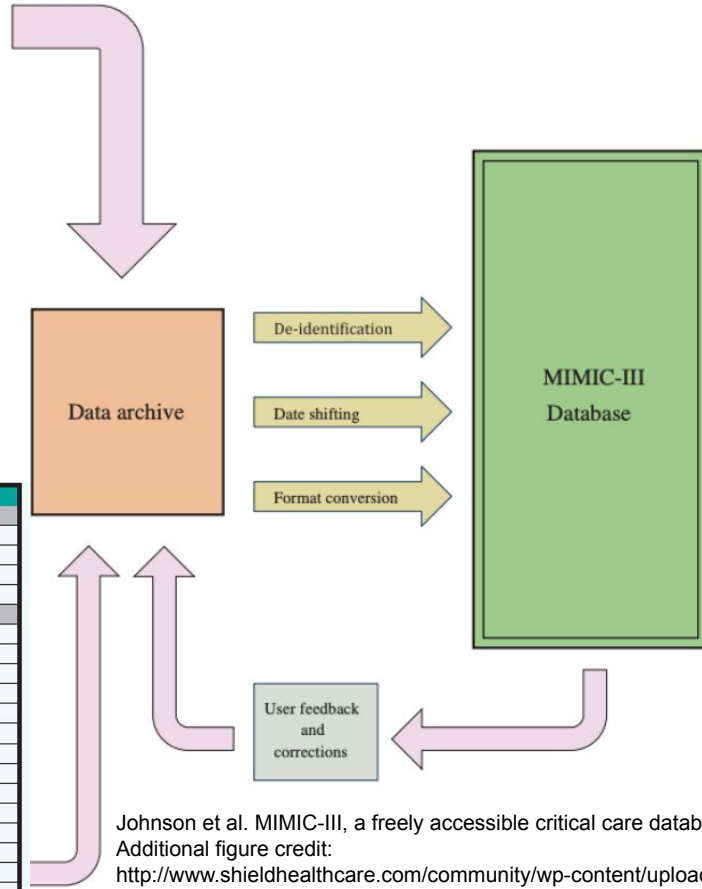


Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.

ICD9 (International classification of diseases):
Diagnosis codes

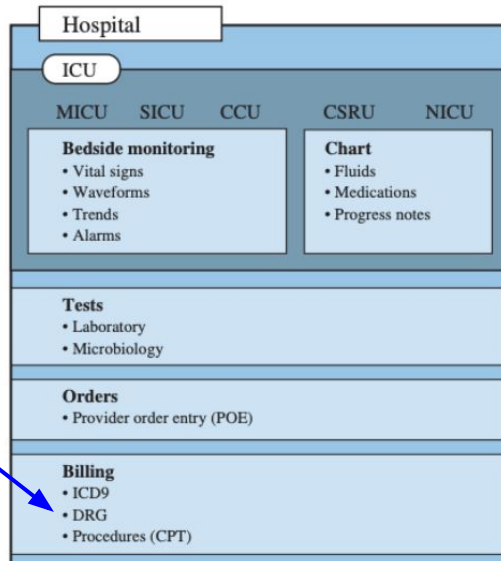


ICD-9	DESCRIPTION	ICD-10	DESCRIPTION
Congenital Malformations and Chromosomal Abnormalities (Including Down's Syndrome)			
753.8	Cystourethral anom NEC	Q64.79	Other congenital malformations of bladder and urethra
758.0	Down's syndrome	Q90.9	Down syndrome, unspecified
741.00	Spin bif w hydroceph NOS	Q05.4	Unspecified spina bifida with hydrocephalus
741.90	Spina bifida	Q05.8	Sacral spina bifida without hydrocephalus
Genitourinary System Diseases (including Incontinence and UTI)			
584.9	Acute kidney failure NOS	N17.9	Acute kidney failure, unspecified
596.9	Bladder disorder NOS	N32.9	Bladder disorder, unspecified
600.01	BPH w urinary obs/LUTS	N40.1	Enlarged prostate with lower urinary tract symptoms
600.00	BPH w/o urinary obs/LUTS	N40.0	Enlarged prostate without lower urinary tract symptoms
585.9	Chronic kidney dis NOS	N18.9	Chronic kidney disease, unspecified
753.19	Cystic kidney diseases NEC	Q61.8	Other cystic kidney diseases
585.6	End stage renal disease	N18.6	End stage renal disease
625.6	Fem stress incontinence	N39.3	Stress incontinence (female) (male)
787.60	Full incontinence-feces	R15.9	Full incontinence of feces
596.51	Hypertonicity of bladder	N32.81	Overactive bladder
788.21	Incomplet bladder emptying	R39.14	Feeling of incomplete bladder emptying
788.34	Inconthce wo sensr aware	N39.42	Incontinence without sensory awareness
788.33	Mixed incontinence	N39.46	Mixed incontinence
596.54	Neurogenic bladder NOS	N31.9	Neuromuscular dysfunction of bladder, unspecified
788.20	Oth urinary incontinence	N39.40R	Other specified urinary incontinence

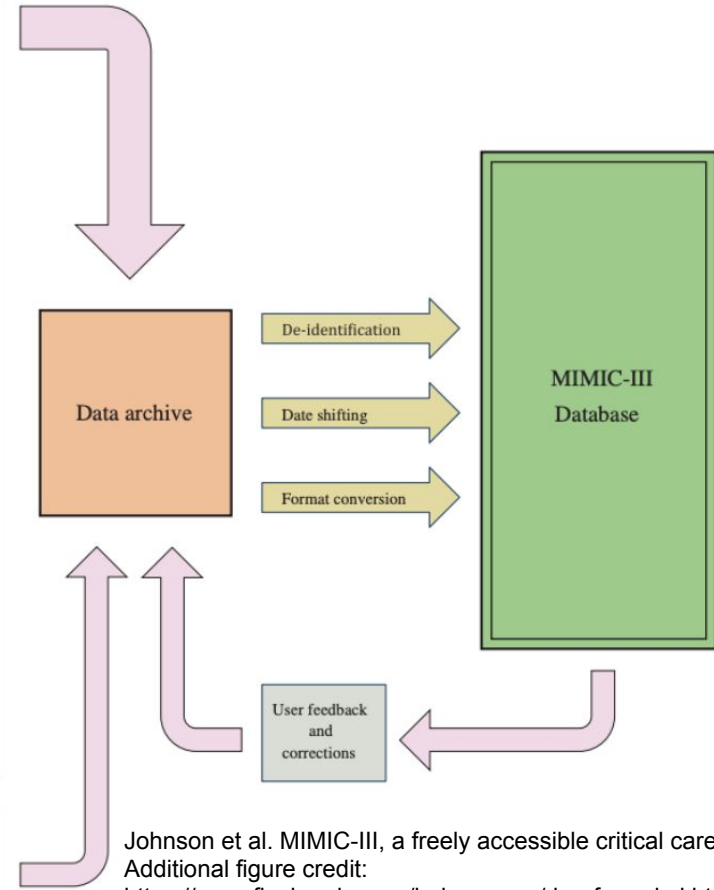


Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.
Additional figure credit:
<http://www.shieldhealthcare.com/community/wp-content/uploads/2015/08/ICD-9-to-ICD-10-Conversion-Guide-Page-1.jpg>

DRG (diagnosis related group): Higher-level codes describing patient groups w/ similar hospital resource use

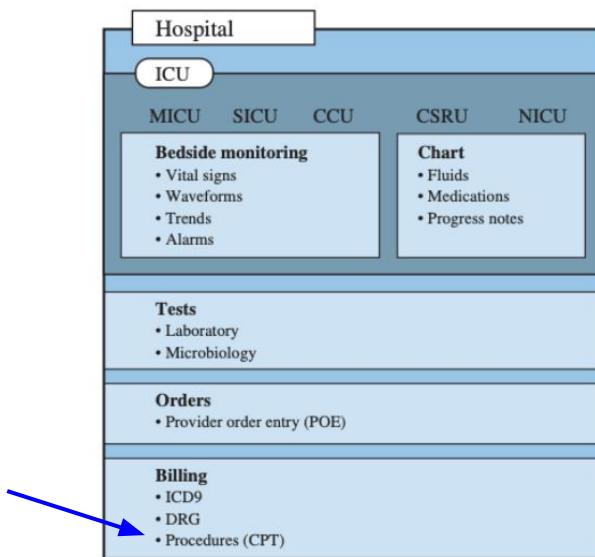


DRG Code and Description	
▶ 079	Respiratory Infections & Inflammations Age >17 w CC
▶ 121	Circulatory Disorders w AMI & Major Comp Discharged Alive
▶ 387	Prematurity w Major Problems
▶ 389	Full Term Neonate w Major Problems
▶ 489	HIV w Major Related Condition
▶ 489	HIV w Major Related Condition
▶ 080	Respiratory Infections & Inflammations Age >17 w/o CC
▶ 081	Respiratory Infections & Inflammations Age 0-17



Johnson et al. MIMIC-III, a freely accessible critical care database. 2016. Additional figure credit: https://www.flashcode.com/help_pages/drg_from_icd.html

CPT (Current procedural terminology): procedures and services codes



BONE DENSITOMETRY/DEXA

DEXA – hips, spine	77080
P-DEXA forearm	77081

CAT SCANS

ABDOMEN

Abdomen w/o contrast	74150
Abdomen w/ contrast	74160
Abdomen w/o & w/ contrast	74170

CHEST/THORAX

Chest/Thorax w/o contrast	71250
Chest/Thorax w/ contrast	74150
Chest/Thorax w/o & w/ contrast	71270

EXTREMITIES

Upper w/o contrast	73200
Upper w/o & w/ contrast	73202

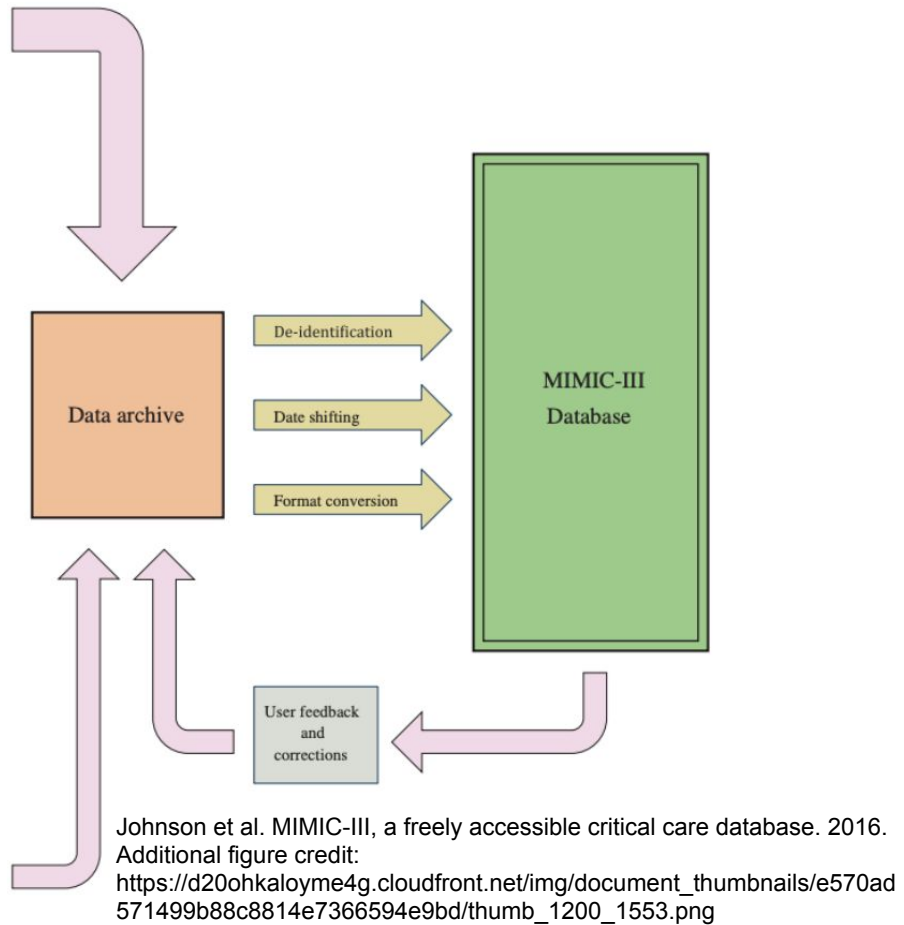
MRI

HEAD/NECK

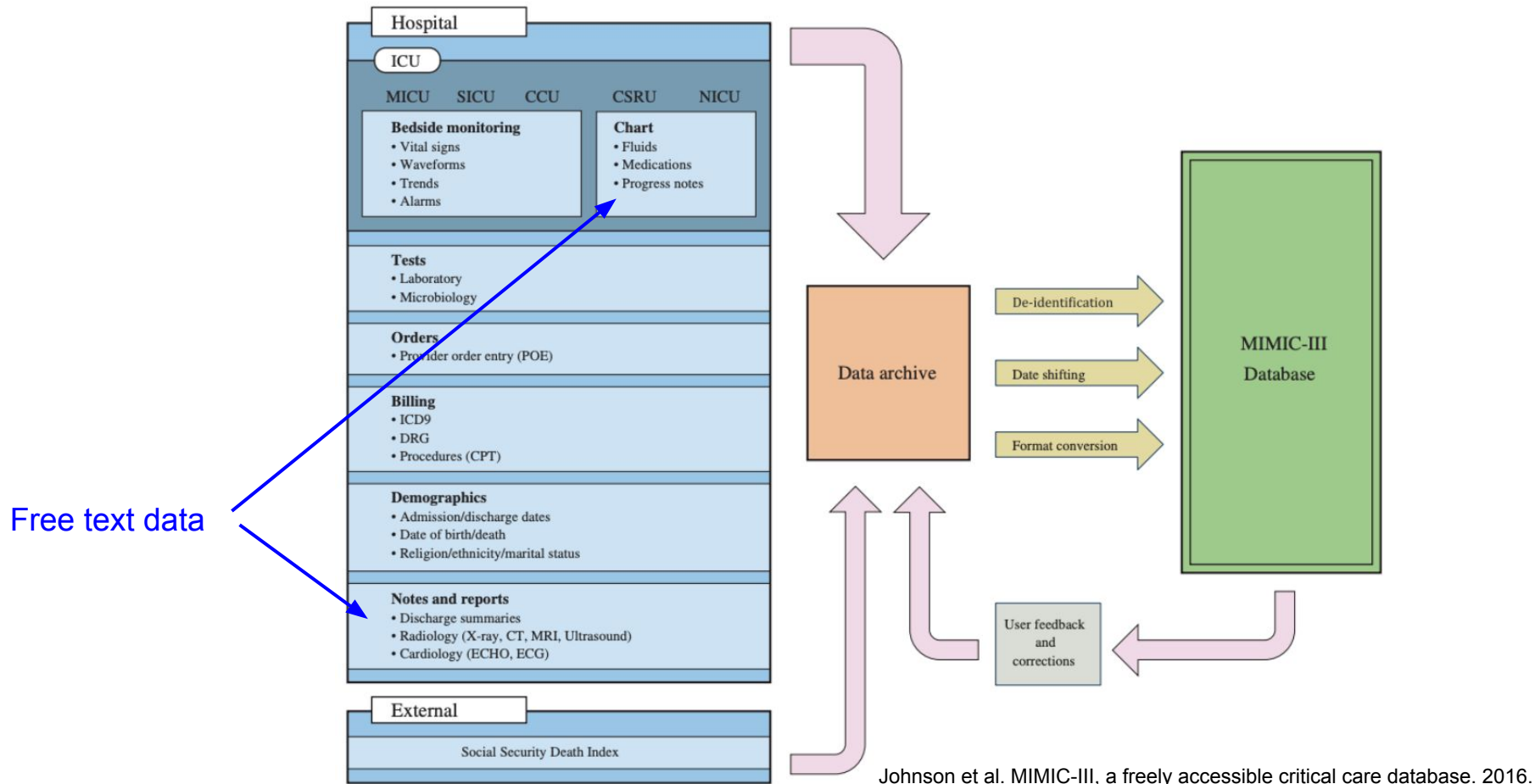
Head/Brain w/o contrast	70551
Head/Brain w/o & w/ contrast	70553
Internal AuditorY Canals (IAC's) w/o & w/	70553
MRA Circle of Willis w/o contrast	70544
MRA Neck (Carotids) w/o contrast	70547
Orbits/Face/Neck w/o	70540
Orbits/Face/Neck w/o & w/ contrast	70543
Pituitary Gland w/o & w/ contrast	70553
TMJ's (Temporomandibular Joints)	70336

SPINE/PELVIS

Cervical w/o contrast	72141
Cervical w/o & w/ contrast	72156
Lumbar w/o contrast	72148
Lumbar w/o & w/ contrast	72158
Pelvis w/o contrast	72195



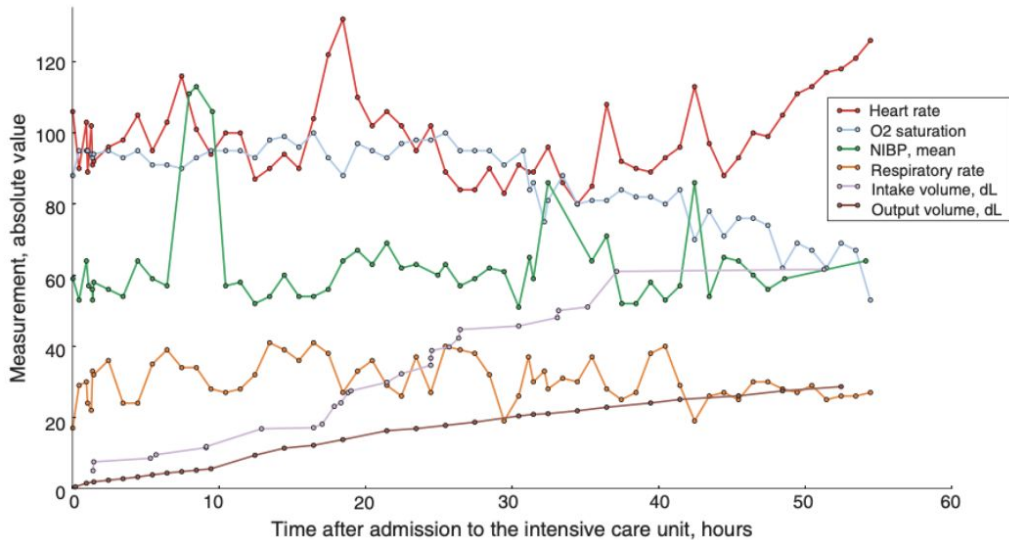
Johnson et al. MIMIC-III, a freely accessible critical care database. 2016. Additional figure credit: https://d20ohkaloyme4g.cloudfront.net/img/document_thumbnails/e570ad571499b88c8814e7366594e9bd/thumb_1200_1553.png



Critical care unit	CCU stays, No. (% by unit)	CSRU stays, No. (% by unit)	MICU stays, No. (% by unit)	SICU stays, No. (% by unit)	TSICU stays, No. (% by unit)	Total stays, No. (% by unit)
Infectious and parasitic diseases, i.e., septicemia, other infectious and parasitic diseases, etc., (001-139)	305 (4.2%)	72 (0.8%)	3,229 (16.7%)	448 (5.6%)	152 (2.8%)	4,206 (8.6%)
Neoplasms of digestive organs and intrathoracic organs, etc., (140-239)	126 (1.8%)	287 (3.2%)	1,415 (7.3%)	1,225 (15.3%)	466 (8.6%)	3,519 (7.2%)
Endocrine, nutritional, metabolic, and immunity (240-279)	104 (1.4%)	36 (0.4%)	985 (5.1%)	178 (2.2%)	54 (1.0%)	1,357 (2.8%)
Diseases of the circulatory system, i.e., ischemic heart diseases, diseases of pulmonary circulation, dysrhythmias, heart failure, cerebrovascular diseases, etc., (390-459)	5,131 (71.4%)	7,138 (78.6%)	2,638 (13.6%)	2,356 (29.5%)	684 (12.6%)	17,947 (36.6%)
Pulmonary diseases, i.e., pneumonia and influenza, chronic obstructive pulmonary disease, etc., (460-519)	416 (5.8%)	141 (1.6%)	3,393 (17.5%)	390 (4.9%)	225 (4.1%)	4,565 (9.3%)
Diseases of the digestive system (520-579)	264 (3.7%)	157 (1.7%)	3,046 (15.7%)	1,193 (14.9%)	440 (8.1%)	5,100 (10.4%)
Diseases of the genitourinary system, i.e., nephritis, nephrotic syndrome, nephrosis, and other diseases of the genitourinary system (580-629)	130 (1.8%)	14 (0.2%)	738 (3.8%)	101 (1.3%)	31 (0.6%)	1,014 (2.1%)
Trauma (800-959)	97 (1.3%)	494 (5.4%)	480 (2.5%)	836 (10.5%)	2,809 (51.7%)	4,716 (9.6%)
Poisoning by drugs and biological substances (960-979)	50 (0.7%)	2 (0.0%)	584 (3.0%)	58 (0.7%)	11 (0.2%)	705 (1.4%)
Other	565 (7.9%)	739 (8.1%)	2,883 (14.9%)	1,204 (15.1%)	563 (10.4%)	5,954 (12.1%)
Total	7,188 (14.6%)	9,080 (18.5%)	19,391 (39.5%)	7,989 (16.3%)	5,435 (11.1%)	49,083 (100%)

Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.

Code status	Full code						Comfort measures
GCS: Verbal	Oriented		Oriented		Oriented		Incomprehensible sounds
GCS: Motor	Obeys commands		Obeys commands		Obeys commands		Flex-withdraws
GCS: Eye	Spontaneously		Spontaneously		To speech		None
Platelet, K/uL	48	53	46		45		
Creatinine, mg/dL	0.7		0.7		0.8		
White blood cell, K/uL	9.1	12.4	16.8		23.2		
Neutrophil, %	37						
Morphine Sulfate							
Vancocycin (1 dose)							
Piperacillin (1 dose)							
NaCl 0.9%	10.0 mL/hour						
Amiodarone			1mg/min		0.5mg/min		
Dextrose 5%			50mL/hour		25mL/hour		



Johnson et al. MIMIC-III, a freely accessible critical care database. 2016.

Examples of prediction tasks: phenotypes

- What conditions a patient has
- Useful for patient treatment and risk monitoring



Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Examples of prediction tasks: in-hospital mortality

- Whether patient will die in the hospital
- Early detection of at-risk patients can improve outcomes

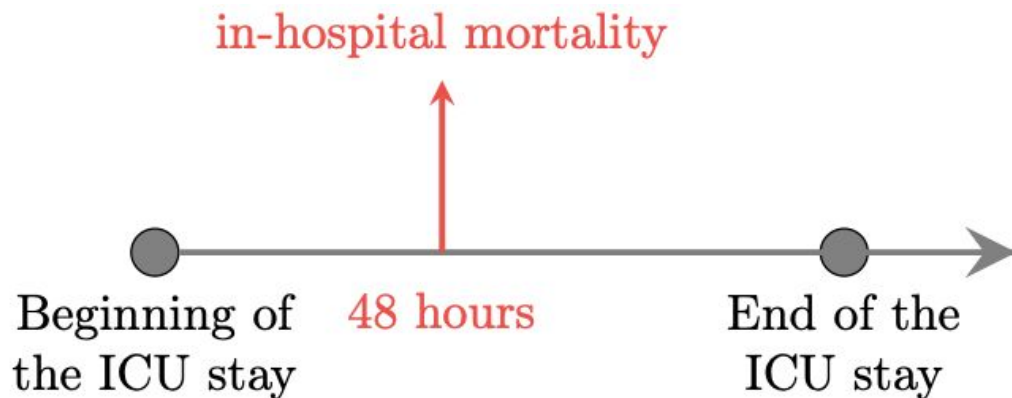


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Examples of prediction tasks: decompensation

- Whether patient will die in the next 24 hours
- Also for early detection, related to in-hospital mortality

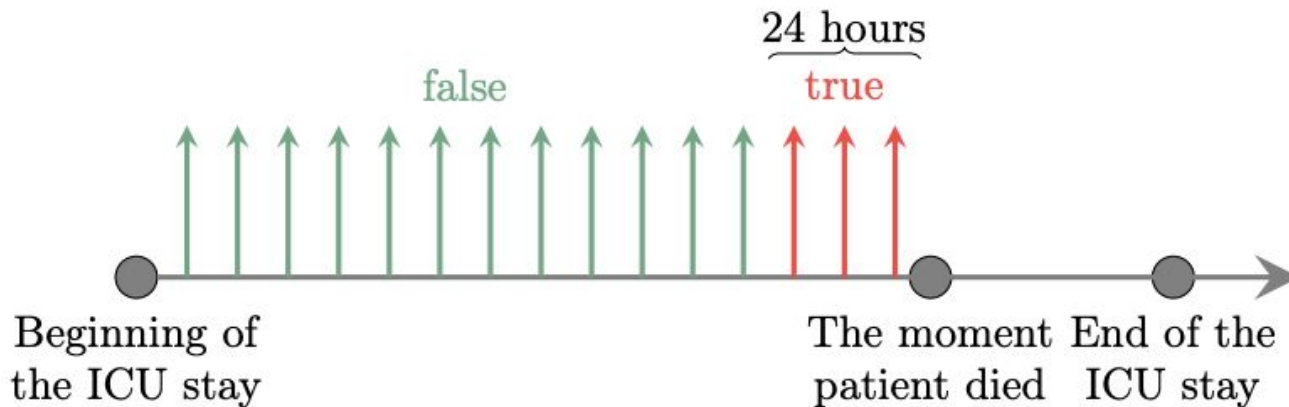


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Examples of prediction tasks: length-of-stay

- How much longer the patient is expected to stay in the ICU
- Useful for measuring patient acuity and resource management

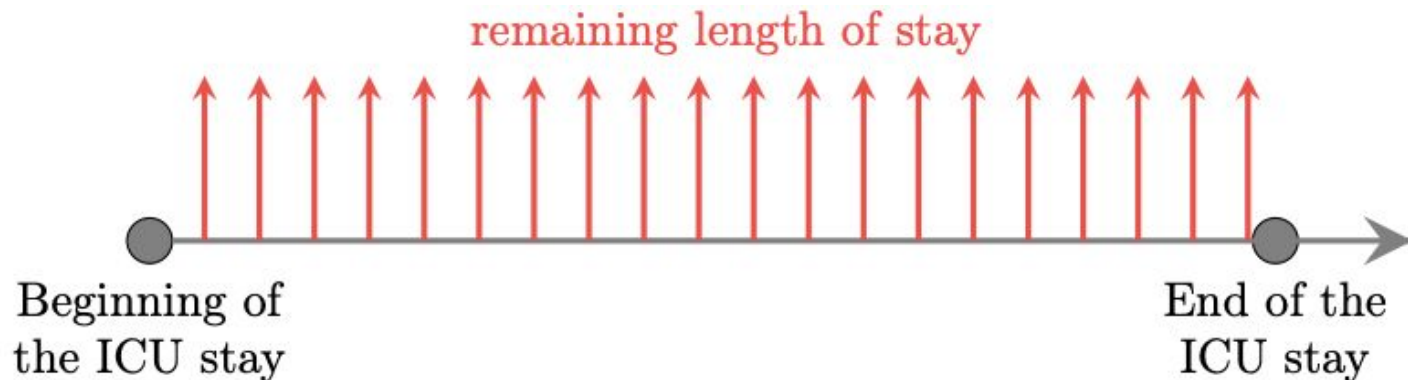


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Remember: “vanilla” neural networks for predictions from clinical variables

Let us consider the task of **regression**: predicting a single real-valued output from input data

Model input: data vector $x = [x_1, x_2, \dots, x_N]$ **Model output:** prediction (single number) \hat{y}

Example: predicting hospital length-of-stay from clinical variables in the electronic health record

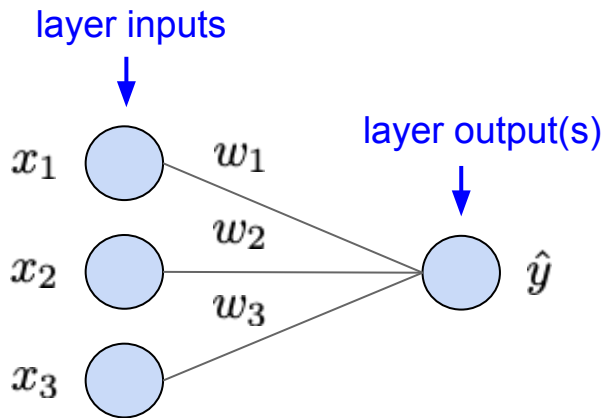
$x =$ [age, weight, ..., temperature, oxygen saturation] $\hat{y} =$ length-of-stay (days)

Remember: “vanilla” neural networks for predictions from clinical variables

Our first architecture: a single-layer, fully connected neural network

For simplicity, use a 3-dimensional input (N = 3)

all inputs of a layer are connected to all outputs of a layer



$$\text{Output: } \hat{y} = w_1x_1 + w_2x_2 + w_3x_3 + b \\ = w^T x + b$$

bias term (allows constant shift)

Neural network parameters:

$$W = \{[w_1, w_2, w_3], b\}$$

layer “weights”

layer bias

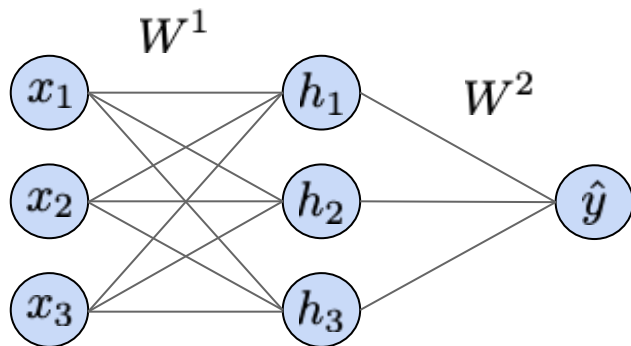
Often refer to all parameters together as just “weights”. Bias is implicitly assumed.

Caveats of our first (simple) neural network architecture:

- Single layer still “shallow”, not yet a “deep” neural network. Will see soon how to stack multiple layers.
- Also equivalent to a linear regression model! But useful base case for deep learning.

Remember: “vanilla” neural networks for predictions from clinical variables

Two-layer fully-connected neural network



$$W^1 = \begin{bmatrix} w_{11}^1 & w_{12}^1 & w_{13}^1 \\ w_{21}^1 & w_{22}^1 & w_{23}^1 \\ w_{31}^1 & w_{32}^1 & w_{33}^1 \end{bmatrix} \quad b^1 = \begin{bmatrix} b_1^1 \\ b_2^1 \\ b_3^1 \end{bmatrix}$$

$$W^2 = [w_{11}^2 \quad w_{12}^2 \quad w_{13}^2] \quad b^2 = [b_1^2]$$

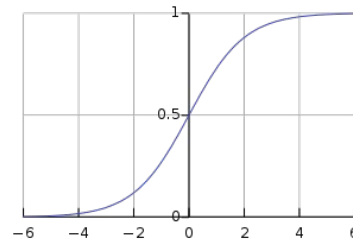
Output: $h = \sigma(W^1x + b^1)$

$$\hat{y} = W^2h + b^2$$

Full function expression:

$$\hat{y} = W^2(\sigma(W^1x + b^1)) + b^2$$

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



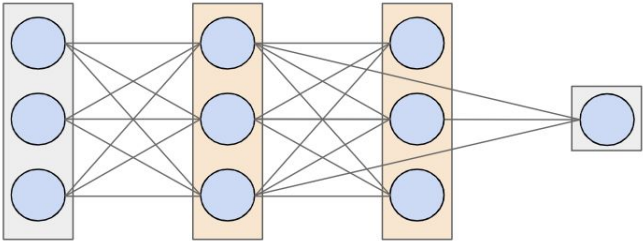
Sigmoid “activation function”

Activation functions

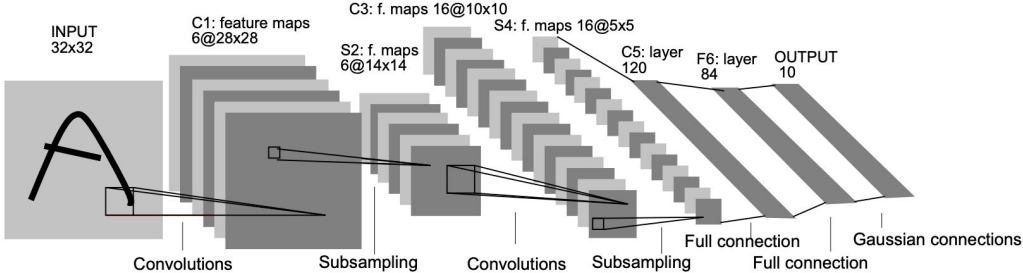
introduce non-linearity into the model -- allowing it to represent highly complex functions.

A fully-connected neural network (also known as multi-layer perceptron) is a stack of [affine transformation + activation function] layers. There is no activation function at the last layer.

Different classes of neural networks

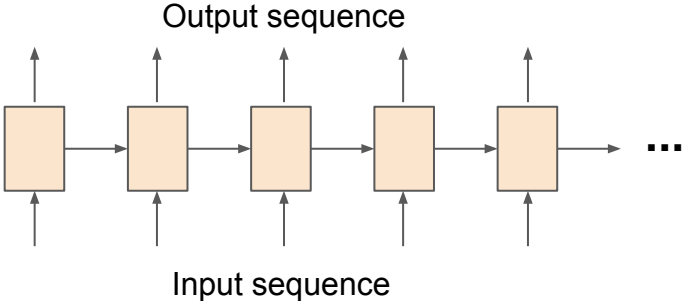


Fully connected neural networks
(linear layers, good for “feature vector” inputs)

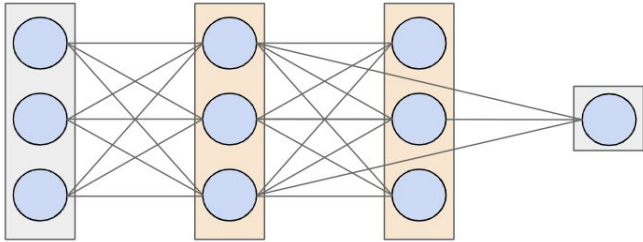


Convolutional neural networks
(convolutional layers, good for image inputs)

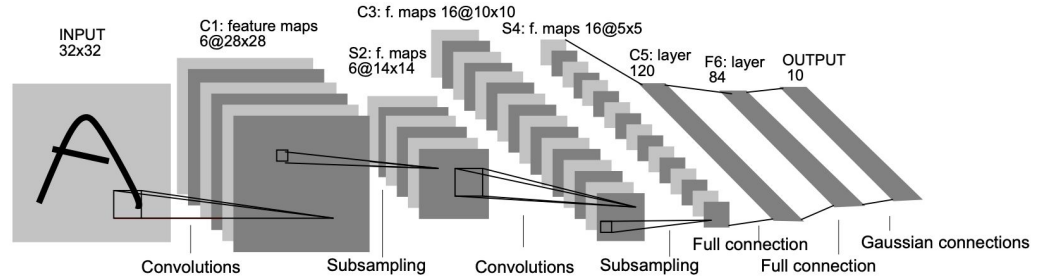
Recurrent neural networks
(linear layers modeling recurrence relation across sequence, good for sequence inputs)



Different classes of neural networks



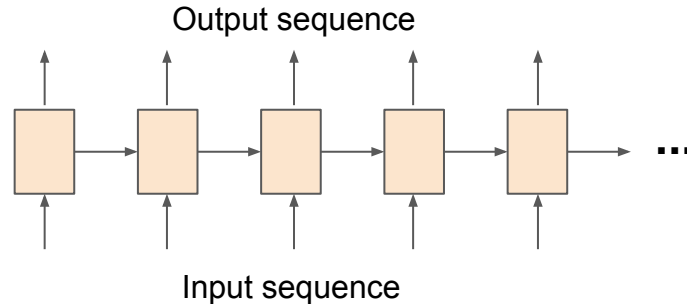
Fully connected neural networks
(linear layers, good for “feature vector” inputs)



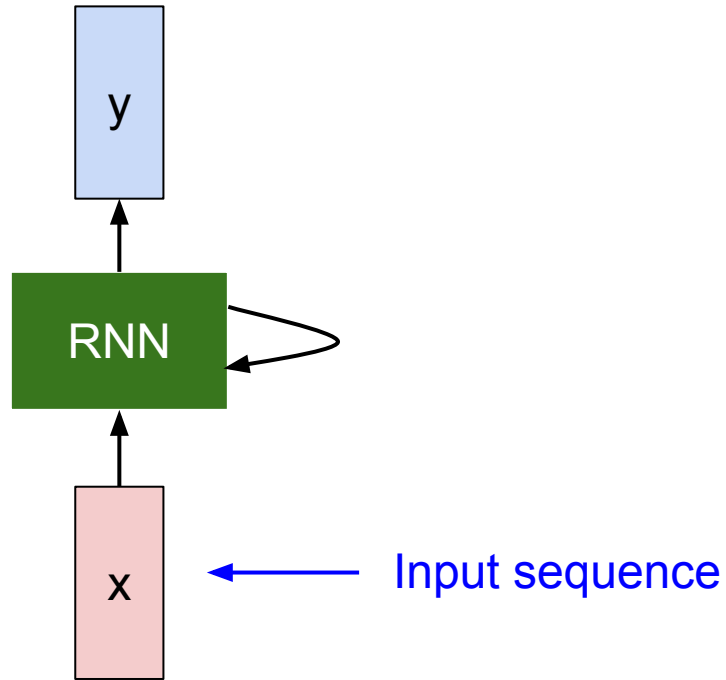
Convolutional neural networks
(convolutional layers, good for image inputs)

Natural fit for clinical data time-series

Recurrent neural networks
(linear layers modeling recurrence relation across sequence, good for sequence inputs)

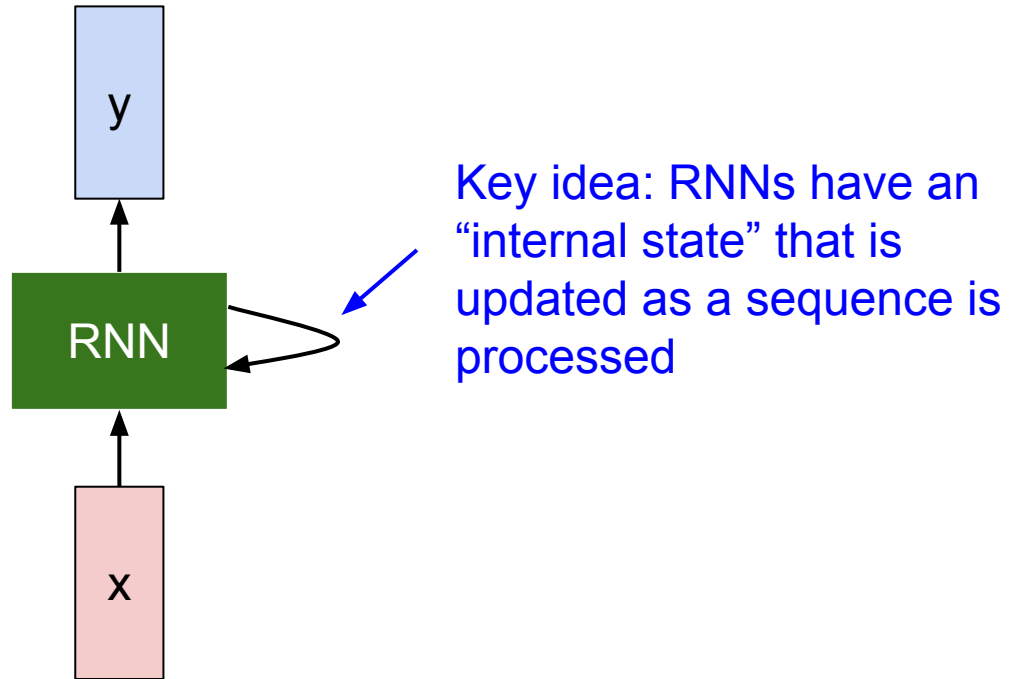


Recurrent Neural Network



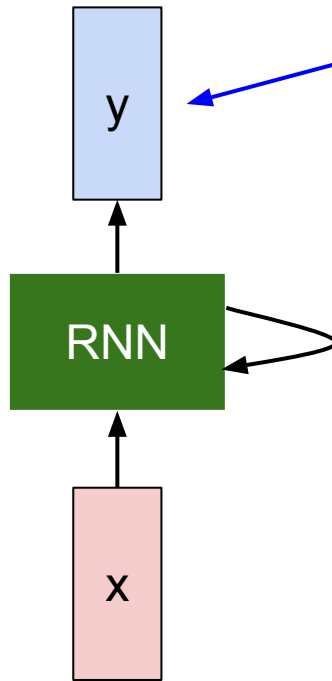
Slide credit: CS231n

Recurrent Neural Network



Slide credit: CS231n

Recurrent Neural Network



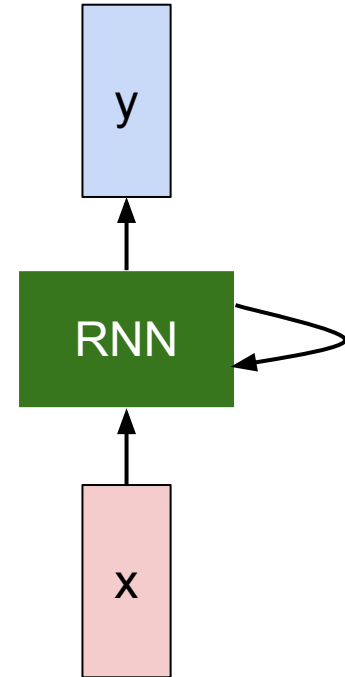
Output
(do not necessarily need
to have an output at every
timestep of the sequence)

Recurrent Neural Network

We can process a sequence of vectors \mathbf{x} by applying a **recurrence formula** at every time step:

$$\boxed{h_t} = \boxed{f_W}(\boxed{h_{t-1}}, \boxed{x_t})$$

new state / some function with parameters W / old state / input vector at some time step

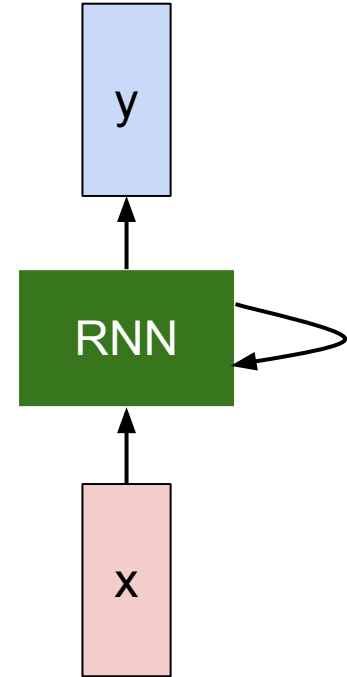


Recurrent Neural Network

We can process a sequence of vectors \mathbf{x} by applying a **recurrence formula** at every time step:

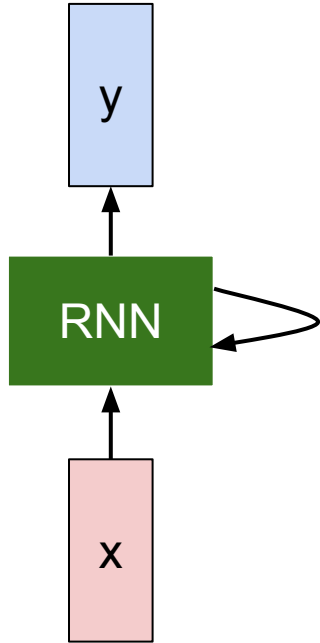
$$h_t = f_W(h_{t-1}, x_t)$$

Notice: the same function and the same set of parameters are used at every time step.



(Vanilla) Recurrent Neural Network

The state consists of a single “hidden” vector h :



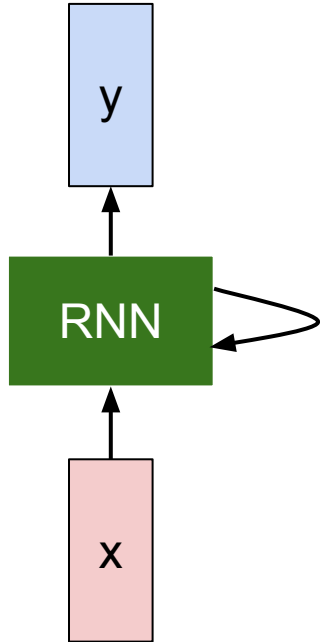
$$h_t = f_W(h_{t-1}, x_t)$$



$$h_t = \tanh(W_{hh}h_{t-1} + W_{xh}x_t)$$

(Vanilla) Recurrent Neural Network

The state consists of a single “hidden” vector h :



$$h_t = f_W(h_{t-1}, x_t)$$

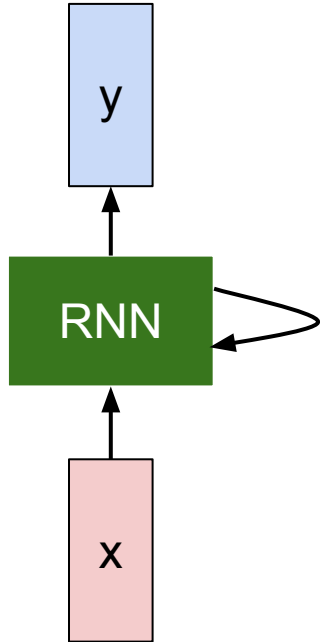


$$h_t = \tanh(W_{hh}h_{t-1} + W_{xh}x_t)$$

$$y_t = W_{hy}h_t$$

(Vanilla) Recurrent Neural Network

The state consists of a single “hidden” vector h :



$$h_t = f_W(h_{t-1}, x_t)$$

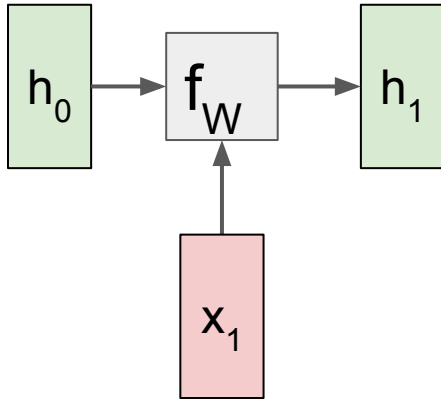


$$h_t = \tanh(W_{hh}h_{t-1} + W_{xh}x_t)$$

$$y_t = W_{hy}h_t$$

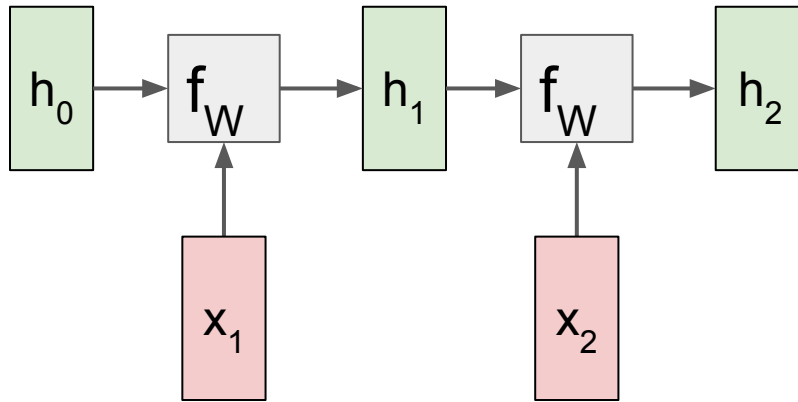
Fully connected layers

RNN: Computational Graph



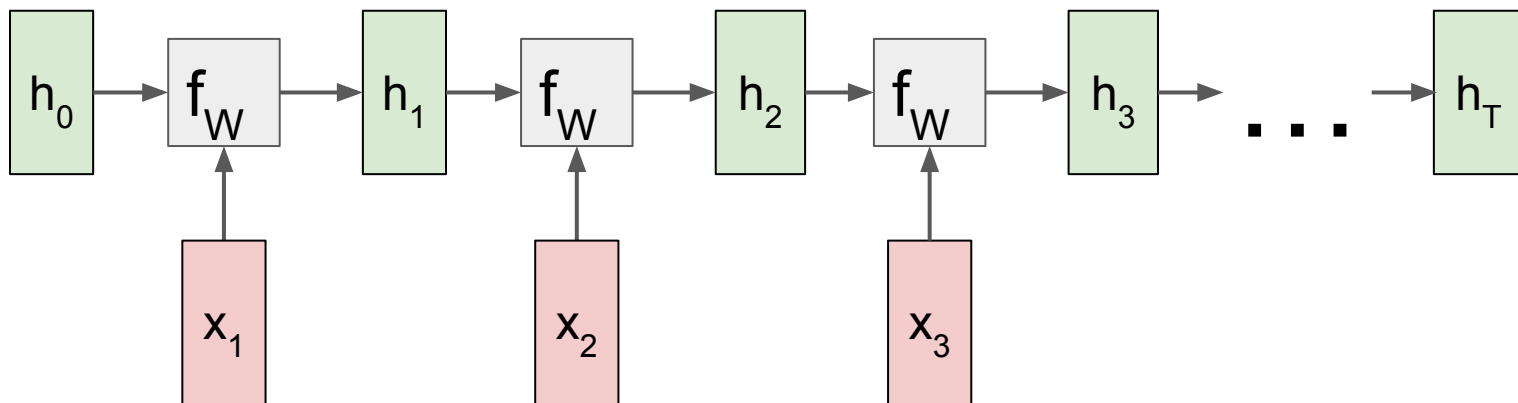
Slide credit: CS231n

RNN: Computational Graph



Slide credit: CS231n

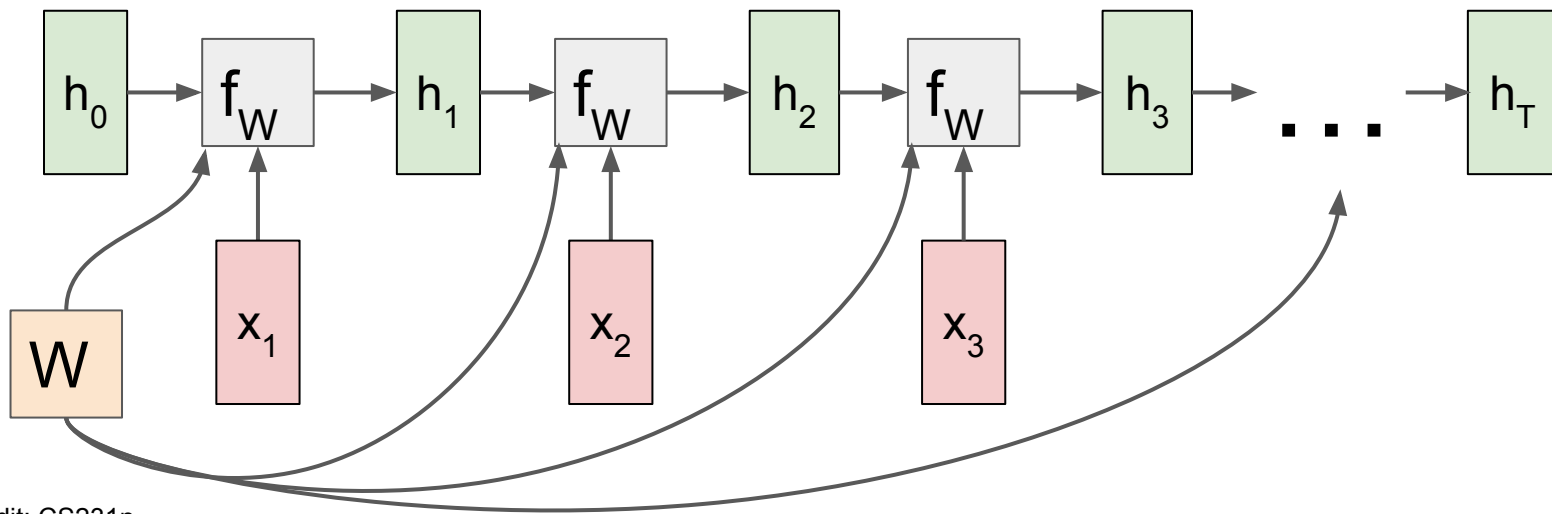
RNN: Computational Graph



Slide credit: CS231n

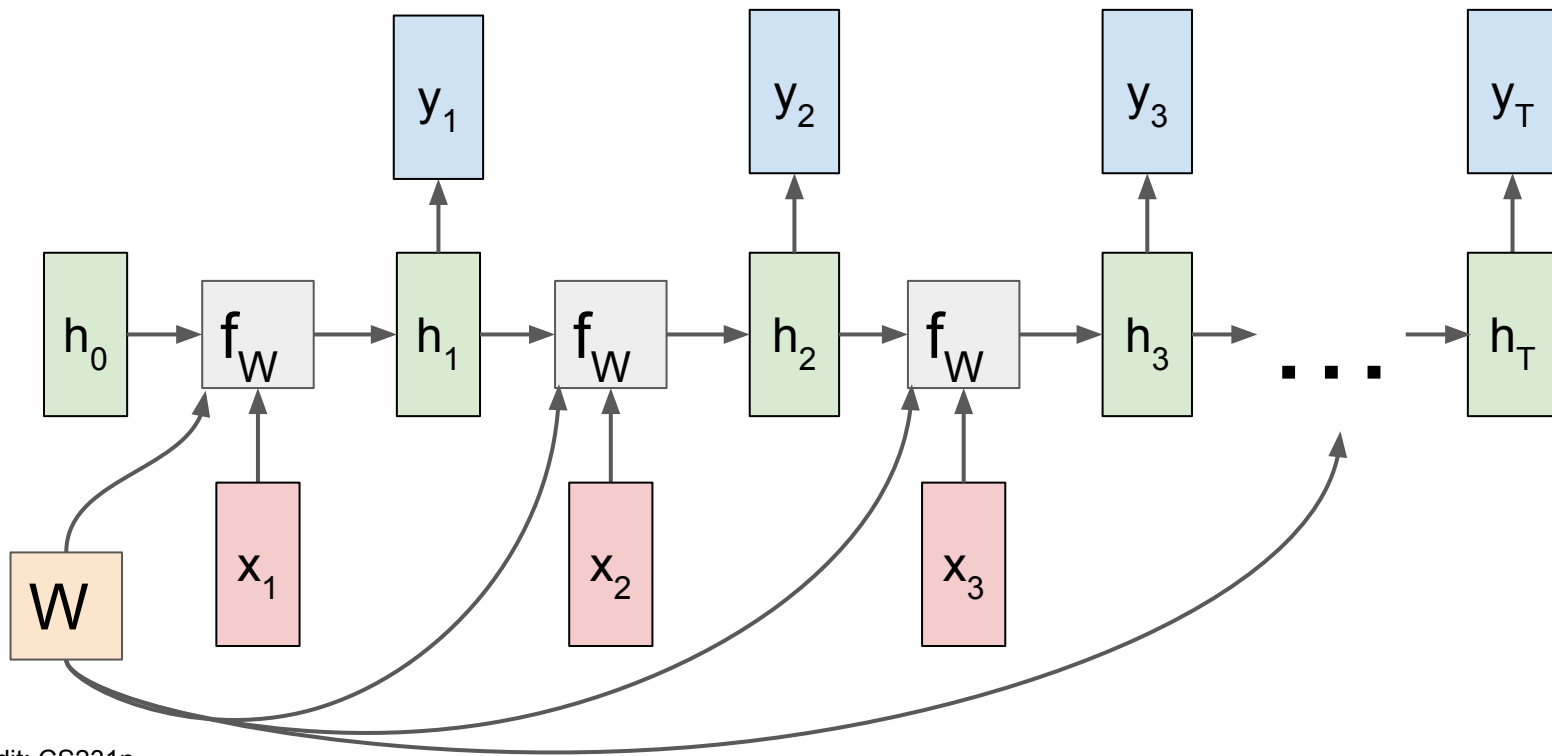
RNN: Computational Graph

Re-use the same weight matrix at every time-step



Slide credit: CS231n

RNN: Computational Graph: Many to Many

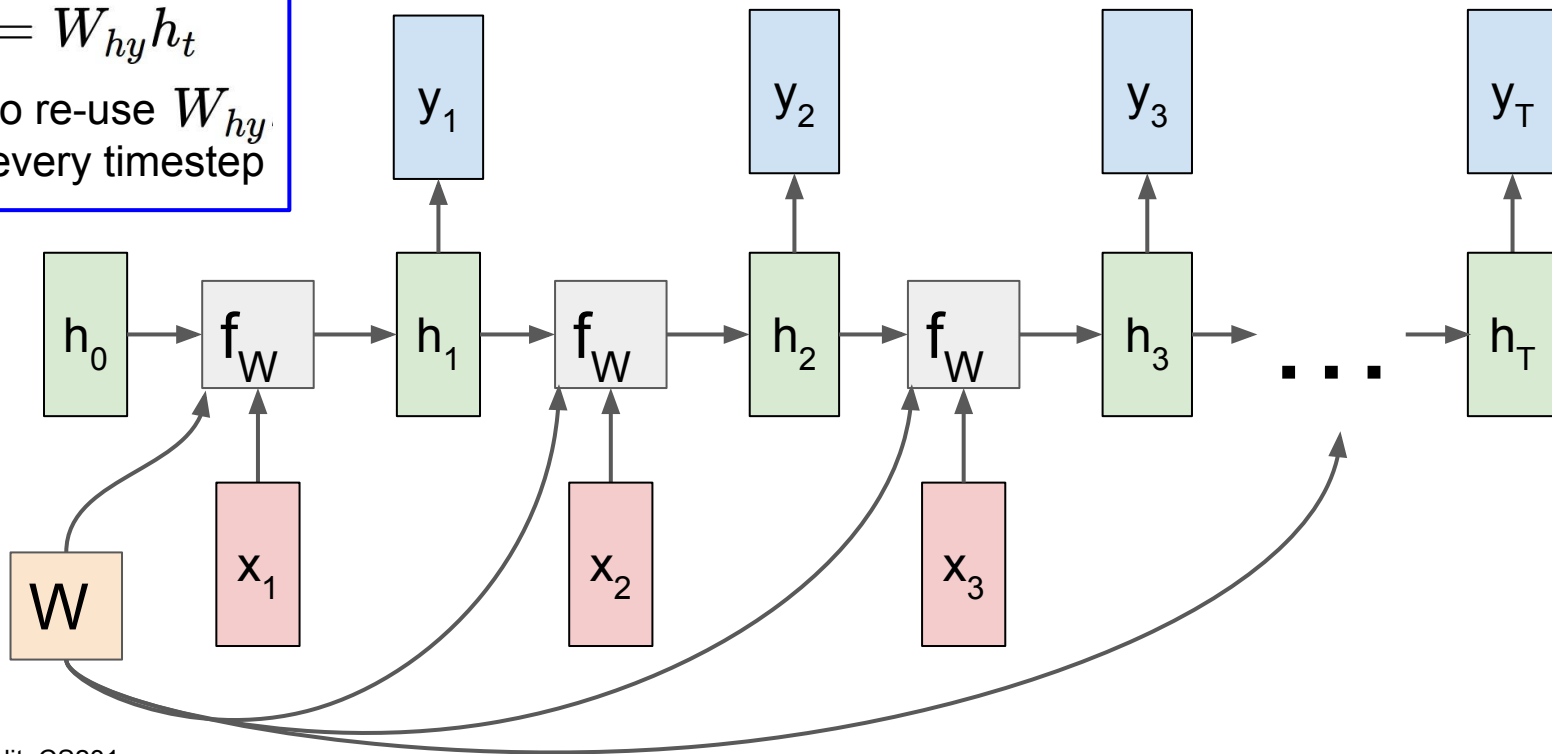


Slide credit: CS231n

RNN: Computational Graph: Many to Many

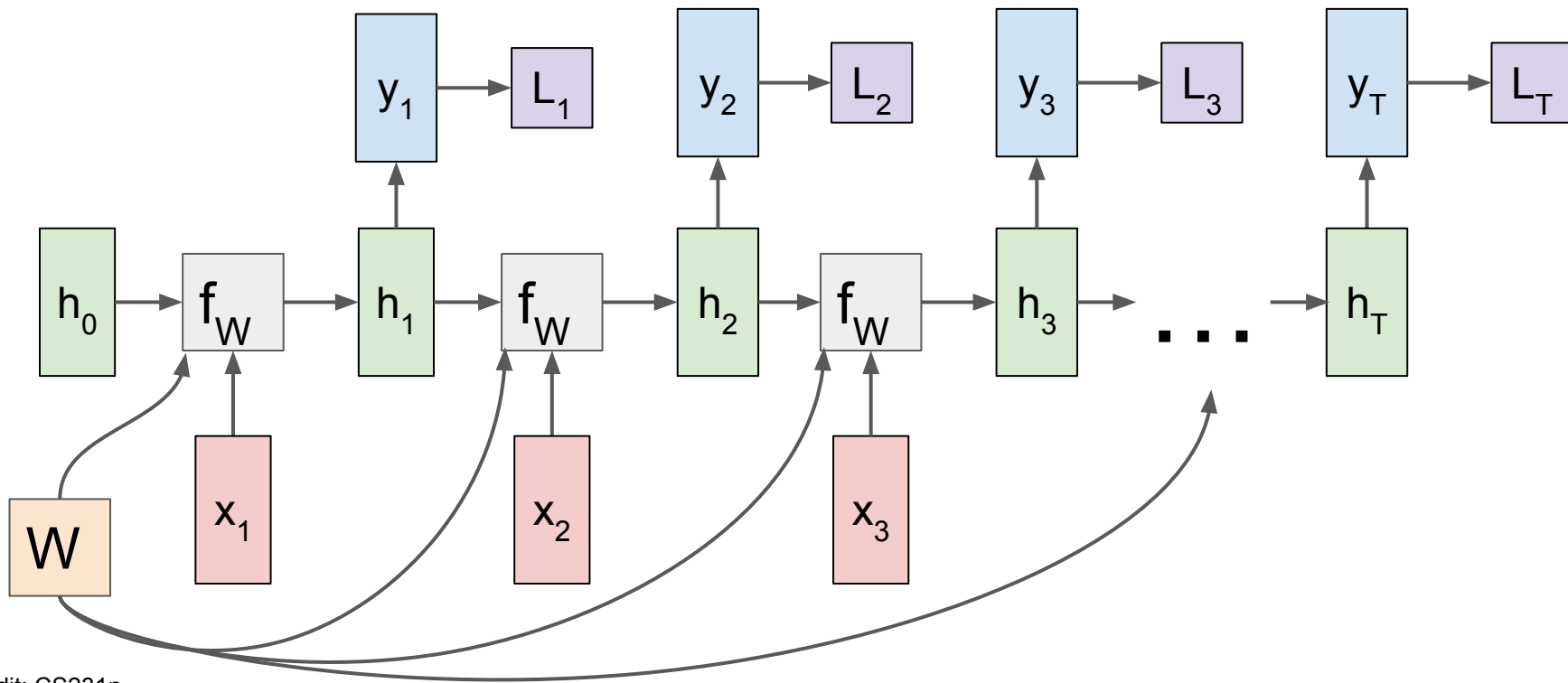
$$y_t = W_{hy}h_t$$

Also re-use W_{hy}
at every timestep



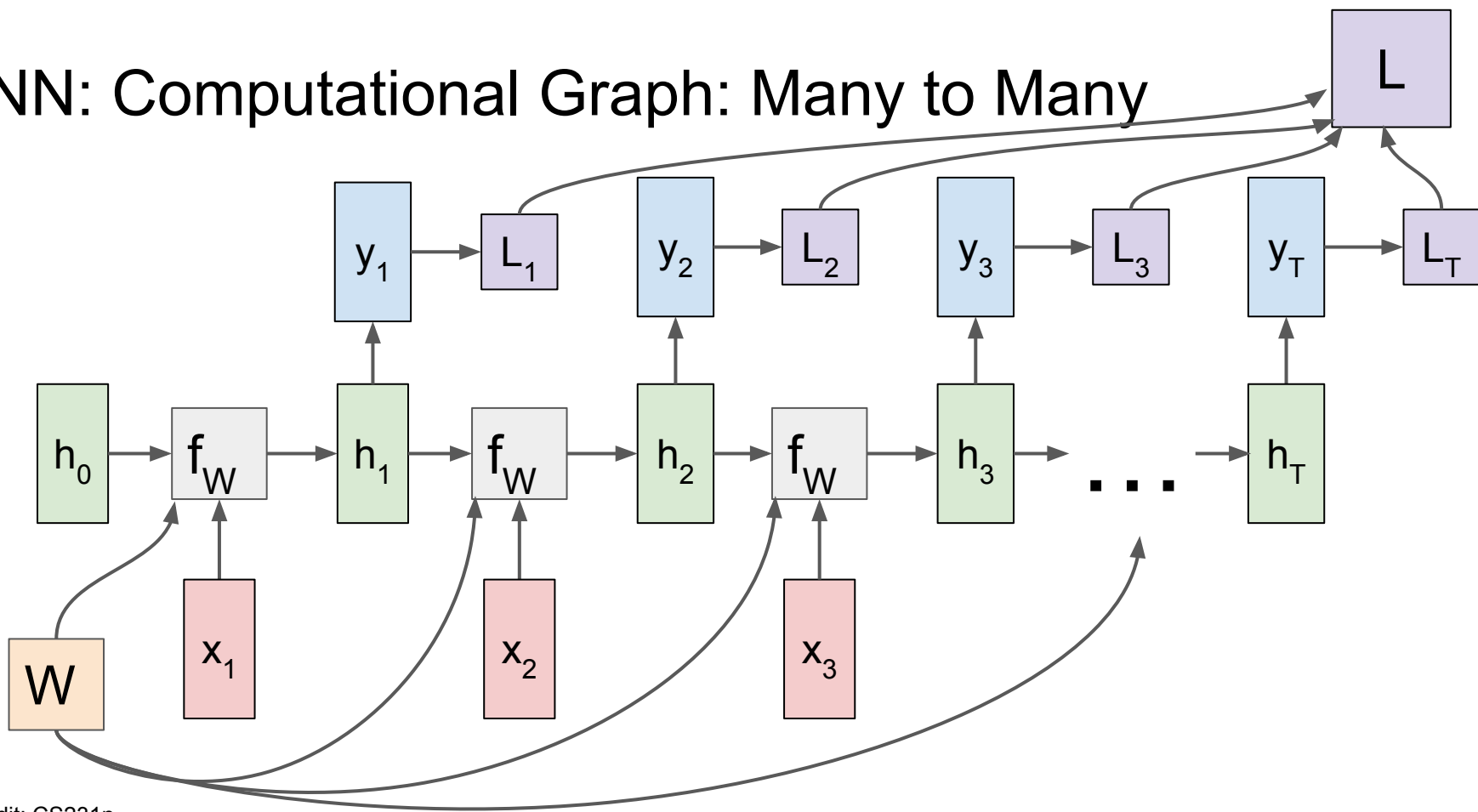
Slide credit: CS231n

RNN: Computational Graph: Many to Many



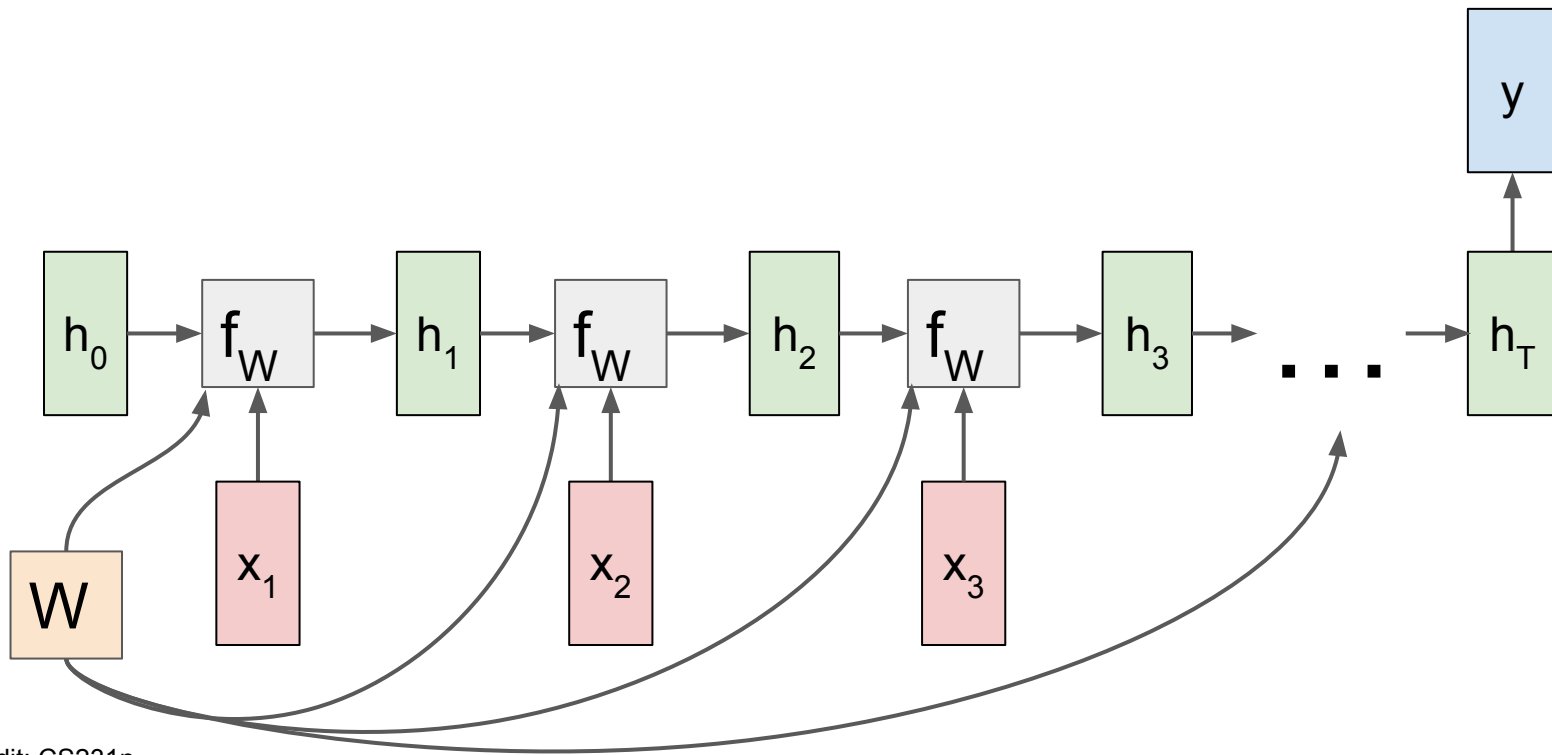
Slide credit: CS231n

RNN: Computational Graph: Many to Many



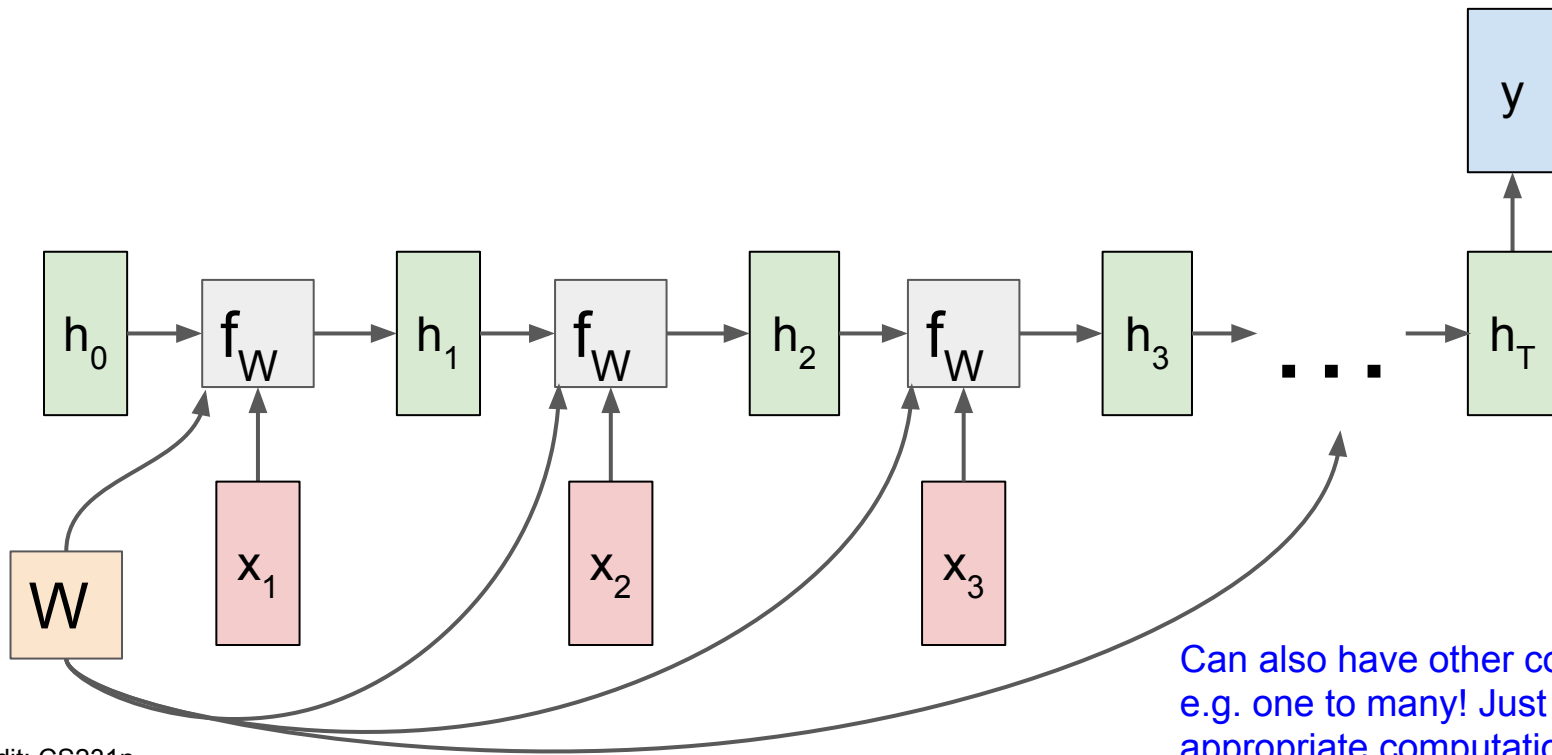
Slide credit: CS231n

RNN: Computational Graph: Many to One



Slide credit: CS231n

RNN: Computational Graph: Many to One

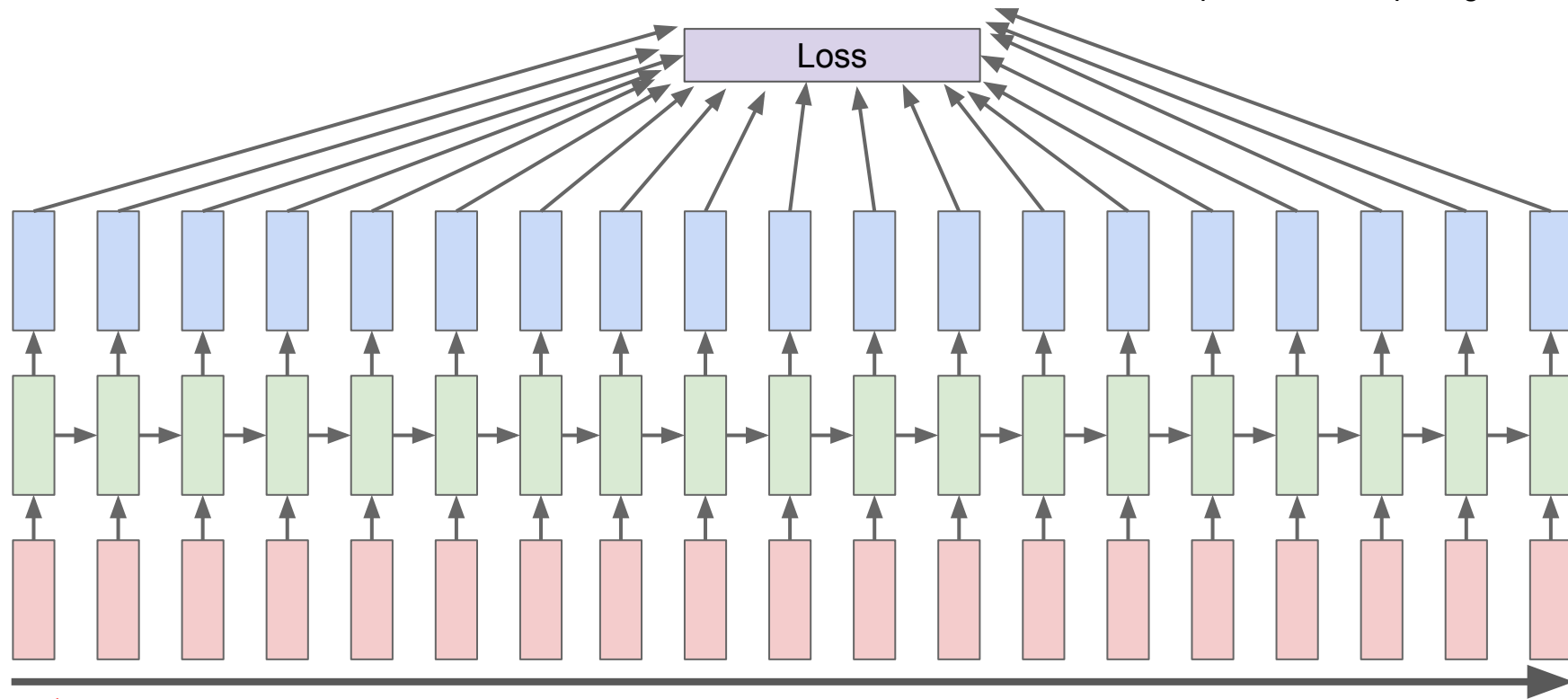


Can also have other configurations
e.g. one to many! Just define
appropriate computational graph.

Slide credit: CS231n

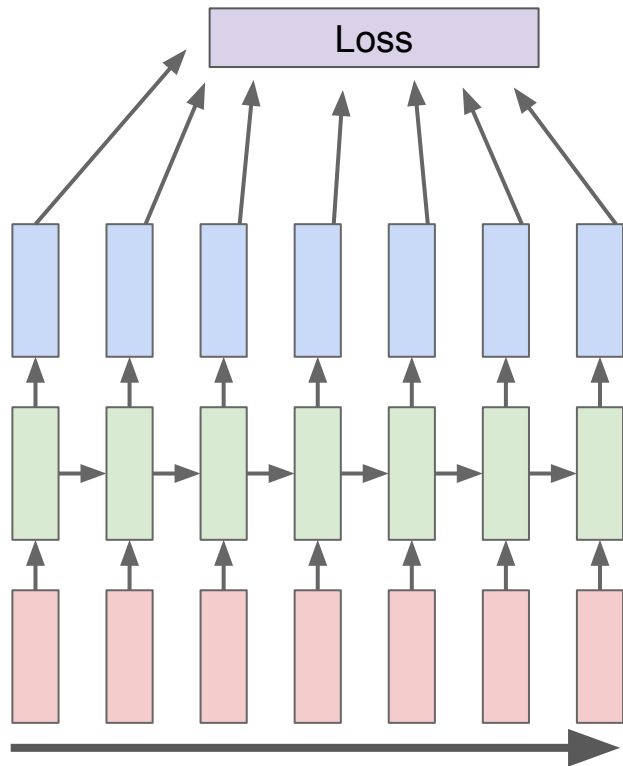
Backpropagation through time

Forward through entire sequence to compute loss, then backward through entire sequence to compute gradient



Slide credit: CS231n

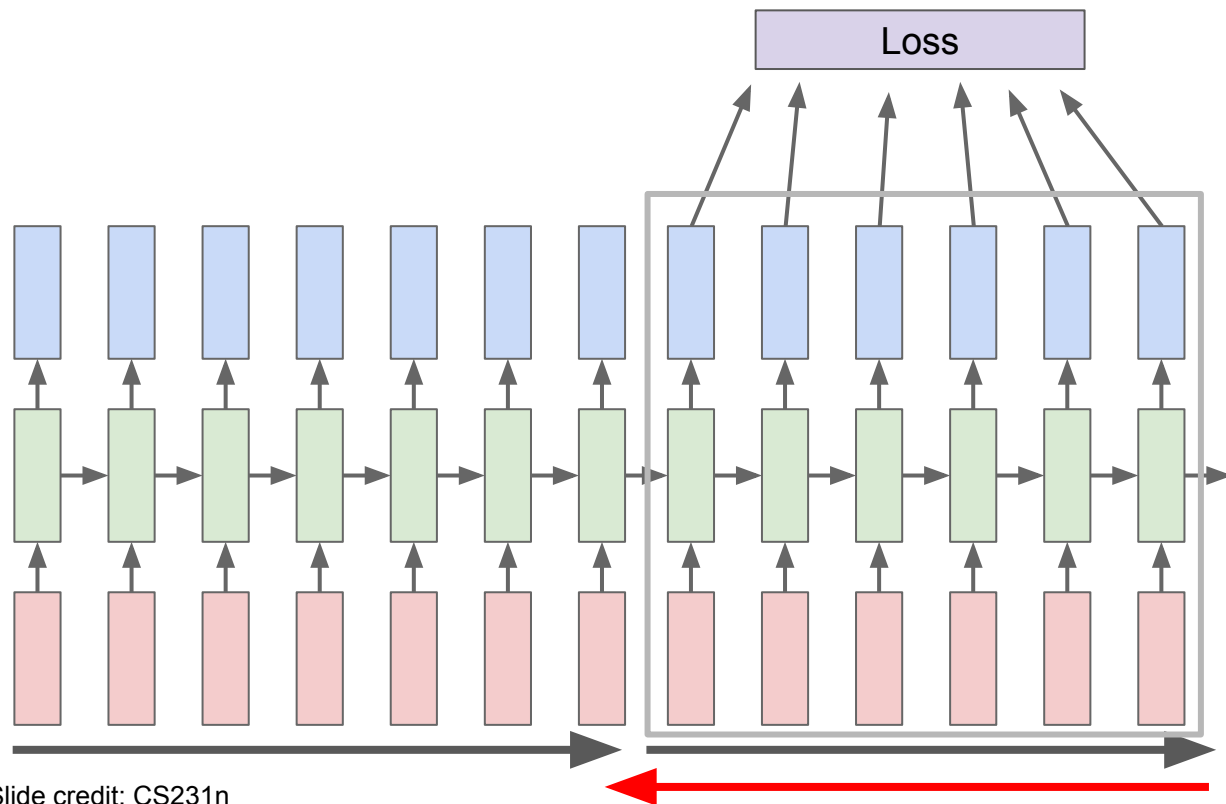
Truncated Backpropagation through time



Run forward and backward through chunks of the sequence instead of whole sequence

Slide credit: CS231n

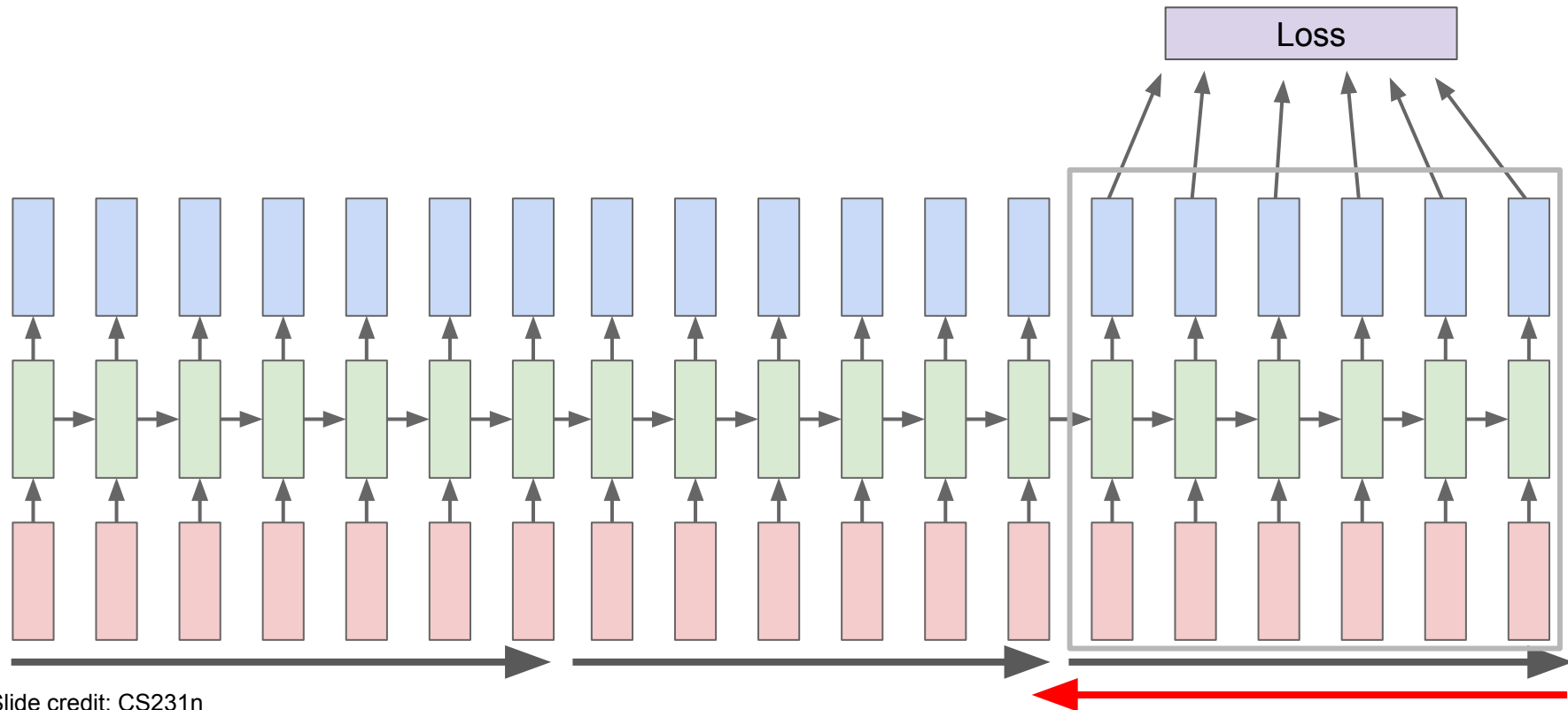
Truncated Backpropagation through time



Carry hidden states forward in time forever, but only backpropagate for some smaller number of steps

Slide credit: CS231n

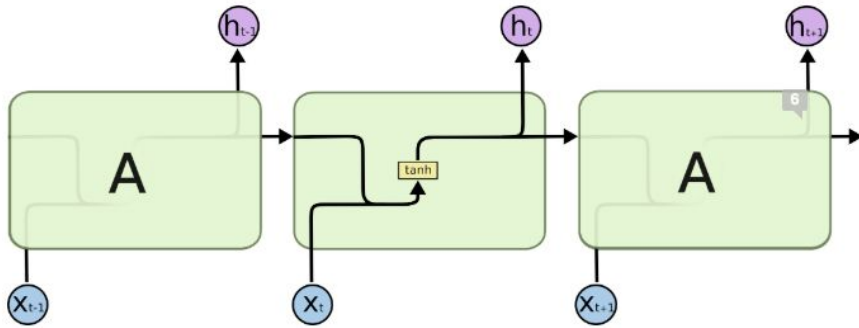
Truncated Backpropagation through time



Slide credit: CS231n

Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



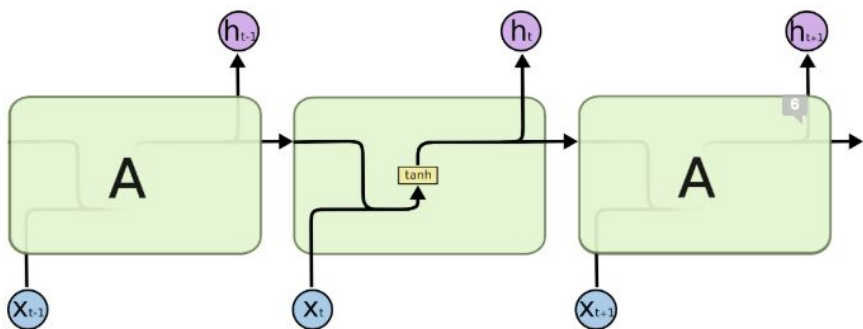
$$h_t = \tanh(W_{hh}h_{t-1} + W_{xh}x_t)$$

$$y_t = W_{hy}h_t$$

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

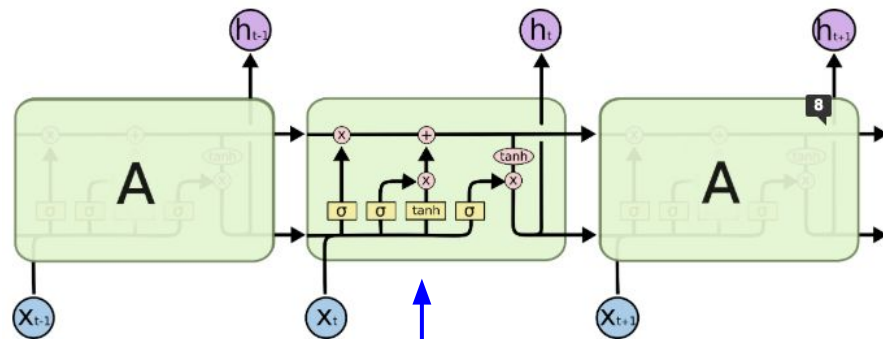
Unrolled Vanilla RNN



$$h_t = \tanh(W_{hh}h_{t-1} + W_{xh}x_t)$$

$$y_t = W_{hy}h_t$$

Unrolled LSTM



Different computation to obtain h_t

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

“Cell state” flows through entire sequence. At each timestep, will be able to modify the cell state.

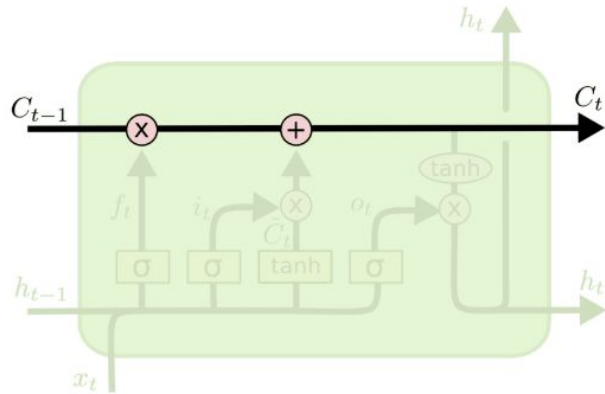
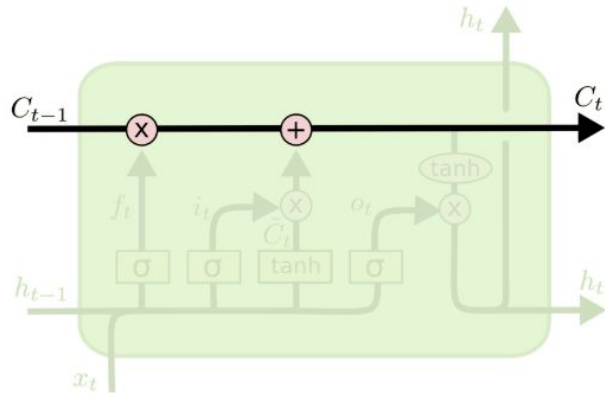


Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

“Cell state” flows through entire sequence. At each timestep, will be able to modify the cell state.



Gates (sigmoid + elementwise multiplication) control passing of information. Sigmoid output of 1 = let everything through; output of 0 = let nothing through.

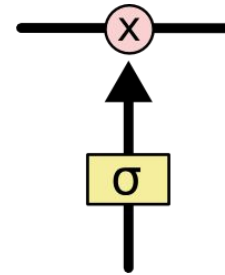
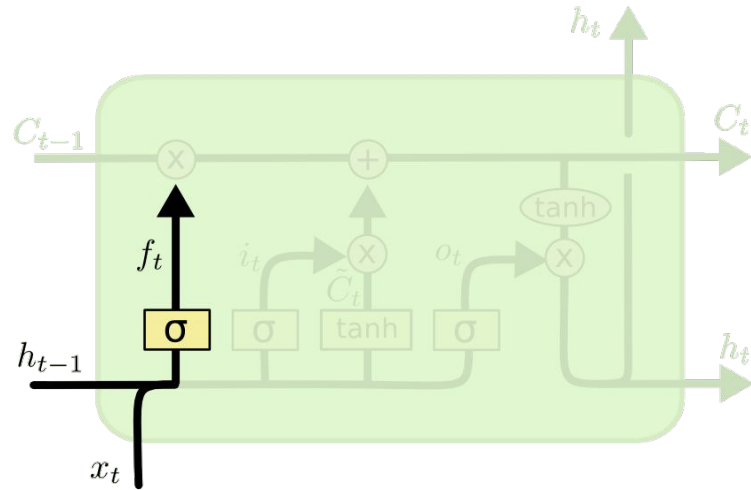


Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks



Fully connected layer

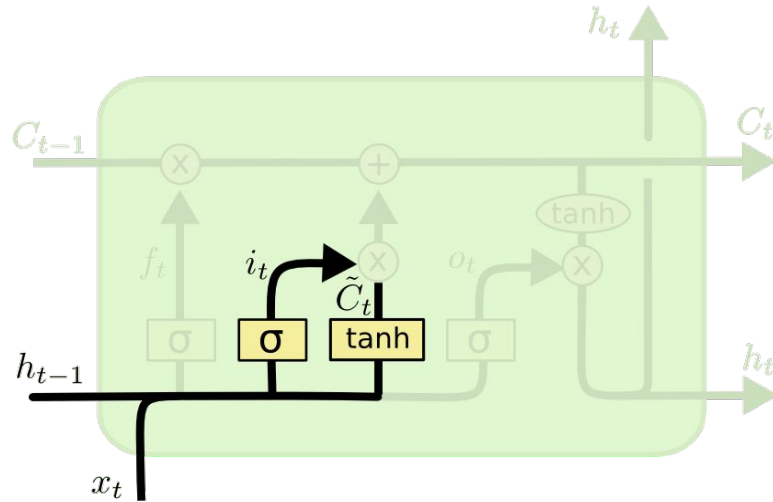
$$f_t = \sigma (W_f \cdot [h_{t-1}, x_t] + b_f)$$

“Forget” gate

Concatenate

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks



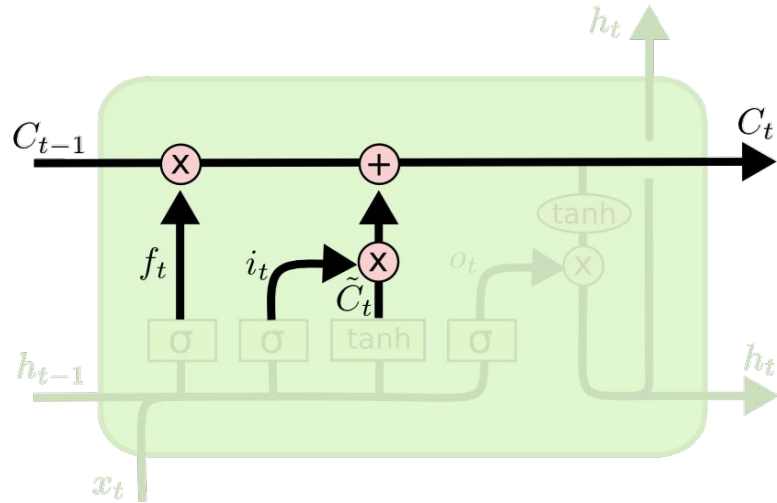
“Input” gate Fully connected layer

$$i_t = \sigma (W_i \cdot [h_{t-1}, x_t] + b_i)$$
$$\tilde{C}_t = \tanh(W_C \cdot [h_{t-1}, x_t] + b_C)$$

New “candidate” values that could be added to modify cell state Fully connected layer

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks



New cell state

$$C_t = f_t * C_{t-1} + i_t * \tilde{C}_t$$

Forget gate

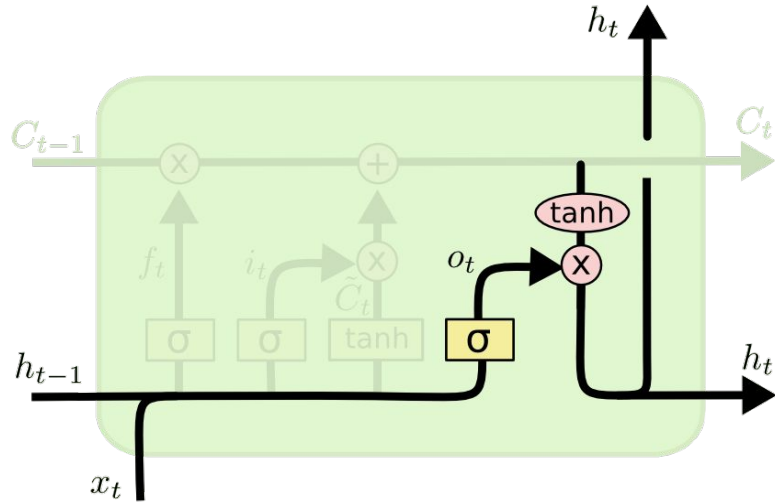
Previous cell state

Input gate

New candidate values

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks



“Output” gate Fully connected layer

$$o_t = \sigma (W_o [h_{t-1}, x_t] + b_o)$$

$$h_t = o_t * \tanh (C_t)$$

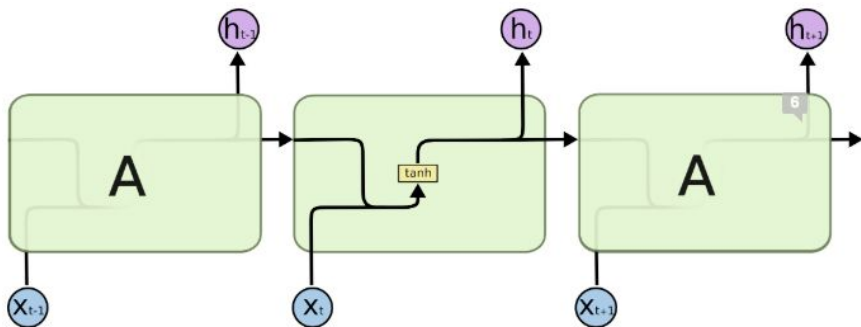
New LSTM
layer output h_t

Current cell
state

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



Unrolled LSTM

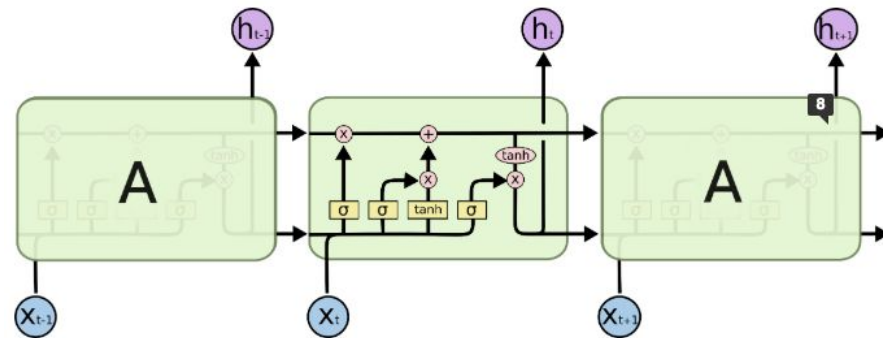
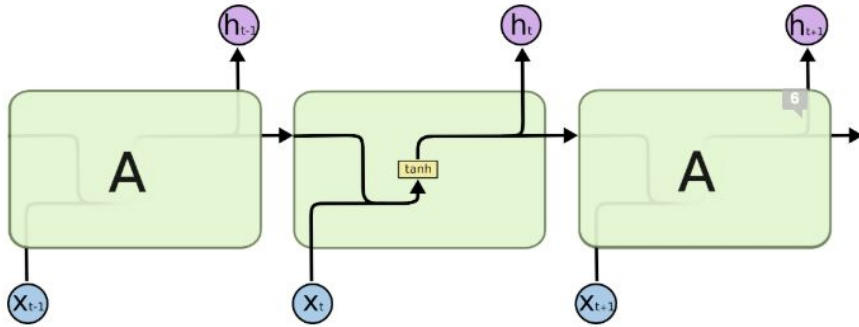


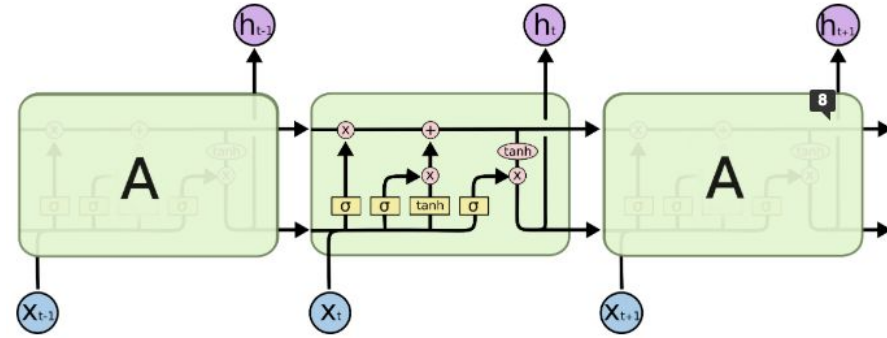
Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



Unrolled LSTM

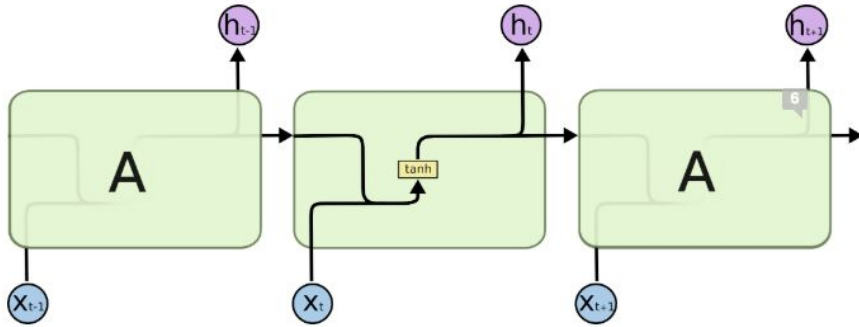


Usage of a “cell state” in the LSTM that is modified through addition allows improved gradient flow through longer sequences.

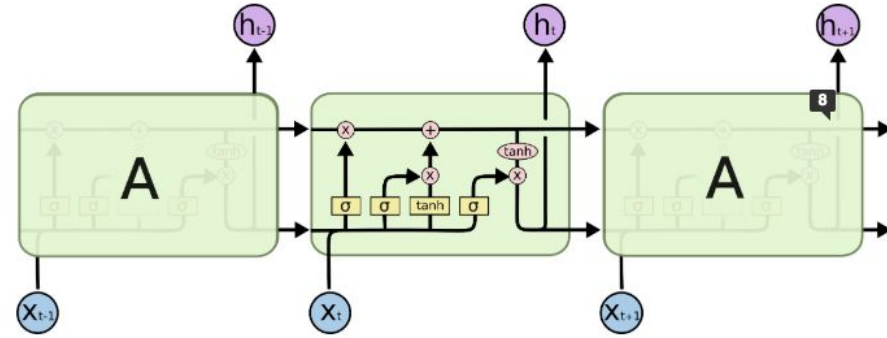
Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



Unrolled LSTM



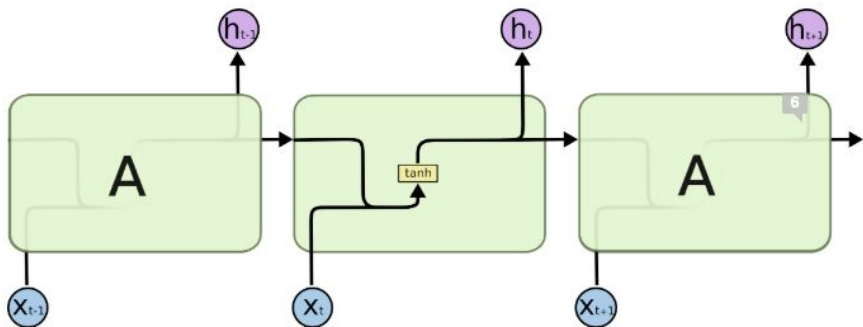
LSTM often used over Vanilla RNN in practice.

Usage of a “cell state” in the LSTM that is modified through addition allows improved gradient flow through longer sequences.

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

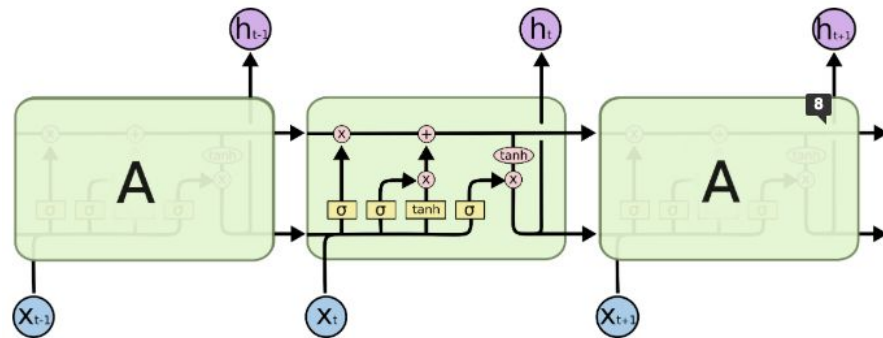
Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



Usage of a “cell state” in the LSTM that is modified through addition allows improved gradient flow through longer sequences.

Unrolled LSTM



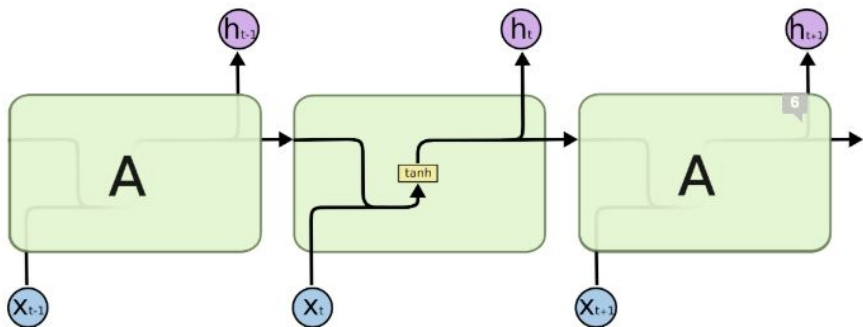
LSTM often used over Vanilla RNN in practice.

Will also see other variants e.g. GRUs with different gating operations.

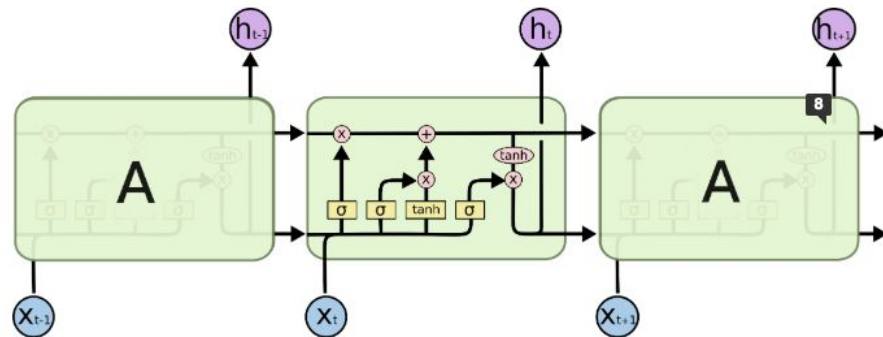
Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Long Short Term Memory (LSTM) Recurrent Networks

Unrolled Vanilla RNN



Unrolled LSTM



Can have multi-layer RNNs and LSTMs, where the $\{h\}$ outputs of one layer form the input sequence for the next layer. One or two layers is common.

Figure credit: <https://colah.github.io/posts/2015-08-Understanding-LSTMs/>

Harutyunyan et al.

- Benchmarked LSTMs vs logistic regression on common prediction tasks using MIMIC-III data
- In-hospital mortality, decompensation, length-of-stay, phenotype classification
- Used a subset of 17 clinical variables from MIMIC-III

Variable	MIMIC-III table	Impute value	Modeled as
Capillary refill rate	chartevents	0.0	categorical
Diastolic blood pressure	chartevents	59.0	continuous
Fraction inspired oxygen	chartevents	0.21	continuous
Glasgow coma scale eye opening	chartevents	4 spontaneously	categorical
Glasgow coma scale motor response	chartevents	6 obeys commands	categorical
Glasgow coma scale total	chartevents	15	categorical
Glasgow coma scale verbal response	chartevents	5 oriented	categorical
Glucose	chartevents, labevents	128.0	continuous
Heart Rate	chartevents	86	continuous
Height	chartevents	170.0	continuous
Mean blood pressure	chartevents	77.0	continuous
Oxygen saturation	chartevents, labevents	98.0	continuous
Respiratory rate	chartevents	19	continuous
Systolic blood pressure	chartevents	118.0	continuous
Temperature	chartevents	36.6	continuous
Weight	chartevents	81.0	continuous
pH	chartevents, labevents	7.4	continuous

Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

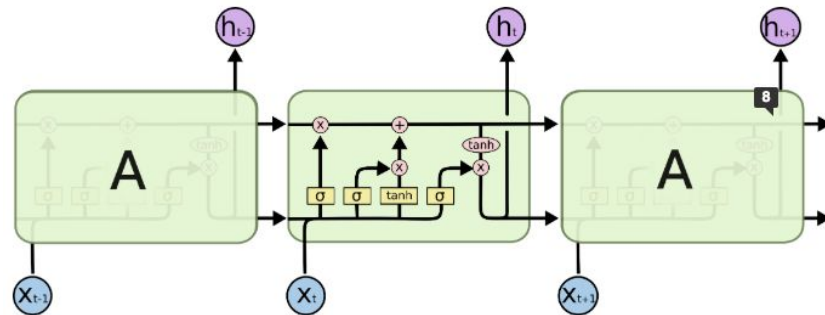
Harutyunyan et al.

- **Logistic regression models**

- Use hand-engineered feature vector to represent a time-series: min, max, mean, std dev, etc. of each feature in several subsequences (full series, first 10% of series, first 50%, last 10%, etc.)
- If feature does not occur in subsequence ([missing data](#)), impute with mean value from training set
- Categorical variables had meaningful numeric values -> no change
- Zero-mean unit-variance standardization of all features

Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.



- LSTM models

- Bucket time series into regularly spaced intervals, take the value (or last value, if multiple) of each variable in the interval to create observation x_t
- Encode categorical variables using a one-hot vector (vector of 0s with a 1 in the observed position).
- If variable is missing in a time bucket, impute using most recent observed measurement if it exists, and mean value from training set otherwise
- Concat the values of each clinical variable with a binary mask indicating presence or not (i.e., missing and needed to impute) to form full observation feature vector x_t

Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: in-hospital mortality

- Input: Time-series data for first 48 hours of ICU stay
- Output: binary classification of in-hospital mortality

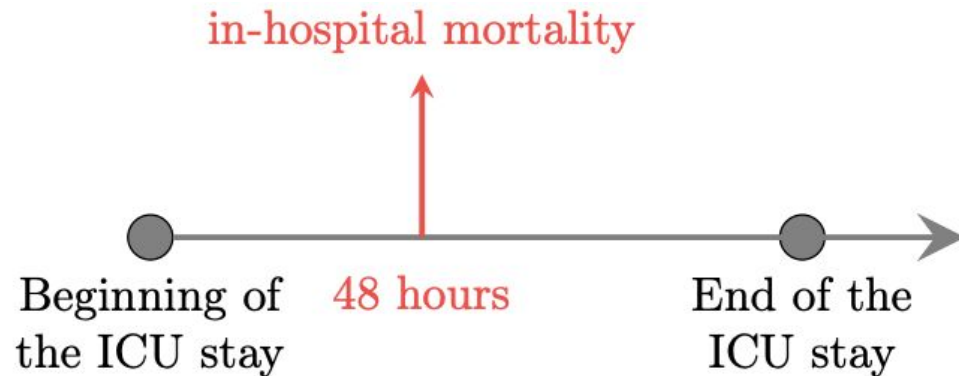


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: in-hospital mortality

- Input: Time-series data for first 48 hours of ICU stay
- Output: binary classification of in-hospital mortality

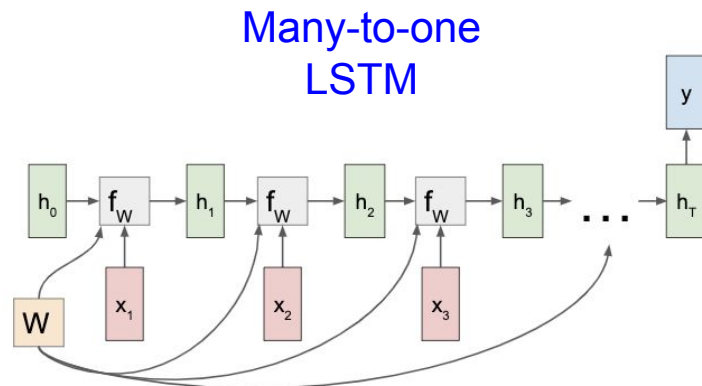
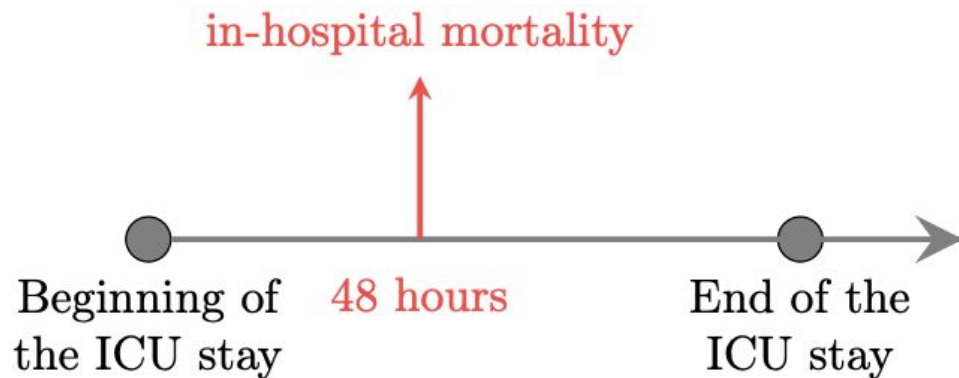


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: decompensation

- Input: Time-series data from beginning of stay until prediction time (every hour)
- Output: Binary classification of mortality in the next 24 hours

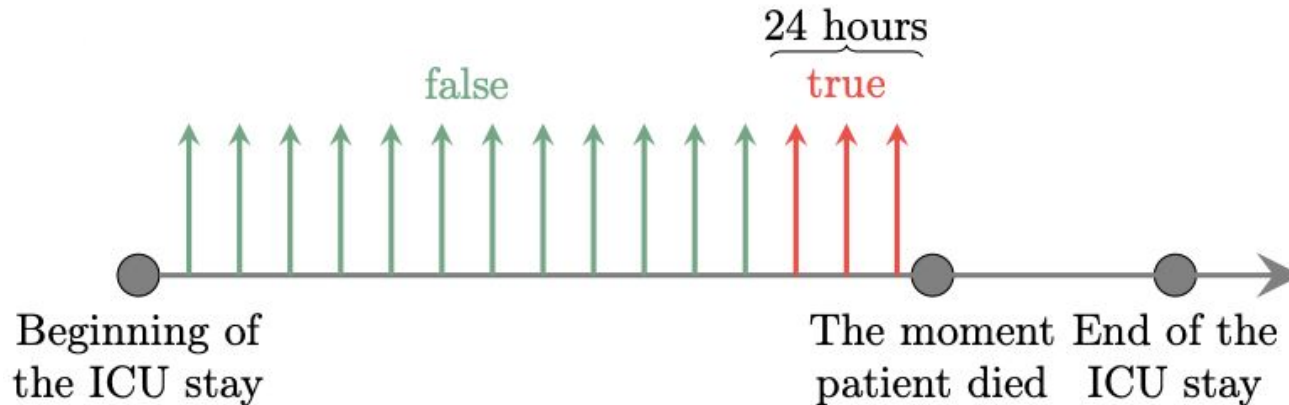


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: decompensation

- Input: Time-series data from beginning of stay until prediction time (every hour)
- Output: Binary classification of mortality in the next 24 hours

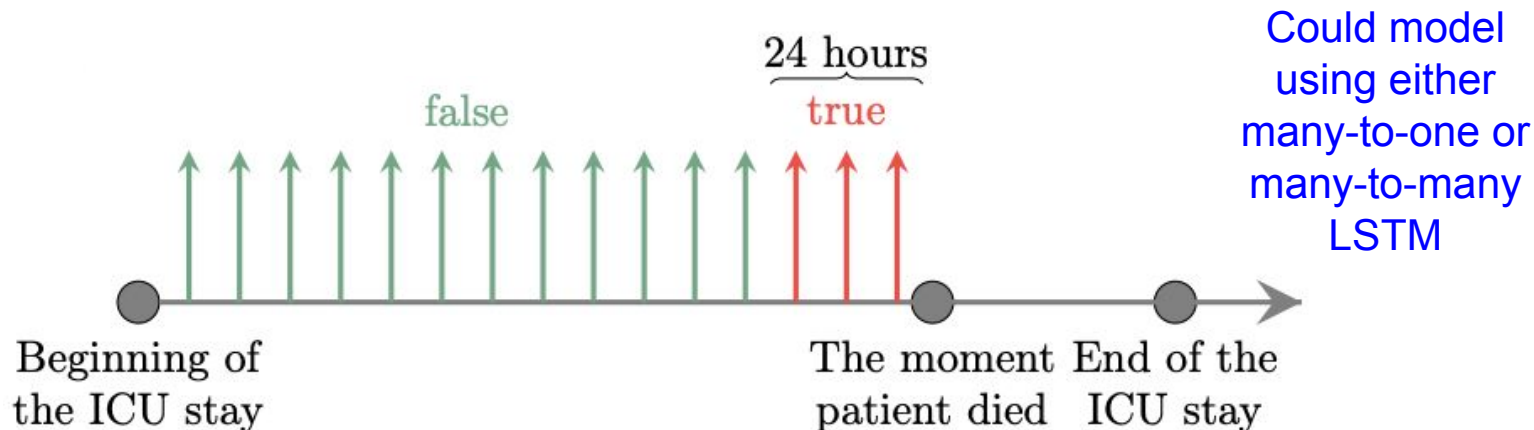
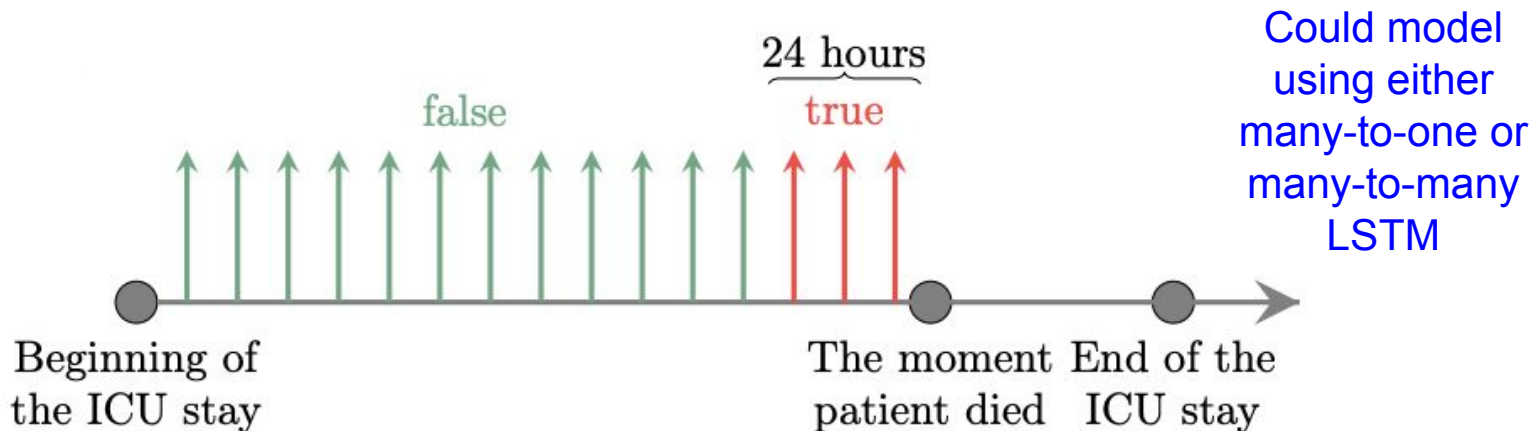


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: decompensation

- Input: Time-series data from beginning of stay until prediction time (every hour)
- Output: Binary classification of mortality in the next 24 hours



Closely related to in-hospital mortality

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: length-of-stay

- Input: Time-series data from beginning of stay until prediction time (every hour)
- Output: remaining time spent in ICU. Model as classification problem: ICU stays < 1 day, each of 7 days, between 1-2 weeks, > 2 weeks

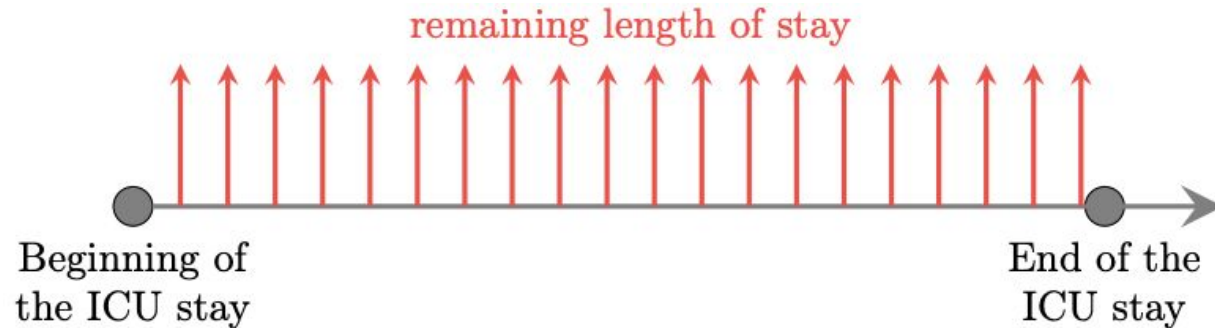
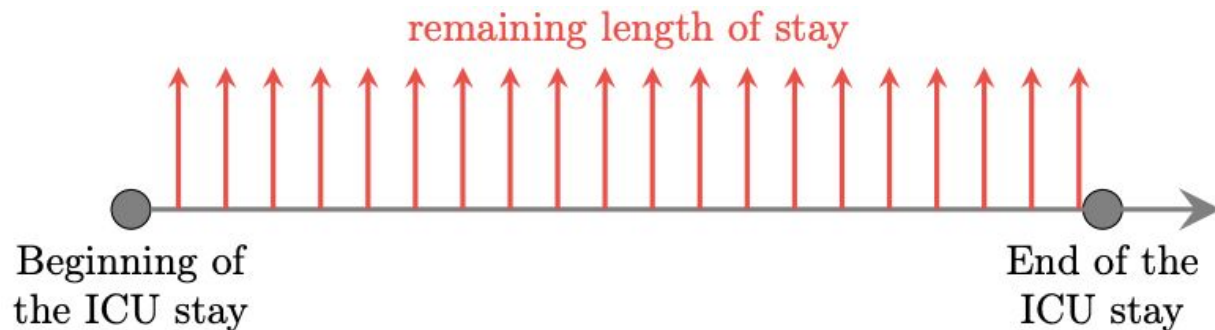


Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: length-of-stay

- Input: Time-series data from beginning of stay until prediction time (every hour)
- Output: remaining time spent in ICU. Model as classification problem: ICU stays < 1 day, each of 7 days, between 1-2 weeks, > 2 weeks



Can model problem in different ways, e.g. directly regress LOS value, or predict meaningful category of extended LOS (>7 days)

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: phenotypes

- Input: Time-series data corresponding to entire ICU stay
- Output: Multilabel classification of the presence of 25 acute care conditions (merged from ICD codes) in stay record



Phenotype

Acute and unspecified renal failure
Acute cerebrovascular disease
Acute myocardial infarction
Cardiac dysrhythmias
Chronic kidney disease
Chronic obstructive pulmonary disease
Complications of surgical/medical care
Conduction disorders
Congestive heart failure; nonhypertensive
Coronary atherosclerosis and related
Diabetes mellitus with complications
Diabetes mellitus without complication
Disorders of lipid metabolism

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: phenotypes

- Input: Time-series data corresponding to entire ICU stay
- Output: Multilabel classification of the presence of 25 acute care conditions (merged from ICD codes) in stay record

Q: Why do we formulate this as a multi-label classification task?



Phenotype

Acute and unspecified renal failure
Acute cerebrovascular disease
Acute myocardial infarction
Cardiac dysrhythmias
Chronic kidney disease
Chronic obstructive pulmonary disease
Complications of surgical/medical care
Conduction disorders
Congestive heart failure; nonhypertensive
Coronary atherosclerosis and related
Diabetes mellitus with complications
Diabetes mellitus without complication
Disorders of lipid metabolism

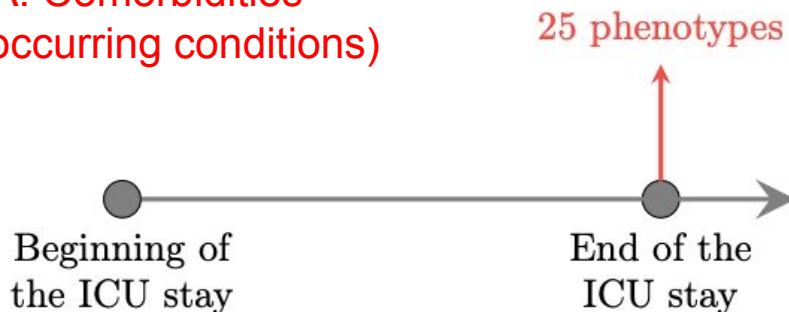
Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: phenotypes

- Input: Time-series data corresponding to entire ICU stay
- Output: Multilabel classification of the presence of 25 acute care conditions (merged from ICD codes) in stay record

Q: Why do we formulate this as a multi-label classification task?

A: Comorbidities (co-occurring conditions)



Phenotype

Acute and unspecified renal failure
Acute cerebrovascular disease
Acute myocardial infarction
Cardiac dysrhythmias
Chronic kidney disease
Chronic obstructive pulmonary disease
Complications of surgical/medical care
Conduction disorders
Congestive heart failure; nonhypertensive
Coronary atherosclerosis and related
Diabetes mellitus with complications
Diabetes mellitus without complication
Disorders of lipid metabolism

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

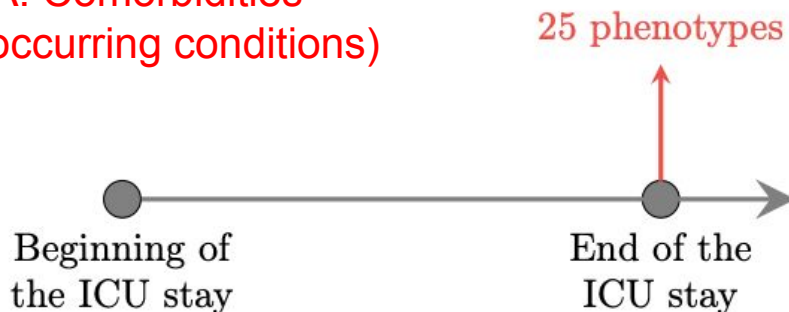
Harutyunyan et al.: phenotypes

- Input: Time-series data corresponding to entire ICU stay
- Output: Multilabel classification of the presence of 25 acute care conditions (merged from ICD codes) in stay record

Q: Why do we formulate this as a multi-label classification task?

A: Comorbidities (co-occurring conditions)

Q: What loss function should we use?



Phenotype

Acute and unspecified renal failure
Acute cerebrovascular disease
Acute myocardial infarction
Cardiac dysrhythmias
Chronic kidney disease
Chronic obstructive pulmonary disease
Complications of surgical/medical care
Conduction disorders
Congestive heart failure; nonhypertensive
Coronary atherosclerosis and related
Diabetes mellitus with complications
Diabetes mellitus without complication
Disorders of lipid metabolism

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: phenotypes

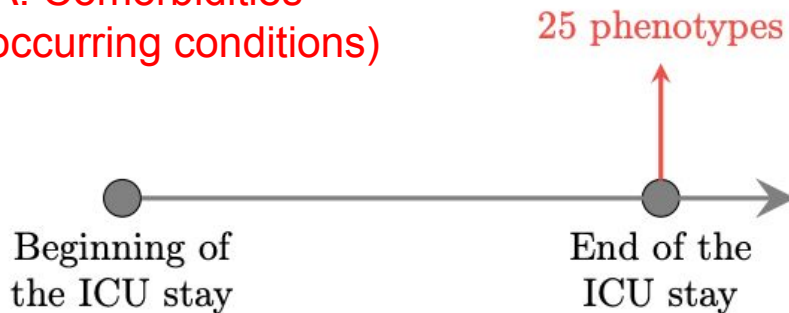
- Input: Time-series data corresponding to entire ICU stay
- Output: Multilabel classification of the presence of 25 acute care conditions (merged from ICD codes) in stay record

Q: Why do we formulate this as a multi-label classification task?

A: Comorbidities (co-occurring conditions)

Q: What loss function should we use?

A: Multiple binary cross-entropy losses



Phenotype

Acute and unspecified renal failure
Acute cerebrovascular disease
Acute myocardial infarction
Cardiac dysrhythmias
Chronic kidney disease
Chronic obstructive pulmonary disease
Complications of surgical/medical care
Conduction disorders
Congestive heart failure; nonhypertensive
Coronary atherosclerosis and related
Diabetes mellitus with complications
Diabetes mellitus without complication
Disorders of lipid metabolism

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: logistic regression vs LSTMs

Found better performance overall for LSTMs (S) vs logistic regression (LR). Also introduced more sophisticated variants and multi-task training (joint training of all tasks together).

In-hospital Mortality	Model	AUC-ROC	Phenotyping	Model	Macro AUC-ROC
	SAPS	0.720 (0.720, 0.720)		LR	0.739 (0.734, 0.743)
	APS-III	0.750 (0.750, 0.750)		S	0.770 (0.766, 0.775)
	OASIS	0.760 (0.760, 0.761)		S + DS	0.774 (0.769, 0.778)
	SAPS-II	0.777 (0.776, 0.777)		C	0.776 (0.772, 0.781)
	LR	0.848 (0.828, 0.868)		C + DS	0.773 (0.769, 0.777)
	S	0.855 (0.835, 0.873)		MS	0.768 (0.763, 0.772)
	S + DS	0.856 (0.836, 0.875)		MC	0.774 (0.770, 0.778)
	C	0.862 (0.844, 0.881)			
	C + DS	0.854 (0.834, 0.873)			
MS	0.861 (0.842, 0.878)				
MC	0.870 (0.852, 0.887)				

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Harutyunyan et al.: logistic regression vs LSTMs

Found better performance overall for LSTMs (S) vs logistic regression (LR). Also introduced more sophisticated variants and multi-task training (joint training of all tasks together).

In-hospital Mortality	Model	AUC-ROC	Phenotyping	Model	Macro AUC-ROC
	SAPS	0.720 (0.720, 0.720)		LR	0.739 (0.734, 0.743)
	APS-III	0.750 (0.750, 0.750)		S	0.770 (0.766, 0.775)
	OASIS	0.760 (0.760, 0.761)		S + DS	0.774 (0.769, 0.778)
	SAPS-II	0.777 (0.776, 0.777)		C	0.776 (0.772, 0.781)
	LR	0.848 (0.828, 0.868)		C + DS	0.773 (0.769, 0.777)
	S	0.855 (0.835, 0.873)		MS	0.768 (0.763, 0.772)
	S + DS	0.856 (0.836, 0.875)		MC	0.774 (0.770, 0.778)
	C	0.862 (0.844, 0.881)			
	C + DS	0.854 (0.834, 0.873)			

Found better performance for phenotyping acute vs chronic conditions -- makes sense!

Figure credit: Harutyunyan et al. Multitask learning and benchmarking with clinical time series data. 2019.

Summary:

- Introduction to EHRs
- EHR prediction tasks
- Recurrent neural networks and LSTMs

Next:

- More on EHR data
- More on feature representations and model interpretability