

**Project Proposal deadline:** tonight, 11:59pm

**Course Notes:** <https://snap-stanford.github.io/cs224w-notes/>  
Help us write the course notes – we will give **generous bonuses!**

# Graph Neural Networks

CS224W: Machine Learning with Graphs

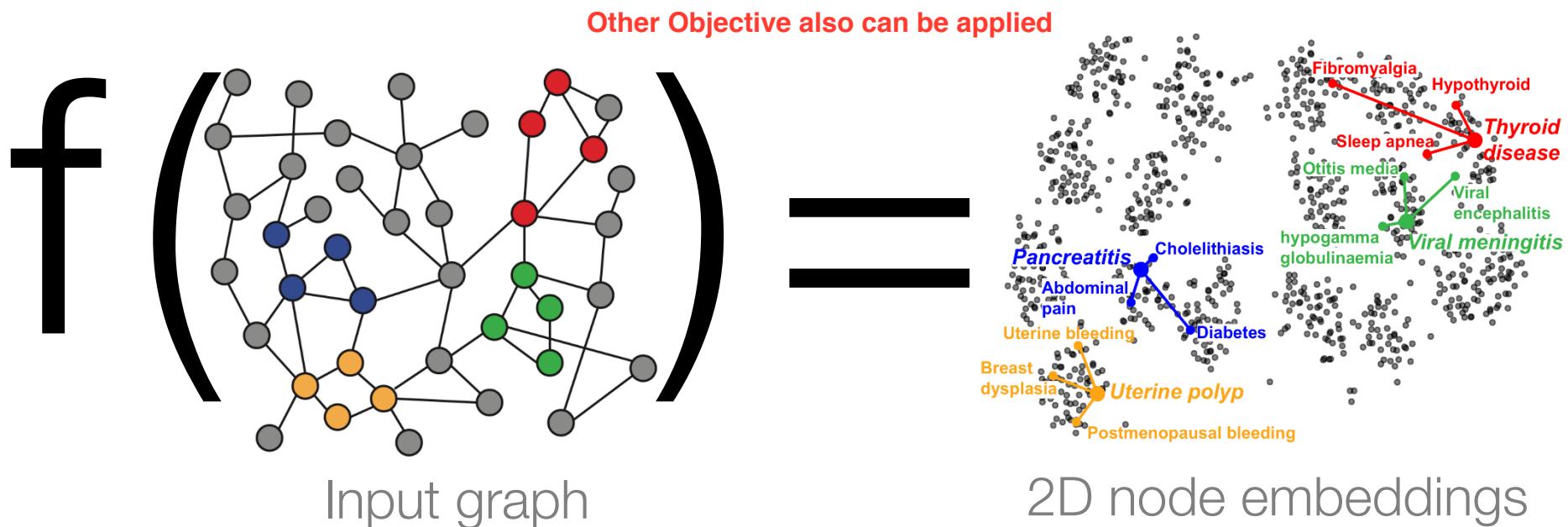
Jure Leskovec, Stanford University

<http://cs224w.stanford.edu>



# Node Embeddings

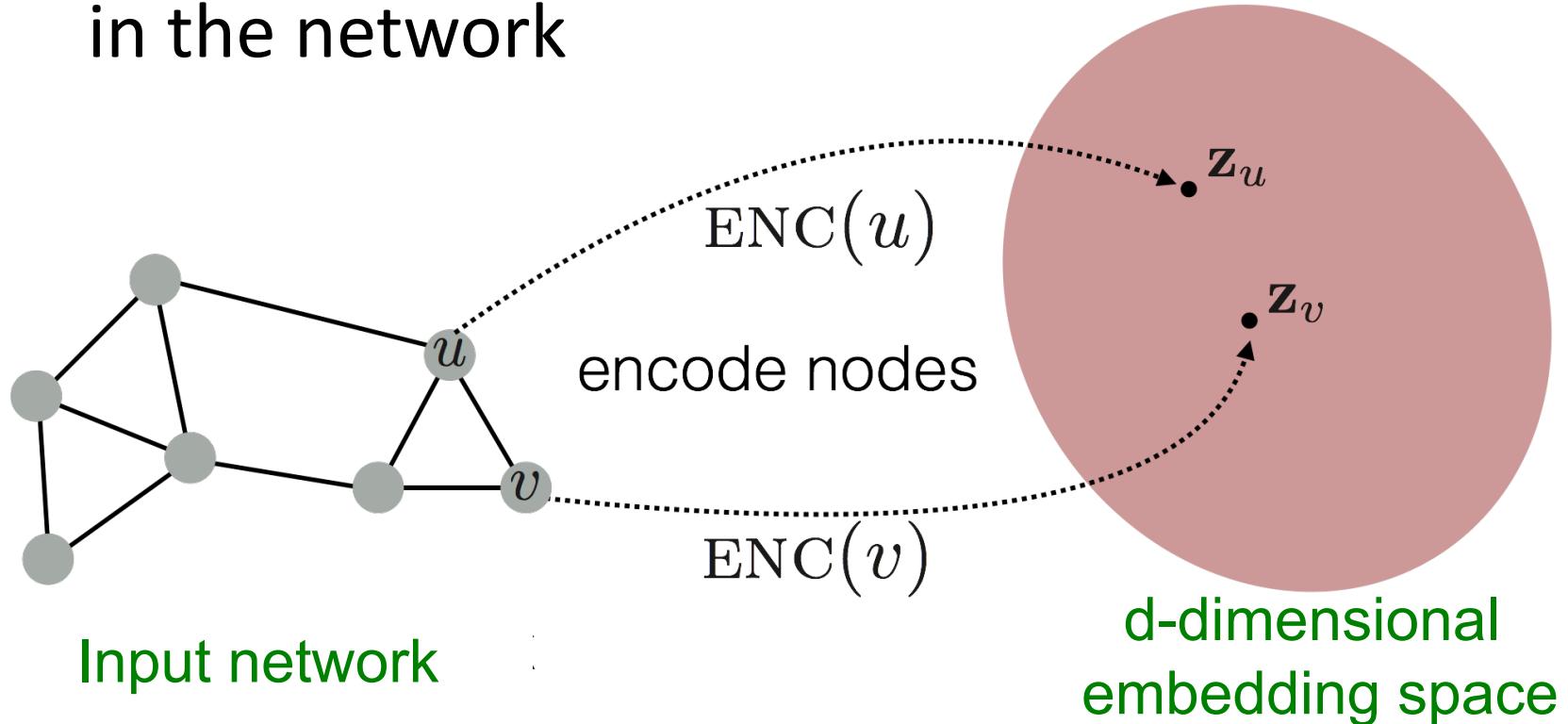
- **Intuition:** Map nodes to  $d$ -dimensional embeddings such that similar nodes in the graph are embedded close together



How to learn mapping function  $f$ ?

# Node Embeddings

- **Goal:** Map nodes so that similarity in the embedding space (e.g., dot product) approximates similarity (e.g., proximity) in the network



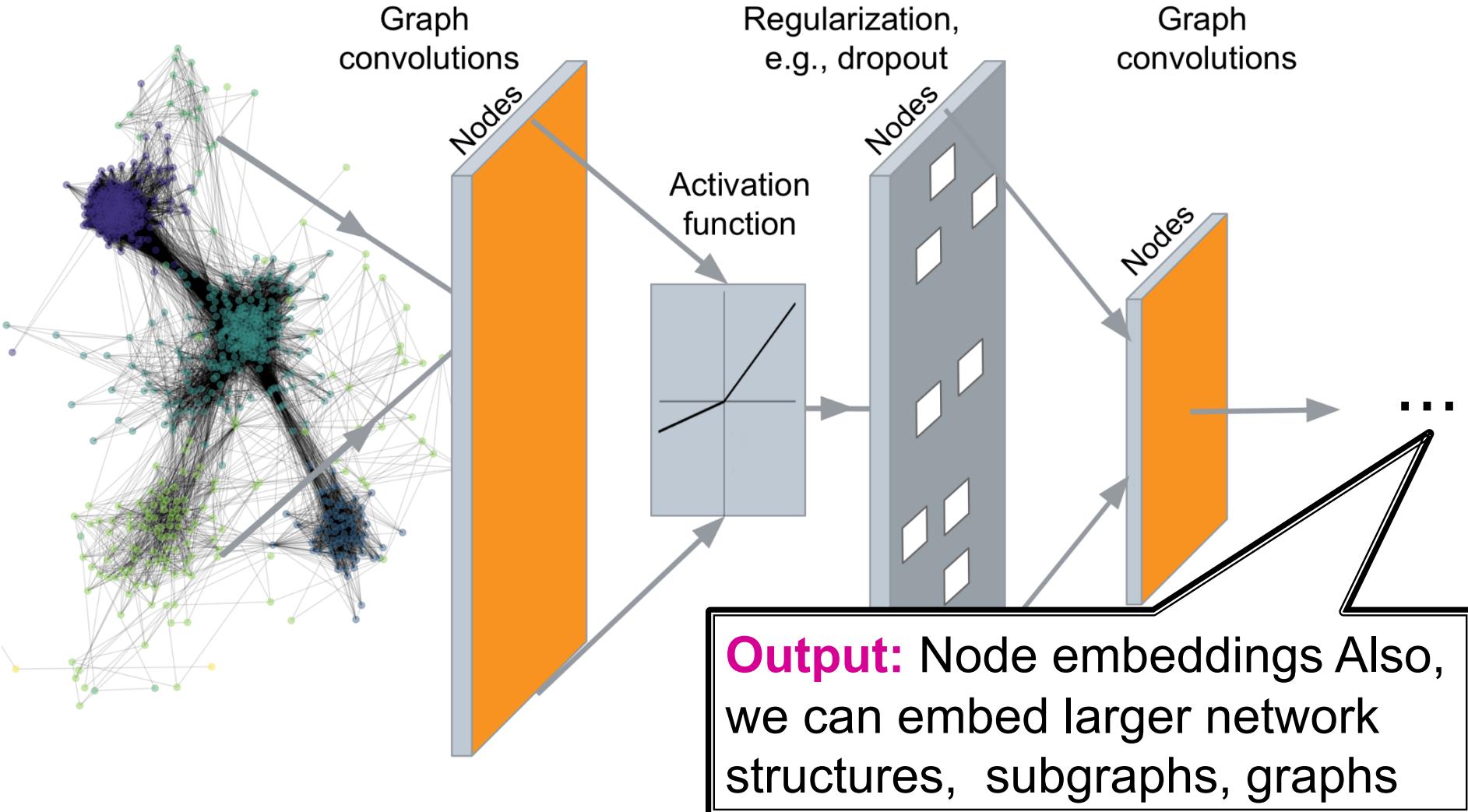
# Today: Deep Graph Encoders

- **Today:** We will now discuss deep methods based on **graph neural networks**:

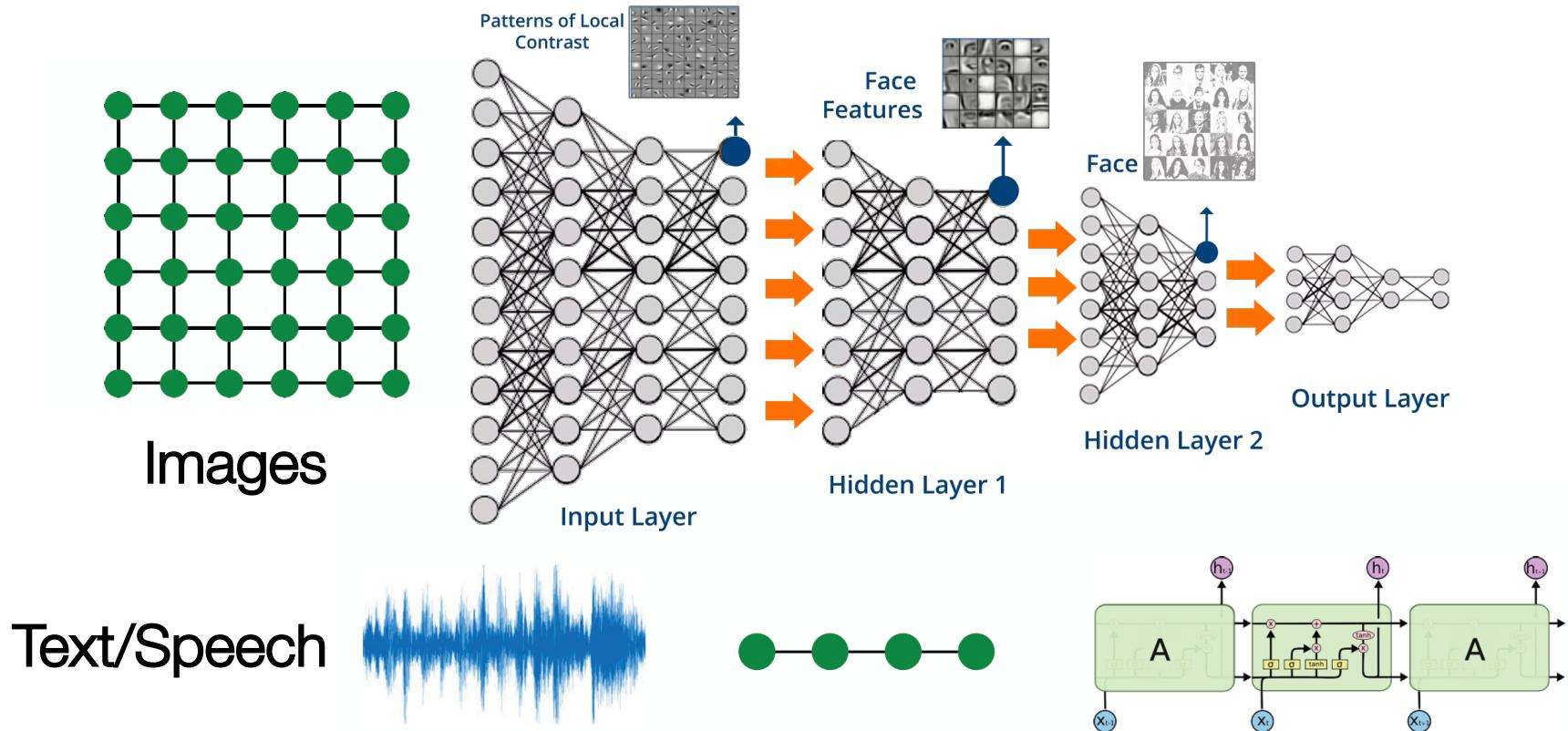
$$\text{ENC}(v) = \begin{matrix} & \text{multiple layers of} \\ & \text{non-linear transformations} \\ & \text{of graph structure} \end{matrix}$$

- **Note:** All these deep encoders can be **combined with node similarity functions** defined in the last lecture

# Deep Graph Encoders



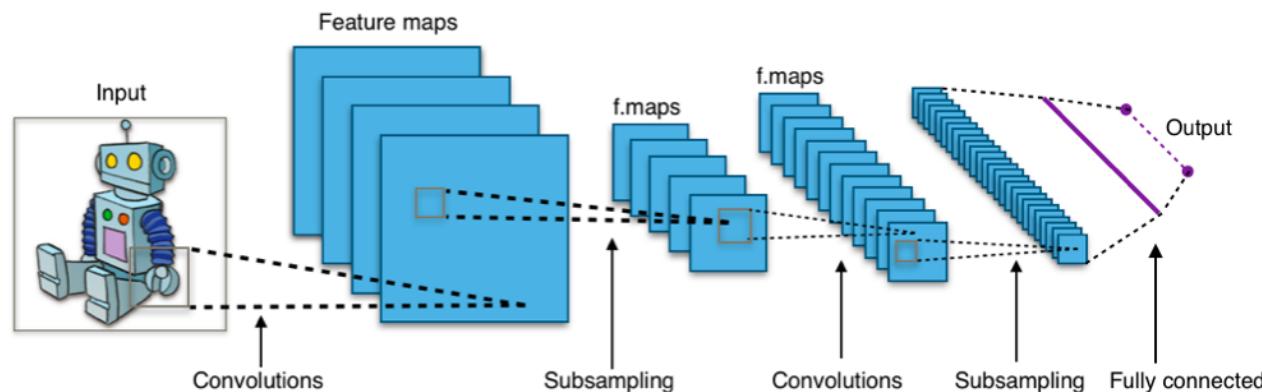
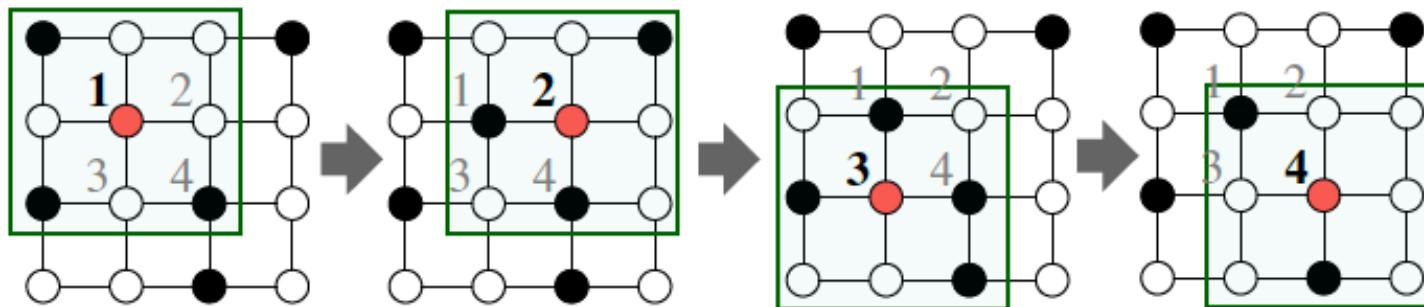
# Modern ML Toolbox



Modern deep learning toolbox is designed for simple sequences & grids

# Idea: Convolutional Networks

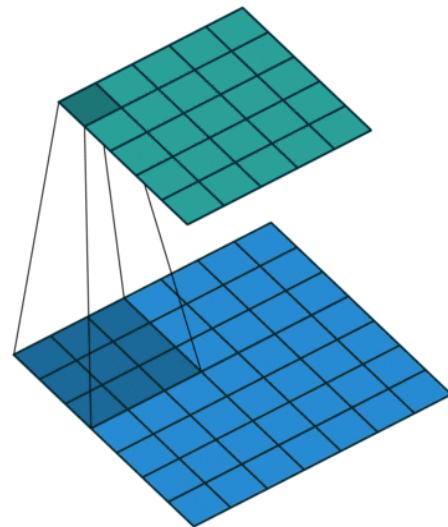
CNN on an image:



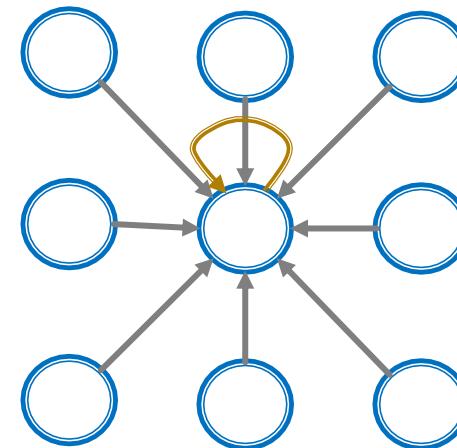
Goal is to generalize convolutions beyond simple lattices  
Leverage node features/attributes (e.g., text, images)

# From Images to Graphs

Single CNN layer with 3x3 filter:



Image



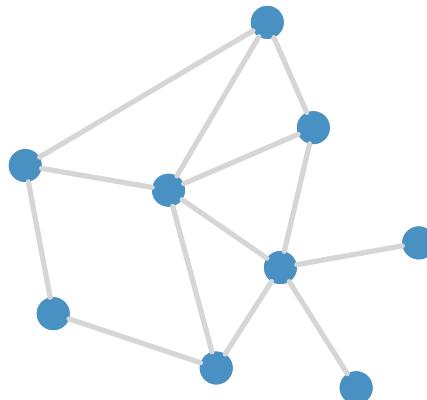
Graph

Transform information at the neighbors and combine it:

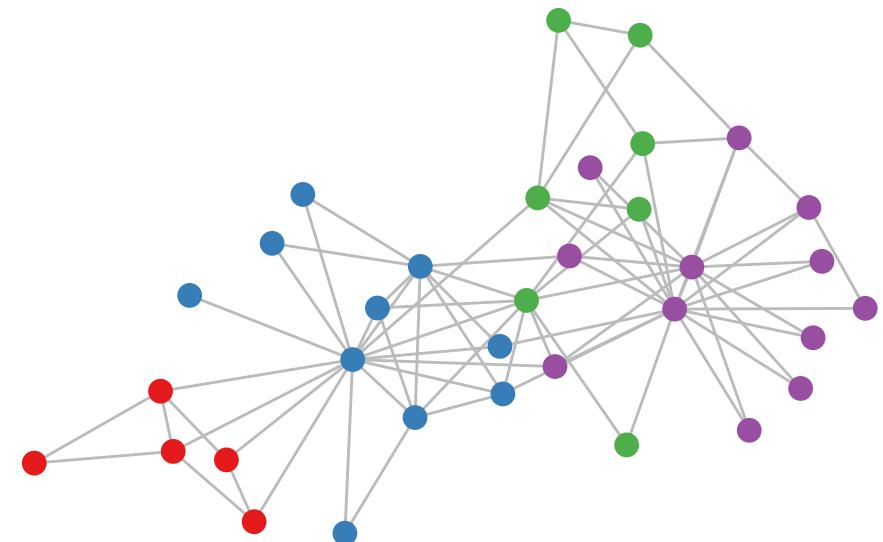
- Transform “messages”  $h_i$  from neighbors:  $W_i h_i$
- Add them up:  $\sum_i W_i h_i$

# Real-World Graphs

But what if your graphs look like this?



or this:



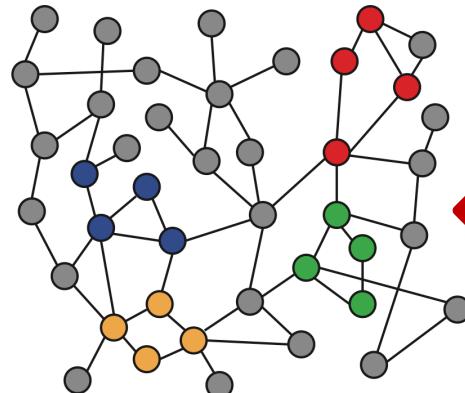
- Examples:

Biological networks, Medical networks, Social networks, Information networks, Knowledge graphs, Communication networks, Web graph, ...

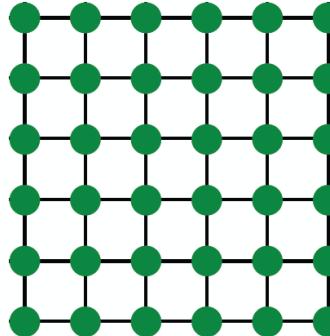
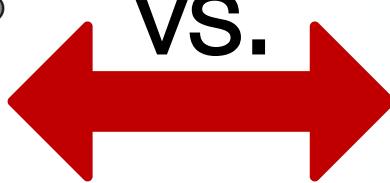
# Why is it Hard?

But networks are far more complex!

- Arbitrary size and complex topological structure (i.e., no spatial locality like grids)



Networks



Images



Text

- No fixed node ordering or reference point
- Often dynamic and have multimodal features

# Outline of Today's Lecture

- 1. Basics of deep learning for graphs**
- 2. Graph Convolutional Networks**
- 3. Graph Attention Networks (GAT)**
- 4. Practical tips and demos**



# Basics of Deep Learning for Graphs

# Content

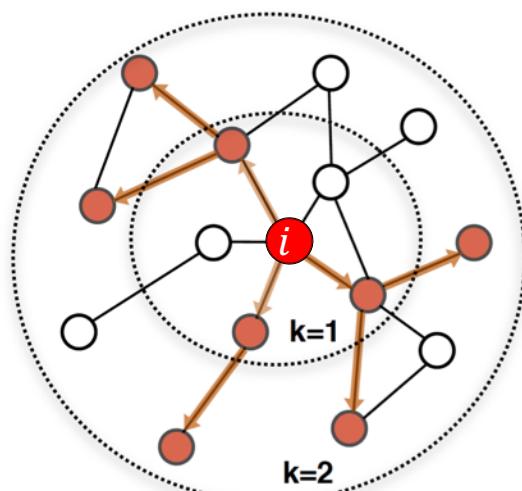
- **Local network neighborhoods:**
  - Describe aggregation strategies
  - Define computation graphs
- **Stacking multiple layers:**
  - Describe the model, parameters, training
  - How to fit the model?
  - Simple example for unsupervised and supervised training

# Setup

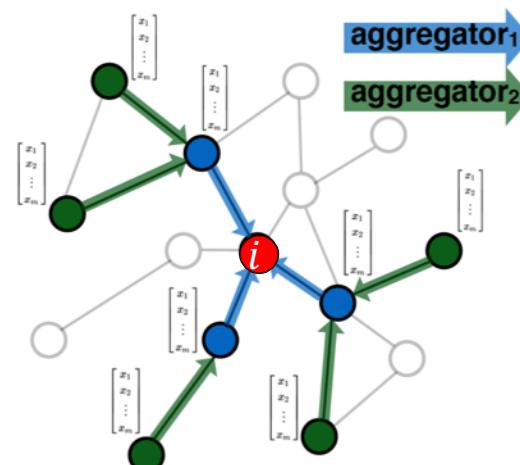
- Assume we have a graph  $G$ :
  - $V$  is the **vertex set**
  - $A$  is the **adjacency matrix** (assume binary)
  - $X \in \mathbb{R}^{m \times |V|}$  is a matrix of **node features**
  - Node features:
    - Social networks: User profile, User image
    - Biological networks: Gene expression profiles, gene functional information
    - No features:
      - Indicator vectors (one-hot encoding of a node)
      - Vector of constant 1:  $[1, 1, \dots, 1]$

# Graph Convolutional Networks

**Idea:** Node's neighborhood defines a computation graph



Determine node computation graph

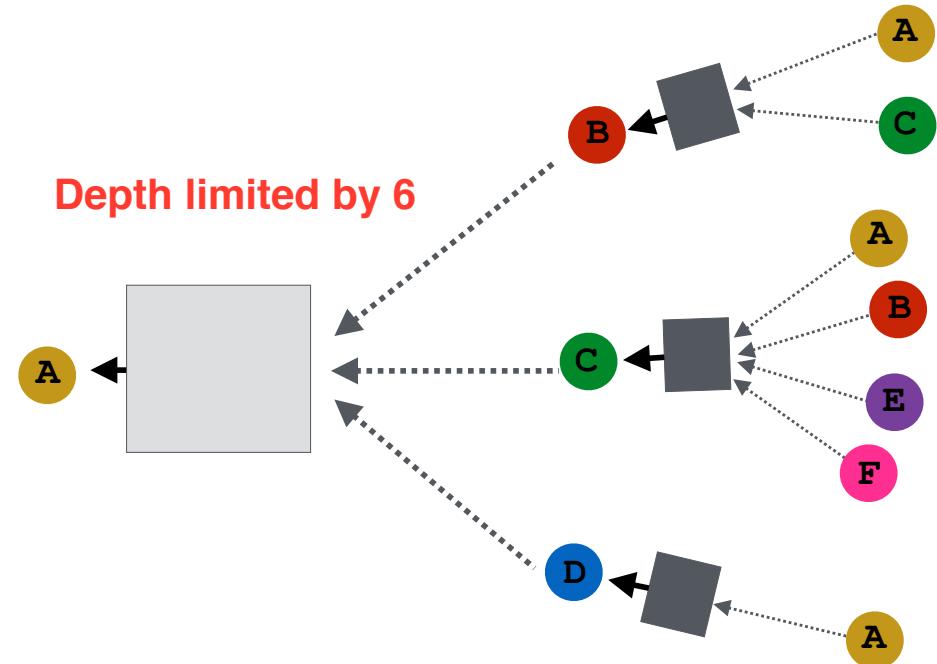
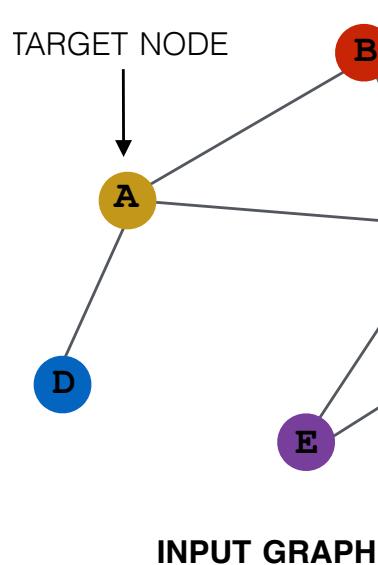


Propagate and transform information

**Learn how to propagate information across the graph to compute node features**

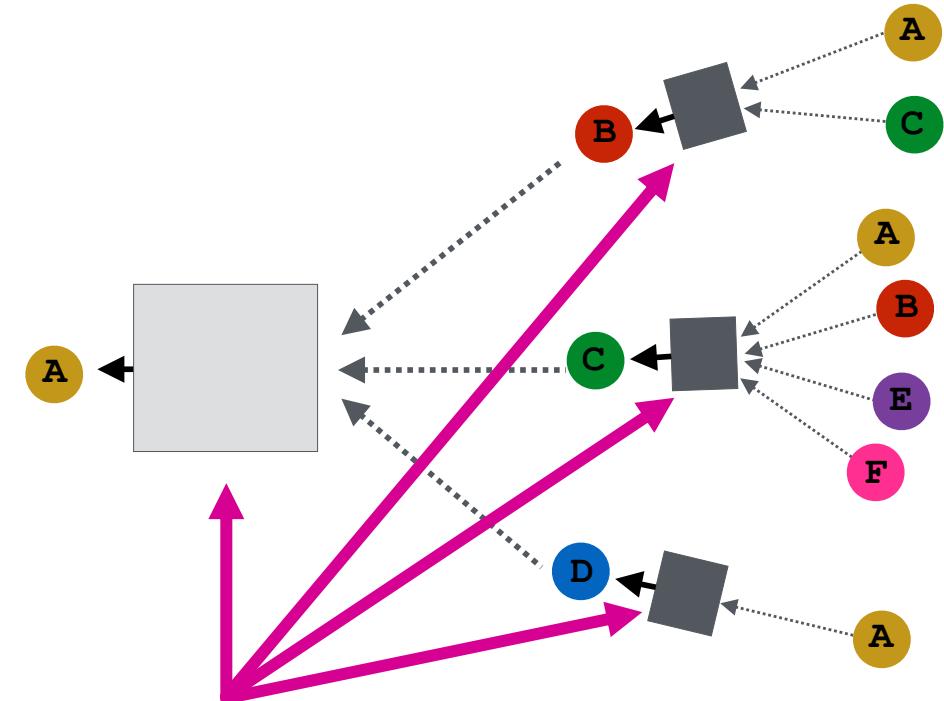
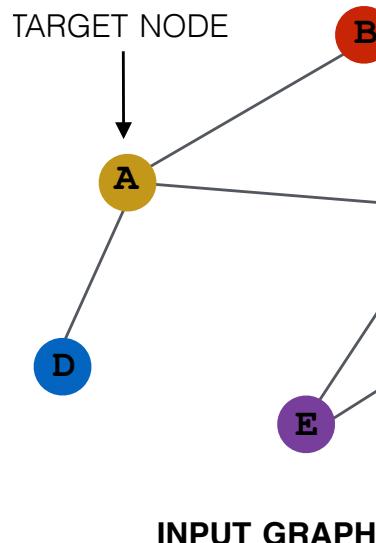
# Idea: Aggregate Neighbors

- **Key idea:** Generate node embeddings based on **local network neighborhoods**



# Idea: Aggregate Neighbors

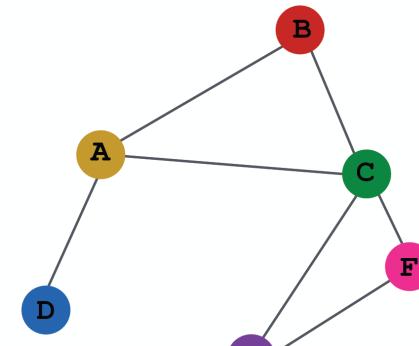
- **Intuition:** Nodes aggregate information from their neighbors using neural networks



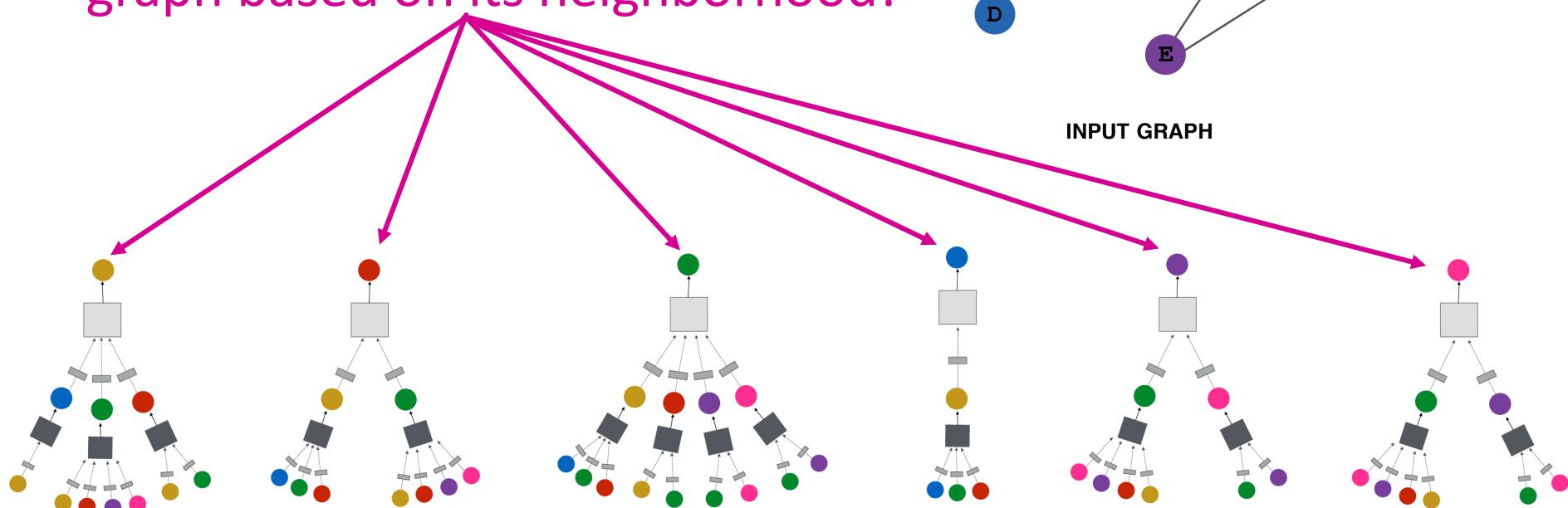
# Idea: Aggregate Neighbors

- **Intuition:** Network neighborhood defines a computation graph

Every node defines a computation graph based on its neighborhood!

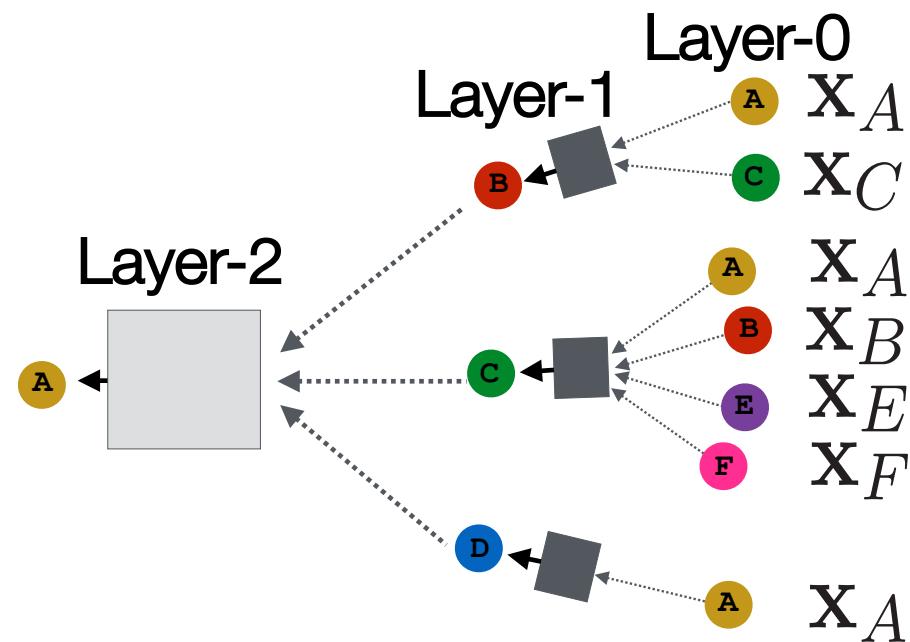
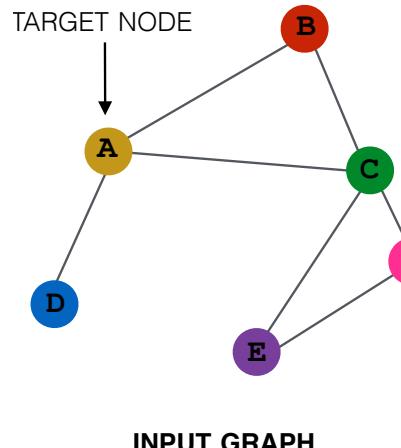


INPUT GRAPH



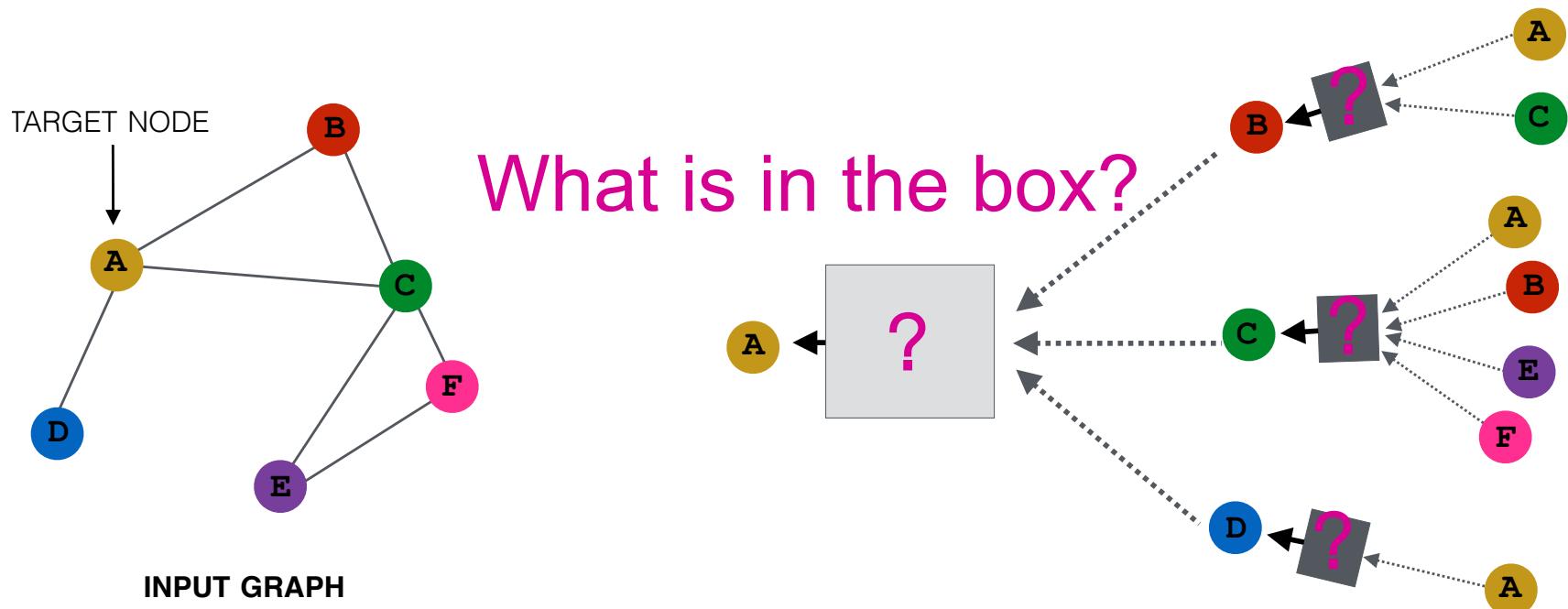
# Deep Model: Many Layers

- Model can be **of arbitrary depth**:
  - Nodes have embeddings at each layer
  - Layer-0 embedding of node  $u$  is its input feature,  $x_u$
  - Layer-K embedding gets information from nodes that are K hops away



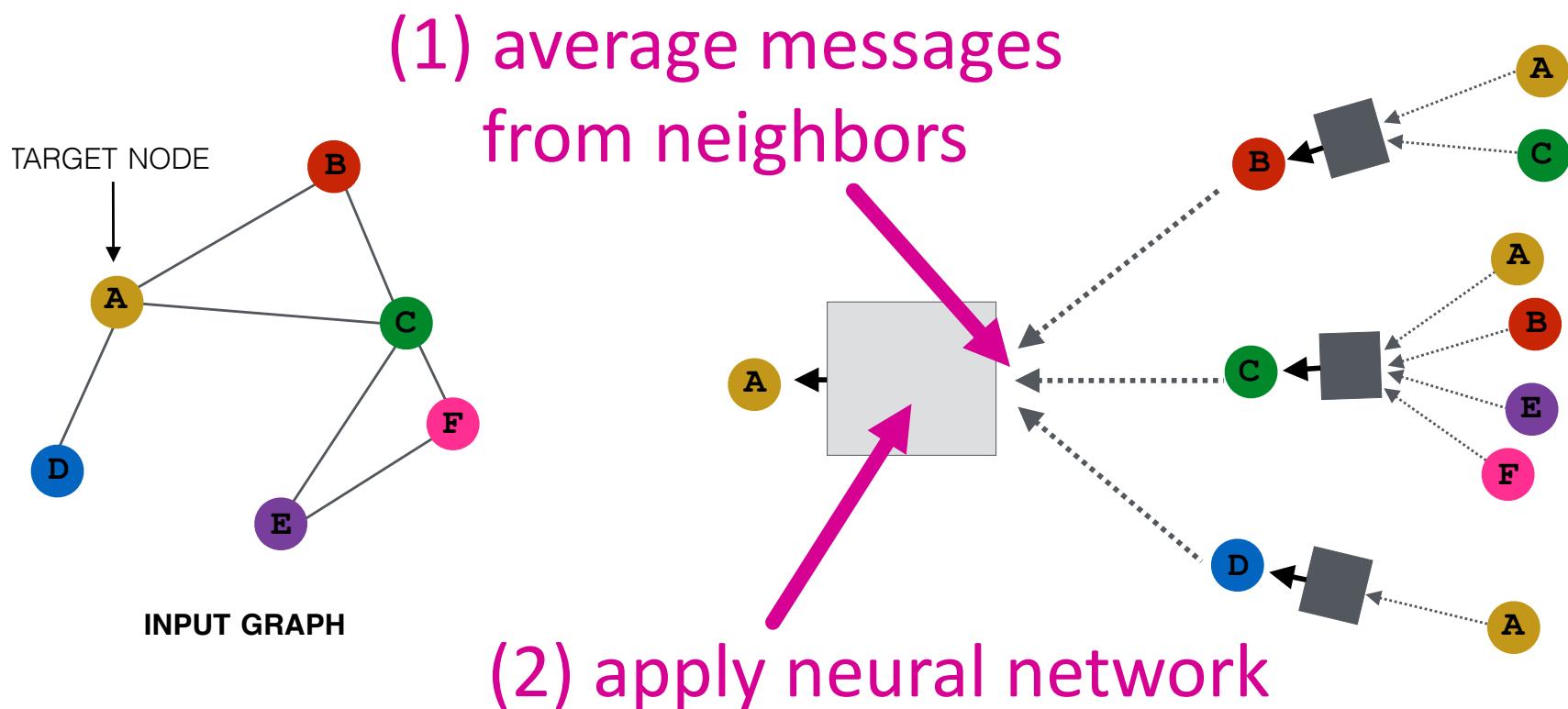
# Neighborhood Aggregation

- **Neighborhood aggregation:** Key distinctions are in how different approaches aggregate information across the layers



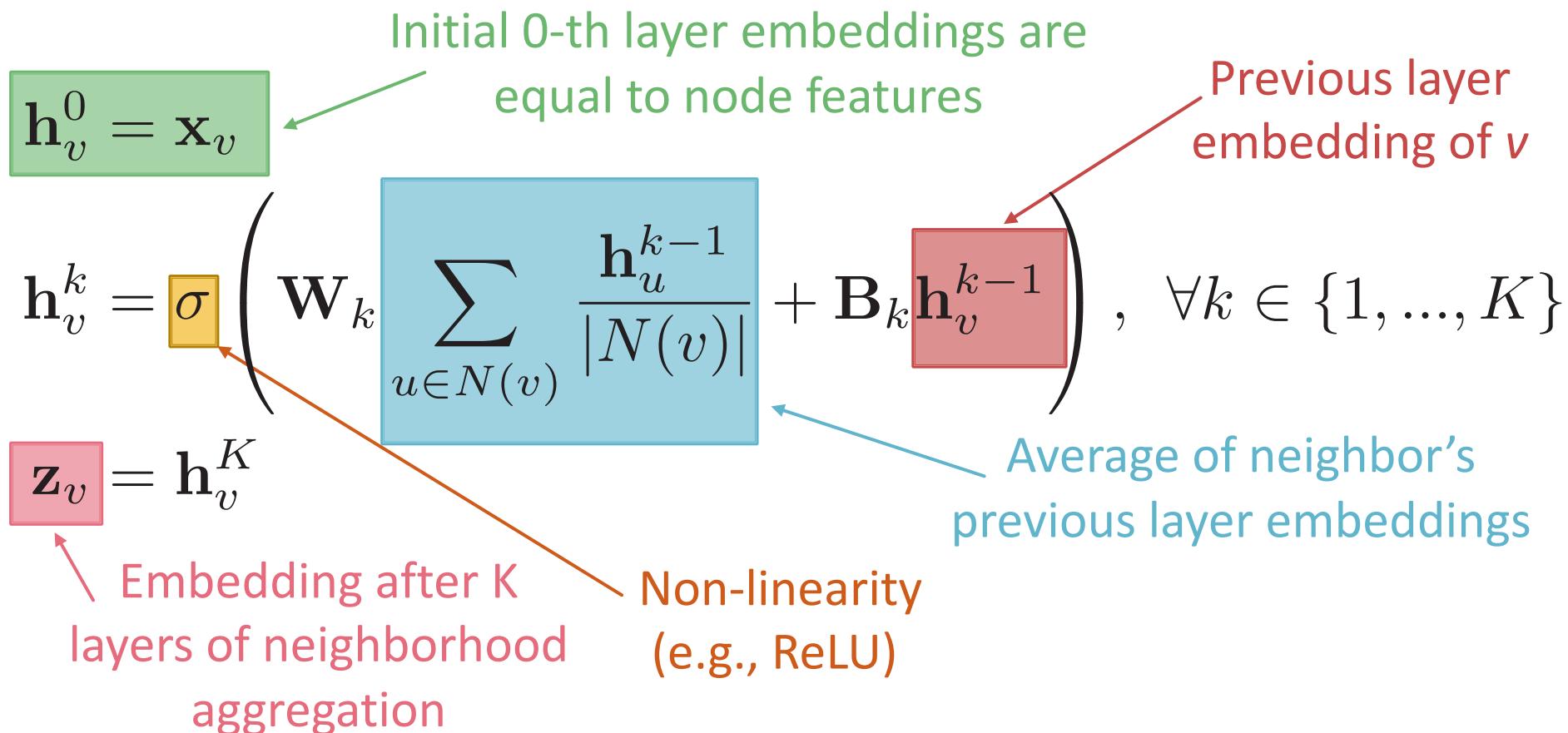
# Neighborhood Aggregation

- **Basic approach:** Average information from neighbors and apply a neural network



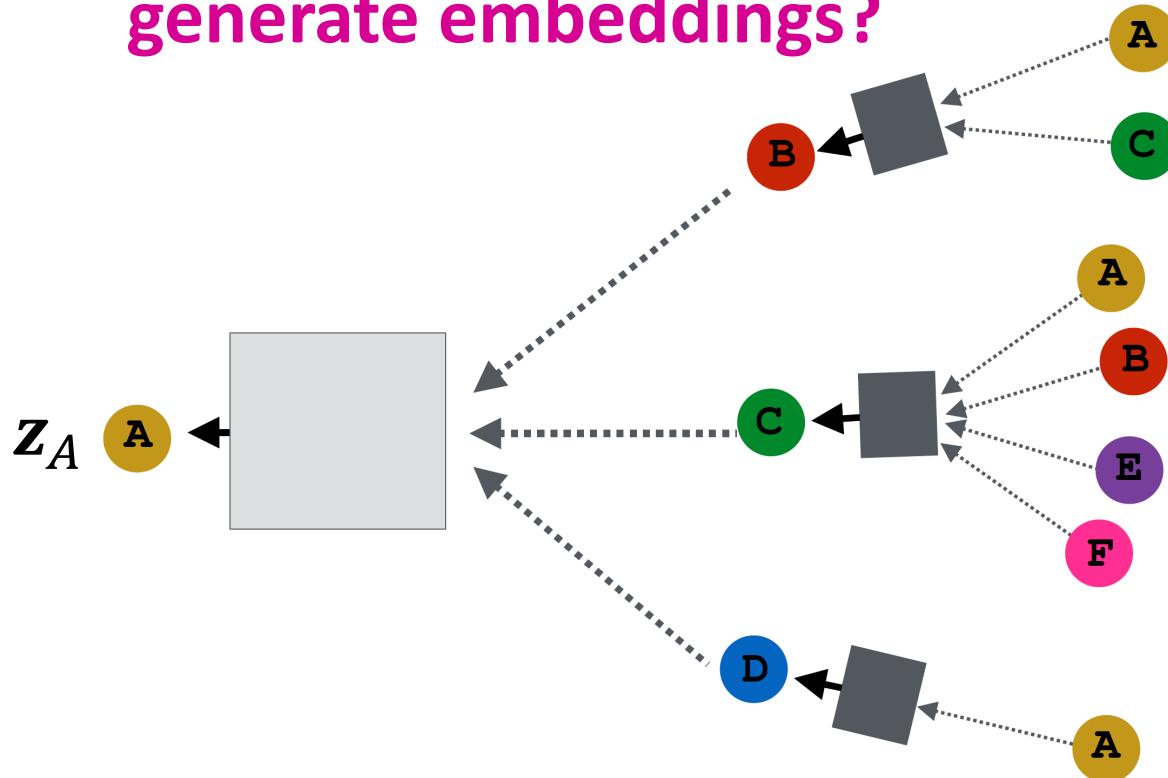
# The Math: Deep Encoder

- **Basic approach:** Average neighbor messages and apply a neural network



# Training the Model

How do we train the model to generate embeddings?



Need to define a loss function on the embeddings

# Model Parameters

$$\mathbf{h}_v^0 = \mathbf{x}_v$$

Trainable weight matrices  
(i.e., what we learn)

$$\mathbf{h}_v^k = \sigma \left( \mathbf{W}_k \sum_{u \in N(v)} \frac{\mathbf{h}_u^{k-1}}{|N(v)|} + \mathbf{B}_k \mathbf{h}_v^{k-1} \right), \quad \forall k \in \{1, \dots, K\}$$

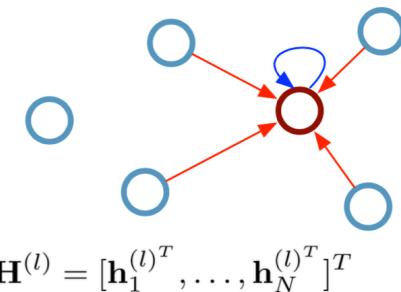
$$\mathbf{z}_v = \mathbf{h}_v^K$$

We can feed these **embeddings into any loss function** and run stochastic gradient descent to **train the weight parameters**

Equivalently  
rewritten in  
vector form:

$$\mathbf{H}^{(l+1)} = \sigma \left( \mathbf{H}^{(l)} \mathbf{W}_0^{(l)} + \tilde{\mathbf{A}} \mathbf{H}^{(l)} \mathbf{W}_1^{(l)} \right)$$

$$\text{with } \tilde{\mathbf{A}} = \mathbf{D}^{-\frac{1}{2}} \mathbf{A} \mathbf{D}^{-\frac{1}{2}}$$



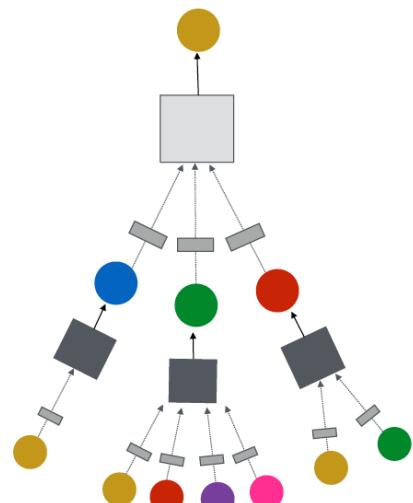
# Unsupervised Training

- Train in an **unsupervised** manner:
  - Use only the graph structure
  - “**Similar**” nodes have **similar embeddings**
- Unsupervised loss function can be anything from the last section, e.g., a loss based on
  - Random walks (node2vec, DeepWalk, struc2vec)
  - Graph factorization
  - Node proximity in the graph

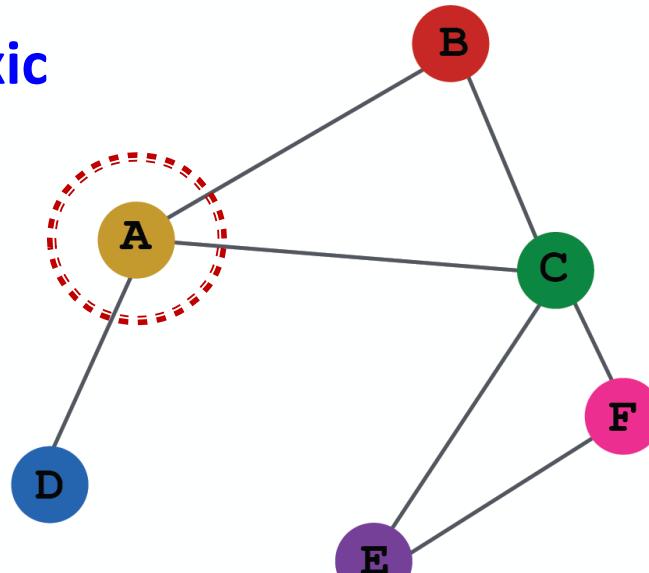
# Supervised Training

**Directly train** the model for a supervised task  
(e.g., node classification)

Safe or toxic  
drug?



Safe or toxic  
drug?



E.g., a drug-drug  
interaction network

# Supervised Training

Directly train the model for a supervised task  
(e.g., **node classification**)

$$\mathcal{L} = \sum_{v \in V} y_v \log(\sigma(\mathbf{z}_v^\top \boldsymbol{\theta})) + (1 - y_v) \log(1 - \sigma(\mathbf{z}_v^\top \boldsymbol{\theta}))$$

Encoder output: node embedding

Classification weights

Node class label

Safe or toxic drug?

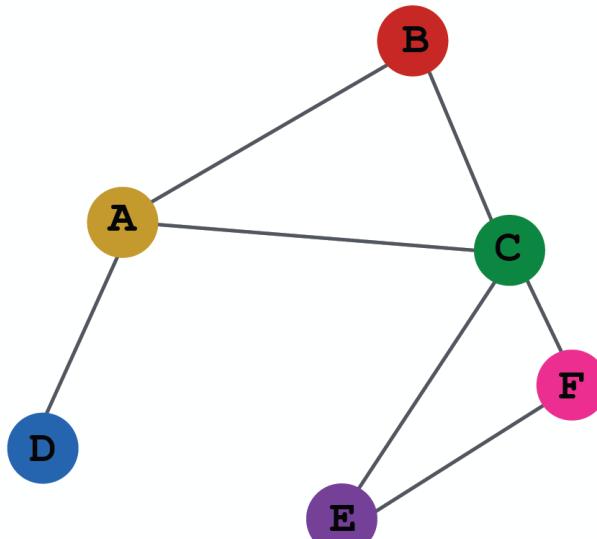
Encoder output: node embedding

Classification weights

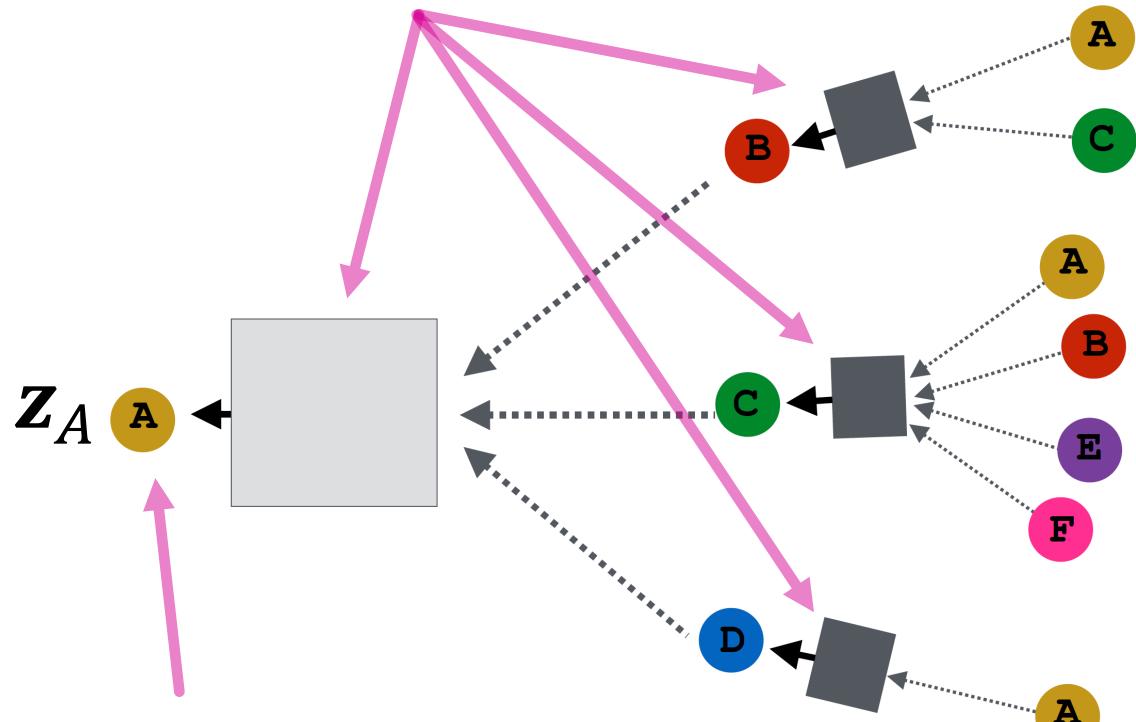
Node class label

Safe or toxic drug?

# Model Design: Overview

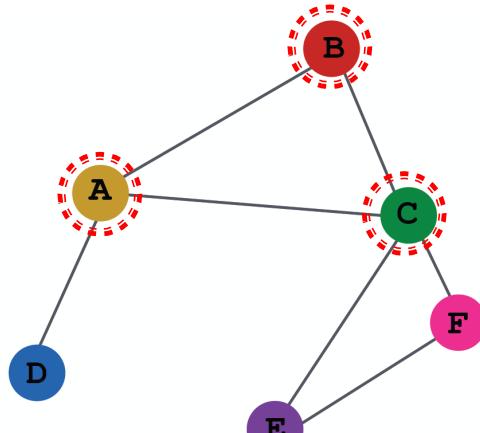


(1) Define a neighborhood aggregation function



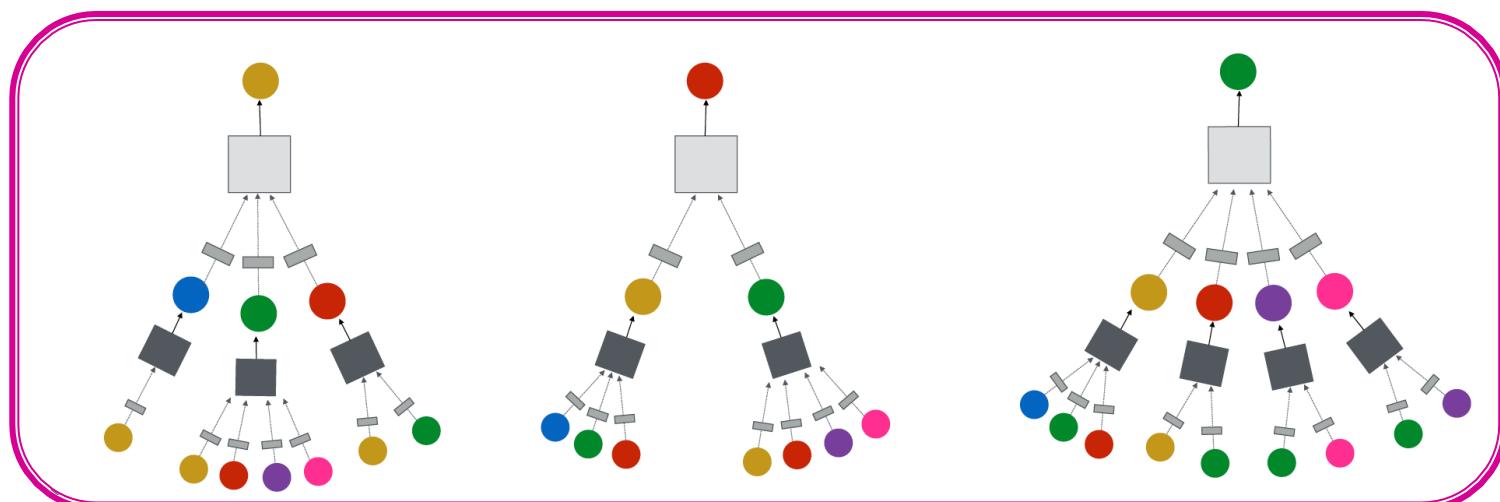
(2) Define a loss function on the embeddings

# Model Design: Overview

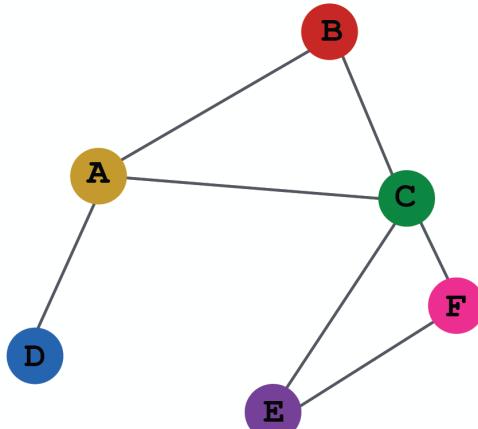


INPUT GRAPH

(3) Train on a set of nodes, i.e.,  
a batch of compute graphs



# Model Design: Overview

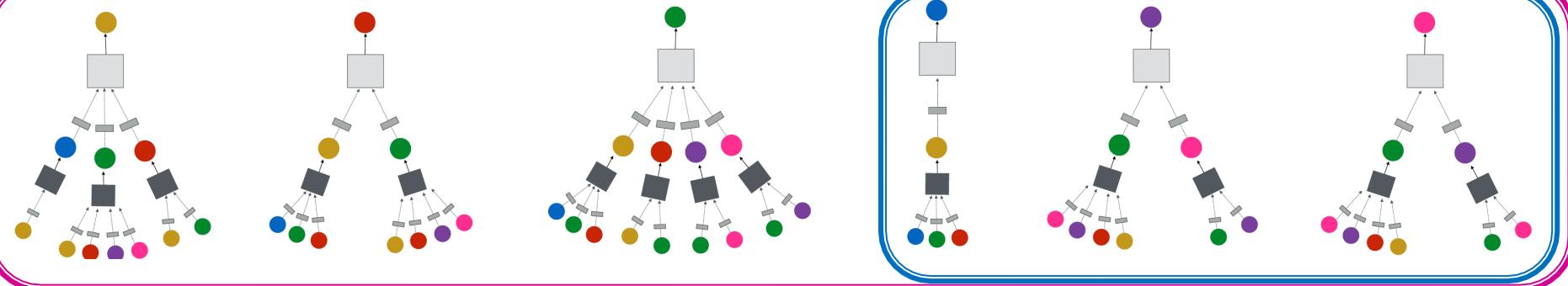


INPUT GRAPH

(4) Generate embeddings  
for nodes as needed

Even for nodes we never  
trained on!

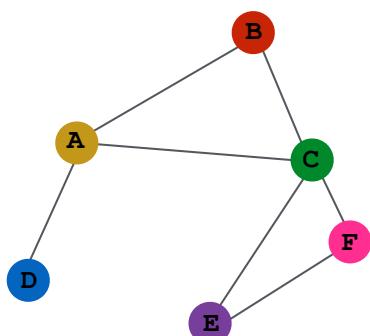
Sharing Parameters



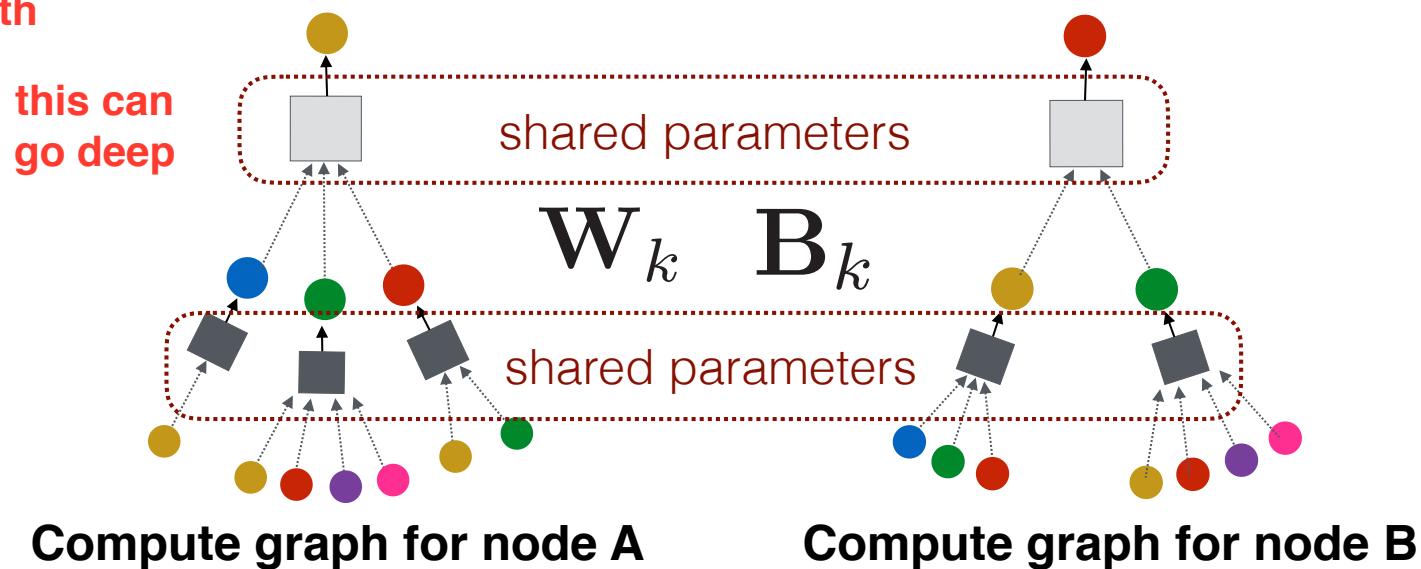
# Inductive Capability

- The same aggregation parameters are shared for all nodes:
  - The number of model parameters is sublinear in  $|V|$  and we can **generalize to unseen nodes!**

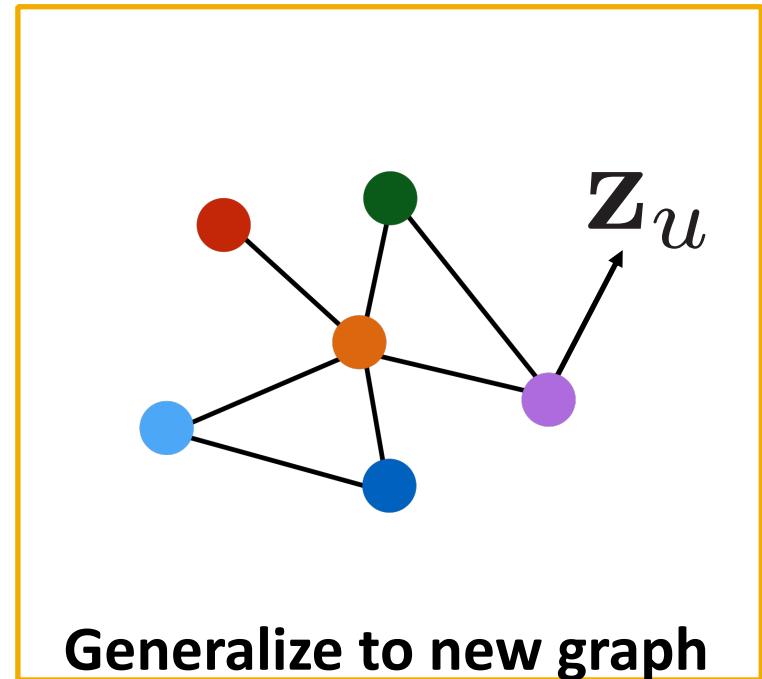
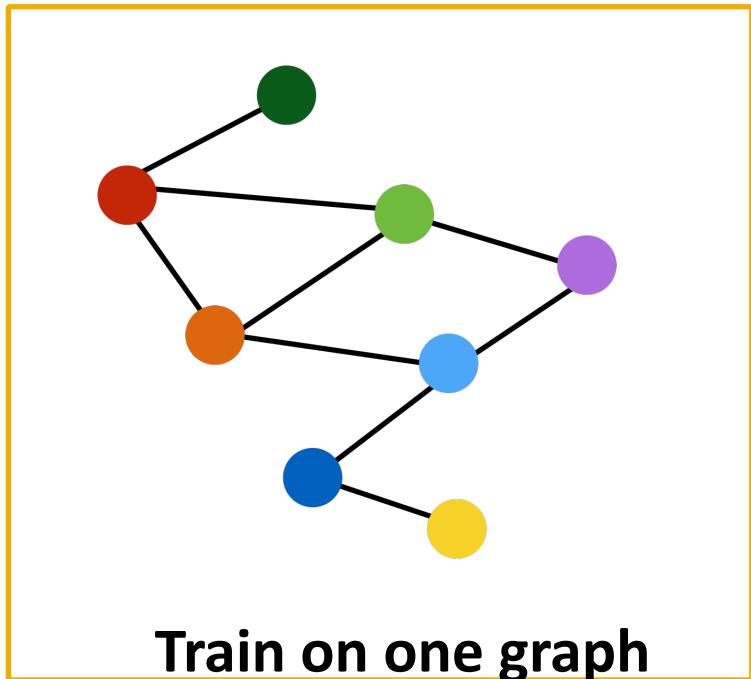
two notions of depth



INPUT GRAPH



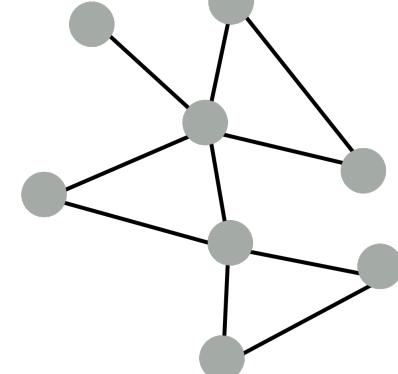
# Inductive Capability: New Graphs



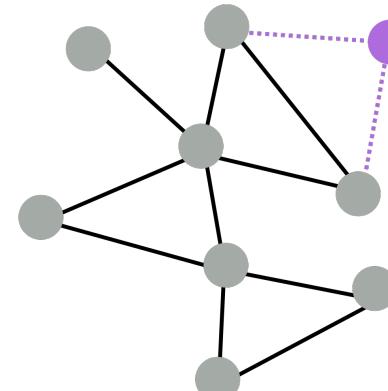
Inductive node embedding → Generalize to entirely unseen graphs

E.g., train on protein interaction graph from model organism A and generate embeddings on newly collected data about organism B

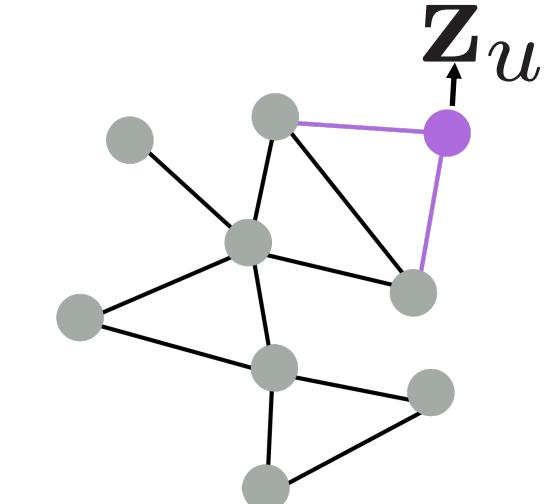
# Inductive Capability: New Nodes



Train with snapshot



New node arrives



Generate embedding  
for new node

- Many application settings constantly encounter previously unseen nodes:
  - E.g., Reddit, YouTube, Google Scholar
- Need to generate new embeddings “on the fly”

# Summary So Far

- **Recap:** Generate node embeddings by aggregating neighborhood information
  - We saw a **basic variant of this idea**
  - Key distinctions are in how different approaches aggregate information across the layers
- **Next:** Describe GraphSAGE graph neural network architecture

# Outline of Today's Lecture

1. Basics of deep learning for graphs 
2. Graph Convolutional Networks
3. Graph Attention Networks (GAT)
4. Practical tips and demos

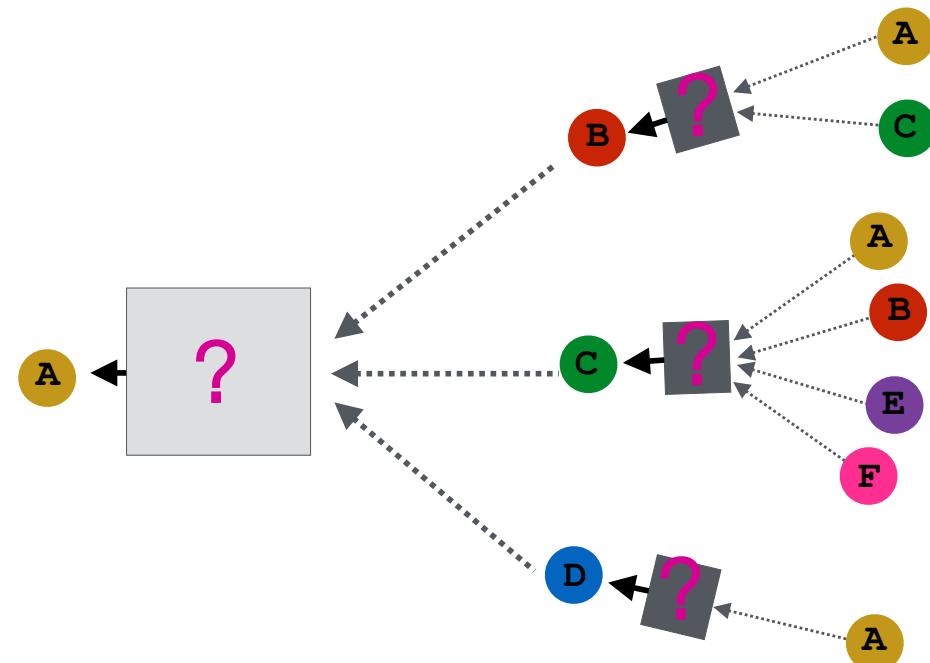
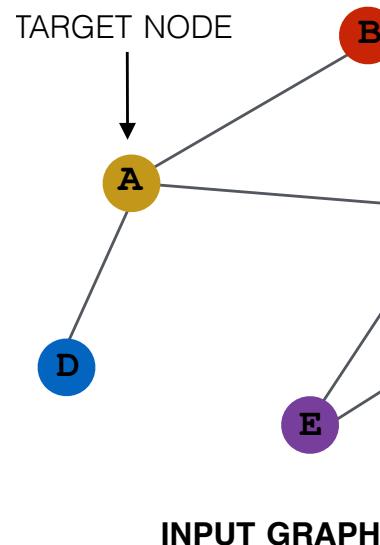


# **Graph Convolutional Networks and GraphSAGE**

# GraphSAGE Idea

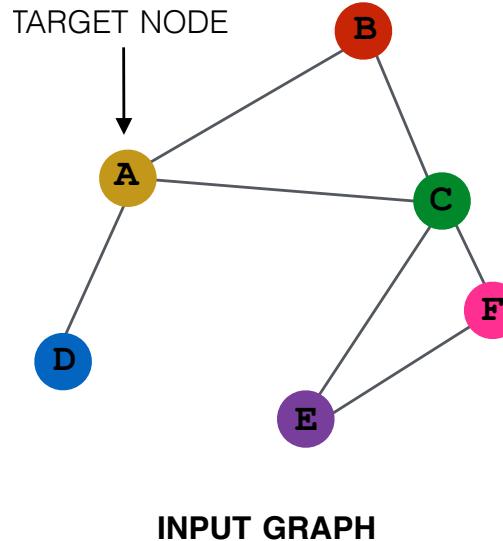
So far we have aggregated the neighbor messages by taking their (weighted) average

**Can we do better?**



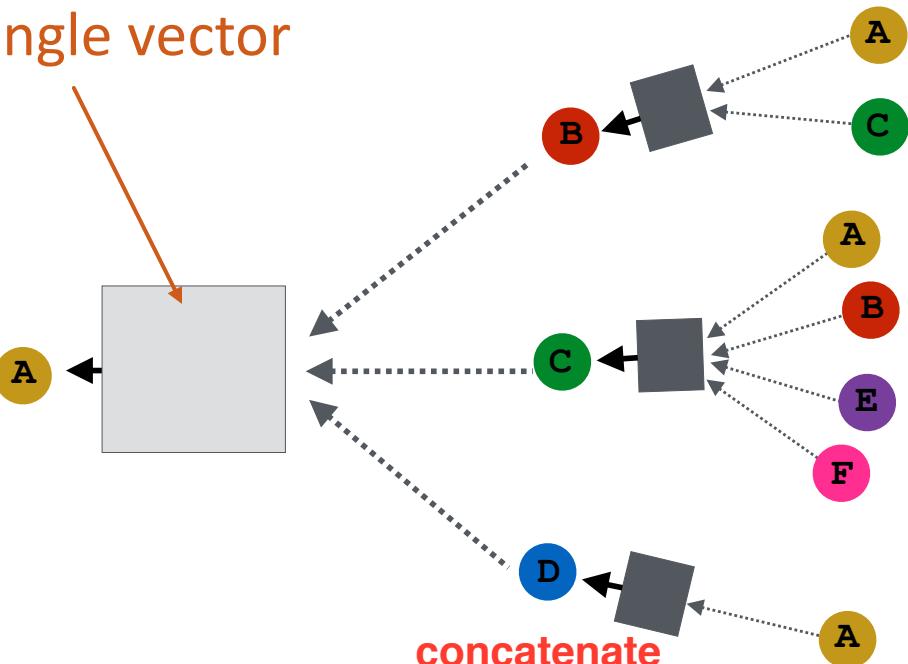
# GraphSAGE Idea

Any differentiable function that  
maps set of vectors in  $N(u)$  to  
a single vector



$$\mathbf{h}_v^k = \sigma \left( [\mathbf{A}_k \cdot \text{AGG}(\{\mathbf{h}_u^{k-1}, \forall u \in N(v)\}), \mathbf{B}_k \mathbf{h}_v^{k-1}] \right)$$

Apply L2 normalization for each node embedding at every layer



# Neighborhood Aggregation

- Simple neighborhood aggregation:

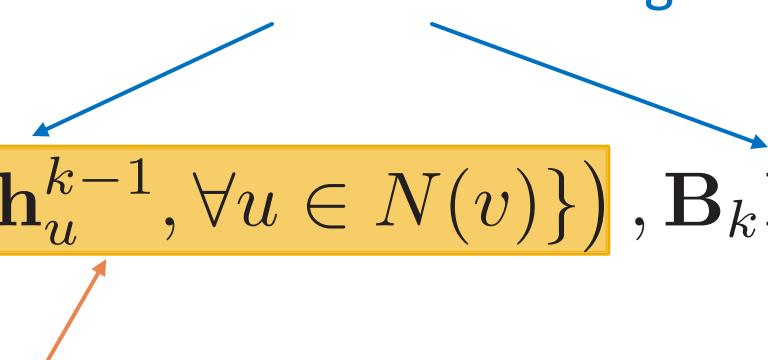
$$\mathbf{h}_v^k = \sigma \left( \mathbf{W}_k \sum_{u \in N(v)} \frac{\mathbf{h}_u^{k-1}}{|N(v)|} + \mathbf{B}_k \mathbf{h}_v^{k-1} \right)$$

- GraphSAGE:

Concatenate neighbor embedding  
and self embedding

$$\mathbf{h}_v^k = \sigma \left( [\mathbf{W}_k \cdot \text{AGG} \left( \{\mathbf{h}_u^{k-1}, \forall u \in N(v)\} \right), \mathbf{B}_k \mathbf{h}_v^{k-1}] \right)$$

Generalized aggregation



# Neighbor Aggregation: Variants

- **Mean:** Take a weighted average of neighbors

$$\text{AGG} = \sum_{u \in N(v)} \frac{\mathbf{h}_u^{k-1}}{|N(v)|}$$

- **Pool:** Transform neighbor vectors and apply symmetric vector function

$$\text{AGG} = \gamma \left( \{\mathbf{Q}\mathbf{h}_u^{k-1}, \forall u \in N(v)\} \right)$$

Element-wise mean/max

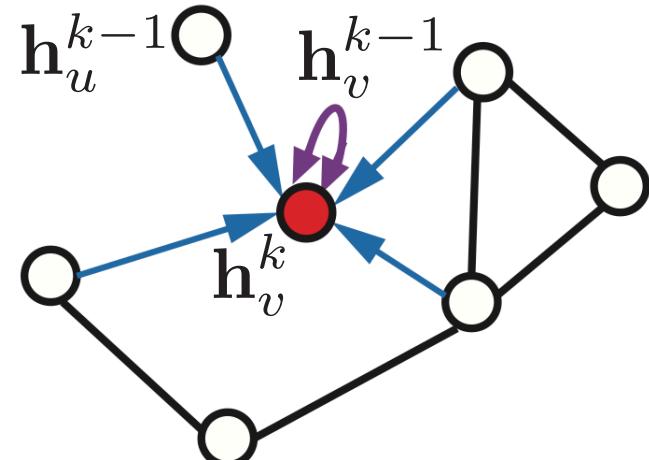
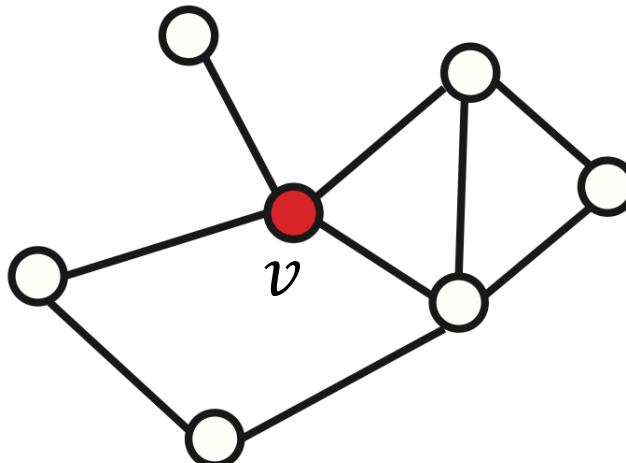
- **LSTM:** Apply LSTM to reshuffled of neighbors

$$\text{AGG} = \text{LSTM} \left( [\mathbf{h}_u^{k-1}, \forall u \in \pi(N(v))] \right)$$

# Recap: GCN, GraphSAGE

**Key idea:** Generate node embeddings based on **local neighborhoods**

- Nodes aggregate “messages” from their neighbors using neural networks
- **Graph convolutional networks:**
  - **Basic variant:** Average neighborhood information and stack neural networks
- **GraphSAGE:**
  - Generalized neighborhood aggregation



# Efficient Implementation

- Many aggregations can be performed efficiently by (sparse) matrix operations
- Let  $H^{k-1} = [\mathbf{h}_1^{k-1} \dots \mathbf{h}_n^{k-1}]$

$$\sum_{u \in N(v)} \frac{\mathbf{h}_u^{k-1}}{|N(v)|} \quad \longrightarrow \quad H^k = D^{-1} A H^{k-1}$$

- Another example: GCN (Kipf *et al.* 2017)

$$H^k = D^{-1/2} A D^{1/2} H^{k-1}$$

# More on Graph Neural Networks

## Tutorials and overviews:

- Relational inductive biases and graph networks (Battaglia et al., 2018)
- Representation learning on graphs: Methods and applications (Hamilton et al., 2017)

## Attention-based neighborhood aggregation:

- Graph attention networks (Hoshen, 2017; Velickovic et al., 2018; Liu et al., 2018)

## Embedding entire graphs:

- Graph neural nets with edge embeddings (Battaglia et al., 2016; Gilmer et. al., 2017)
- Embedding entire graphs (Duvenaud et al., 2015; Dai et al., 2016; Li et al., 2018) and graph pooling (Ying et al., 2018, Zhang et al., 2018)
- Graph generation and relational inference (You et al., 2018; Kipf et al., 2018)
- How powerful are graph neural networks(Xu et al., 2017)

## Embedding nodes:

- Varying neighborhood: Jumping knowledge networks (Xu et al., 2018), GeniePath (Liu et al., 2018)
- Position-aware GNN (You et al. 2019)

## Spectral approaches to graph neural networks:

- Spectral graph CNN & ChebNet (Bruna et al., 2015; Defferrard et al., 2016)
- Geometric deep learning (Bronstein et al., 2017; Monti et al., 2017)

## Other GNN techniques:

- Pre-training Graph Neural Networks (Hu et al., 2019)
- GNNExplainer: Generating Explanations for Graph Neural Networks (Ying et al., 2019)

# Outline of Today's Lecture

- 1. Basics of deep learning for graphs** 
- 2. Graph Convolutional Networks** 
- 3. Graph Attention Networks (GAT)** 
- 4. Practical tips and demos**

# Graph Attention Networks

# Simple Neighborhood Aggregation

- **Recap: Simple neighborhood aggregation:**

$$\mathbf{h}_v^k = \sigma \left( \mathbf{W}_k \sum_{u \in N(v)} \frac{\mathbf{h}_u^{k-1}}{|N(v)|} + \mathbf{B}_k \mathbf{h}_v^{k-1} \right)$$

- Graph convolutional operator:
  - Aggregates messages across neighborhoods,  $N(v)$
  - $\alpha_{vu} = 1/|N(v)|$  is the **weighting factor (importance)** of node  $u$ 's message to node  $v$
  - $\Rightarrow \alpha_{vu}$  is defined **explicitly** based on the structural properties of the graph
  - $\Rightarrow$  All neighbors  $u \in N(v)$  are equally important to node  $v$

# Graph Attention Networks

Can we do better than simple neighborhood aggregation?

Can we let weighting factors  $\alpha_{vu}$  to be implicitly defined?

- **Goal:** Specify arbitrary importances to different neighbors of each node in the graph
- **Idea:** Compute embedding  $h_v^k$  of each node in the graph following an **attention strategy**:
  - Nodes attend over their neighborhoods' message
  - Implicitly specifying different weights to different nodes in a neighborhood

# Attention Mechanism (1)

- Let  $\alpha_{vu}$  be computed as a byproduct of an **attention mechanism**  $a$ :
  - Let  $a$  compute **attention coefficients**  $e_{vu}$  across pairs of nodes  $u, v$  based on their messages:
$$e_{vu} = a(\mathbf{W}_k \mathbf{h}_u^{k-1}, \mathbf{W}_k \mathbf{h}_v^{k-1})$$
    - $e_{vu}$  indicates the importance of node  $u$ 's message to node  $v$
  - Normalize coefficients** using the softmax function in order to be comparable across different neighborhoods:

$$\alpha_{vu} = \frac{\exp(e_{vu})}{\sum_{k \in N(v)} \exp(e_{vk})}$$

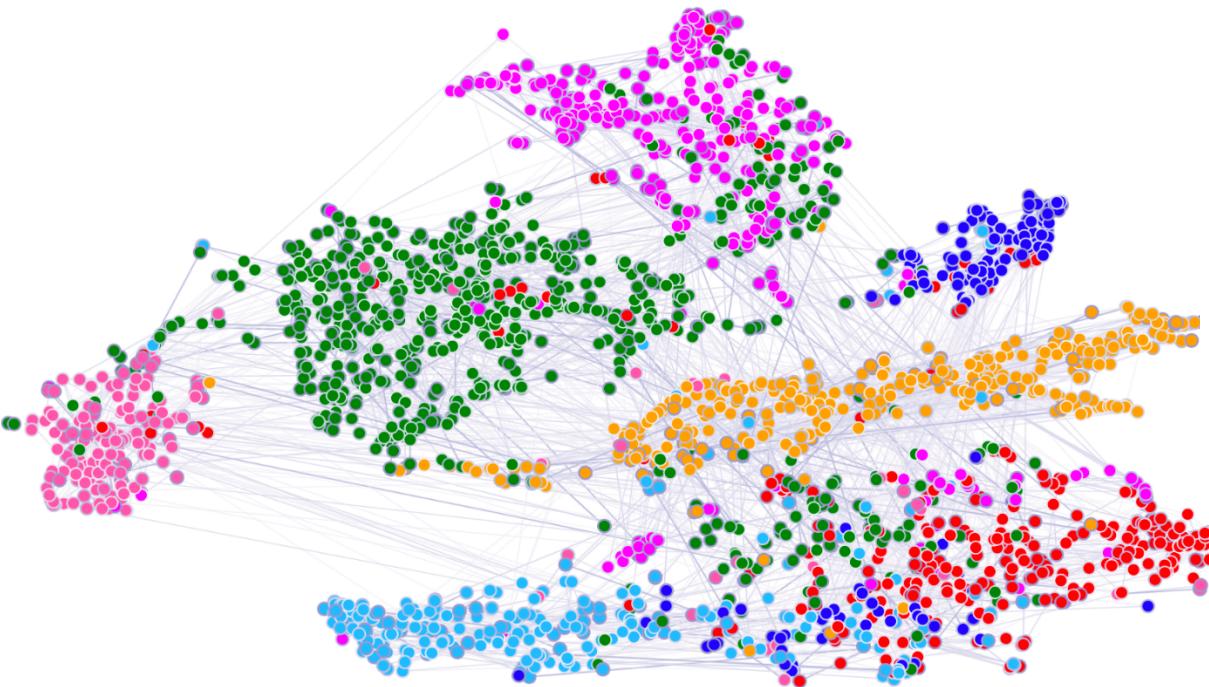
$$\mathbf{h}_v^k = \sigma\left(\sum_{u \in N(v)} \alpha_{vu} \mathbf{W}_k \mathbf{h}_u^{k-1}\right)$$

**Next:** What is the form of attention mechanism  $a$ ?

# Attention Mechanism (2)

- Attention mechanism  $\alpha$ :
  - The approach is agnostic to the choice of  $\alpha$ 
    - E.g., use a simple single-layer neural network
    - $\alpha$  can have parameters, which need to be estimates
  - Parameters of  $\alpha$  are trained jointly:
    - Learn the parameters together with weight matrices (i.e., other parameter of the neural net) in an end-to-end fashion
- **Multi-head attention:** Stabilize the learning process of attention mechanism [Velickovic et al., ICLR 2018]:
  - Attention operations in a given layer are independently replicated R times (each replica with different parameters)
  - Outputs are aggregated (by concatenating or adding)

# GAT Example: Cora Citation Net



Method	Cora
MLP	55.1%
ManiReg (Belkin et al., 2006)	59.5%
SemiEmb (Weston et al., 2012)	59.0%
LP (Zhu et al., 2003)	68.0%
DeepWalk (Perozzi et al., 2014)	67.2%
ICA (Lu & Getoor, 2003)	75.1%
Planetoid (Yang et al., 2016)	75.7%
Chebyshev (Defferrard et al., 2016)	81.2%
GCN (Kipf & Welling, 2017)	81.5%
<b>GAT</b>	<b>83.3%</b>
improvement w.r.t GCN	1.8%

Attention mechanism can be used with many different graph neural network models

In many cases, attention leads to performance gains

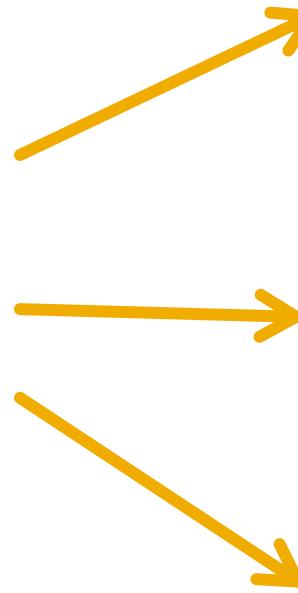
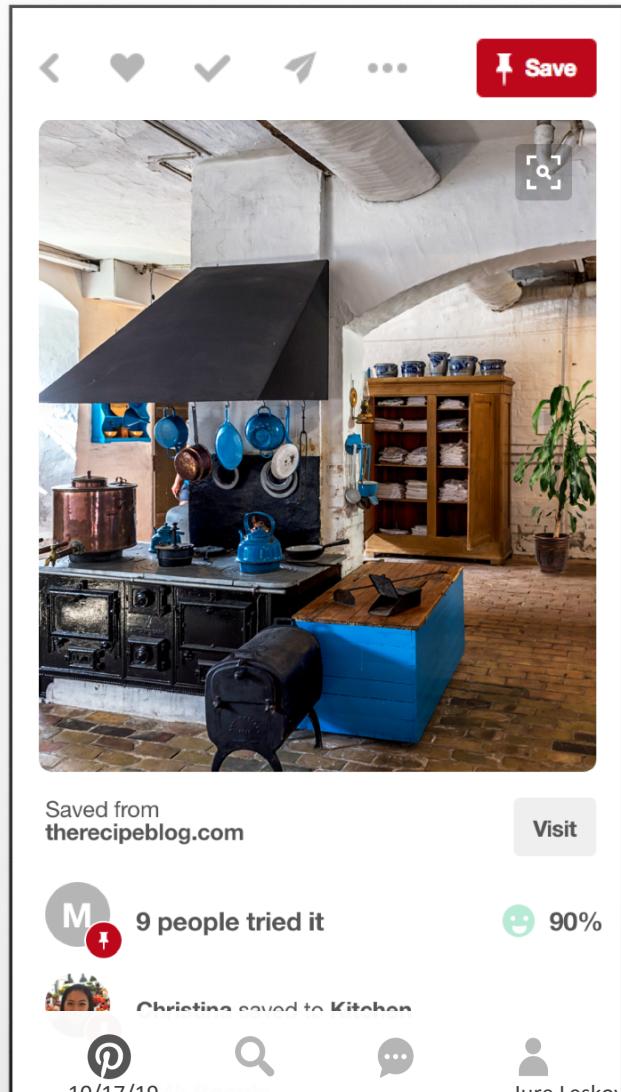
- t-SNE plot of GAT-based node embeddings:
  - Node color: 7 publication classes
  - Edge thickness: Normalized attention coefficients between nodes  $i$  and  $j$ , across eight attention heads,  $\sum_k (\alpha_{ij}^k + \alpha_{ji}^k)$

# Outline of Today's Lecture

1. Basics of deep learning for graphs 
2. Graph Convolutional Networks (GCN) 
3. Graph Attention Networks (GAT) 
4. Practical tips and demos 

# Example Application

# Application: Pinterest



Blue accents  
219 Pins



Vintage kitchen  
377 Pins



- 300M users
- 4+B pins, 2+B boards

# Application: Pinterest

Human curated collection of pins

The image shows a mobile screen of the Pinterest app. At the top, there are three pin cards:

- A person wearing a dark blue jacket with 'VERY APE' and an ape logo on the back.
- A Hans Wegner chair next to a lamp.
- A green plant with the caption "This is just a beautiful image for thoughts. Yay or nay, your choice."

Below these are two explanatory text blocks:

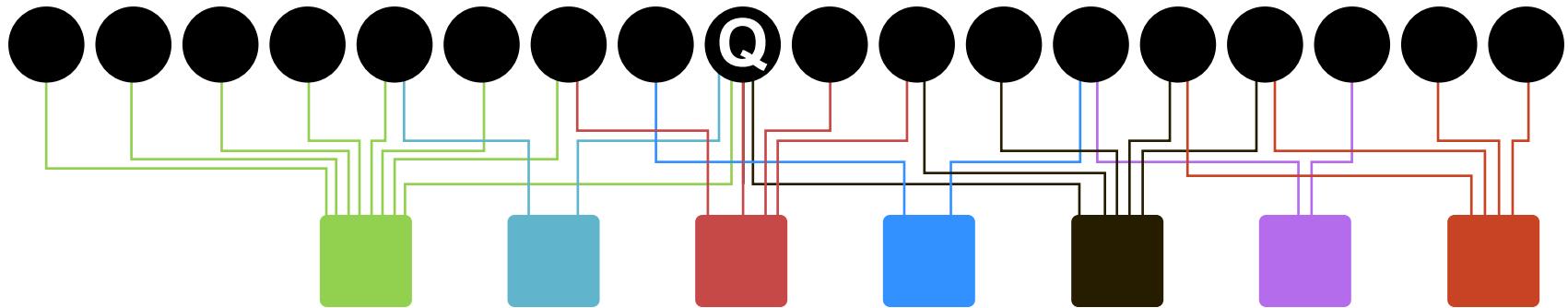
- Pin:** A visual bookmark someone has saved from the internet to a board they've created.
- Pin:** Image, text, link

At the bottom, there are several boards displayed as grid thumbnails:

- mid century modern ... MJL I -
- Man Style Gavin Jones
- men + style I FIG+SALT
- Plants HelloSandwich
- Men's Style Andrea Sempli
- Mid century modern Tyler Goodro
- Plants Moorea Seal
- Mid century modern ... Prettygreenetea

**Board:** A collection of ideas (pins having something in common)

# Pinterest Graph



**Graph:** 2B pins, 1B boards, 20B edges

- **Graph is dynamic:** Need to apply to new nodes without model retraining
- **Rich node features:** Content, images

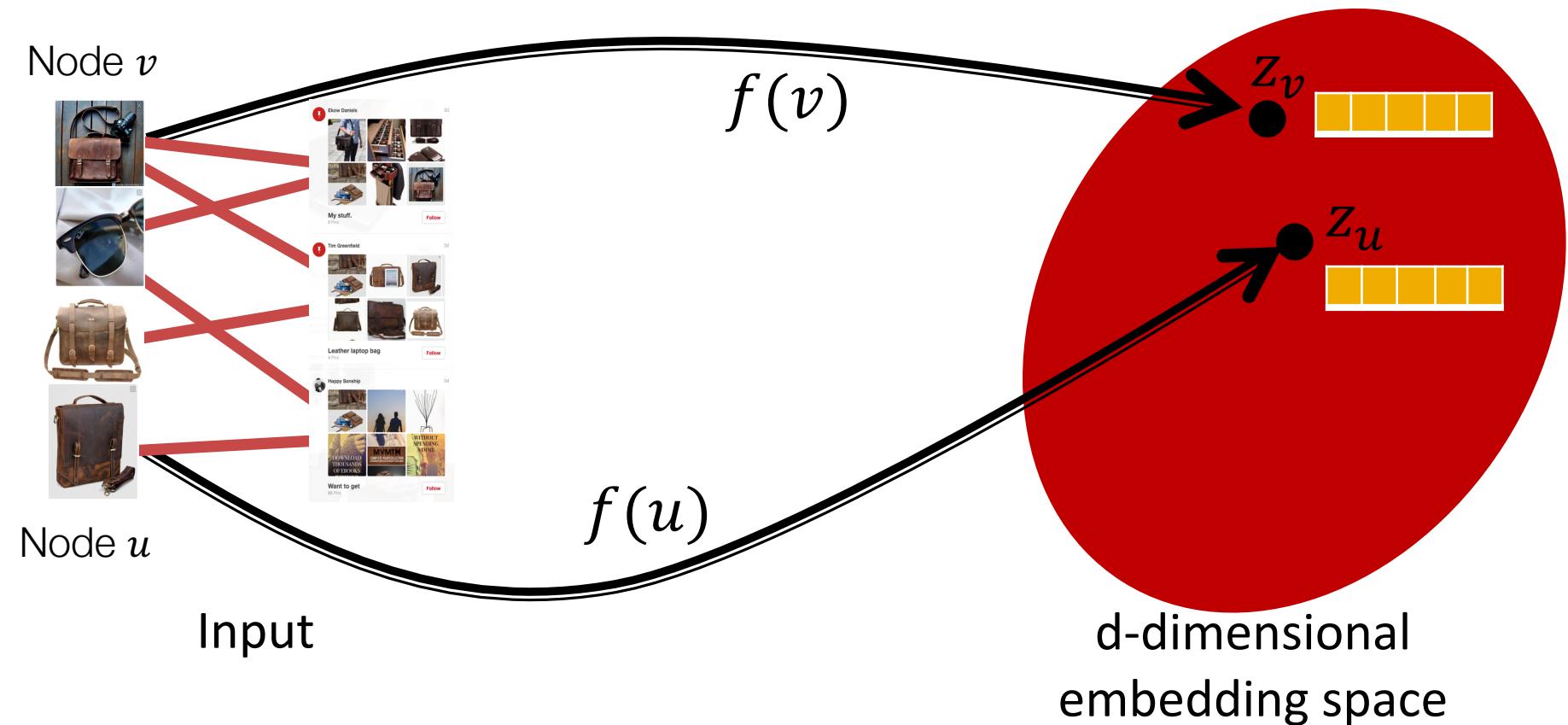
# PinSage: Overview

- **PinSage** graph convolutional network:
  - **Goal:** Generate embeddings for nodes (e.g., Pins/images) in a web-scale Pinterest graph containing billions of objects
  - **Key Idea:** Borrow information from nearby nodes
    - E.g., bed rail Pin might look like a garden fence, but gates and beds are rarely adjacent in the graph



- Pin embeddings are essential to various tasks like recommendation of Pins, classification, clustering, ranking
  - Services like “Related Pins”, “Search”, “Shopping”, “Ads”

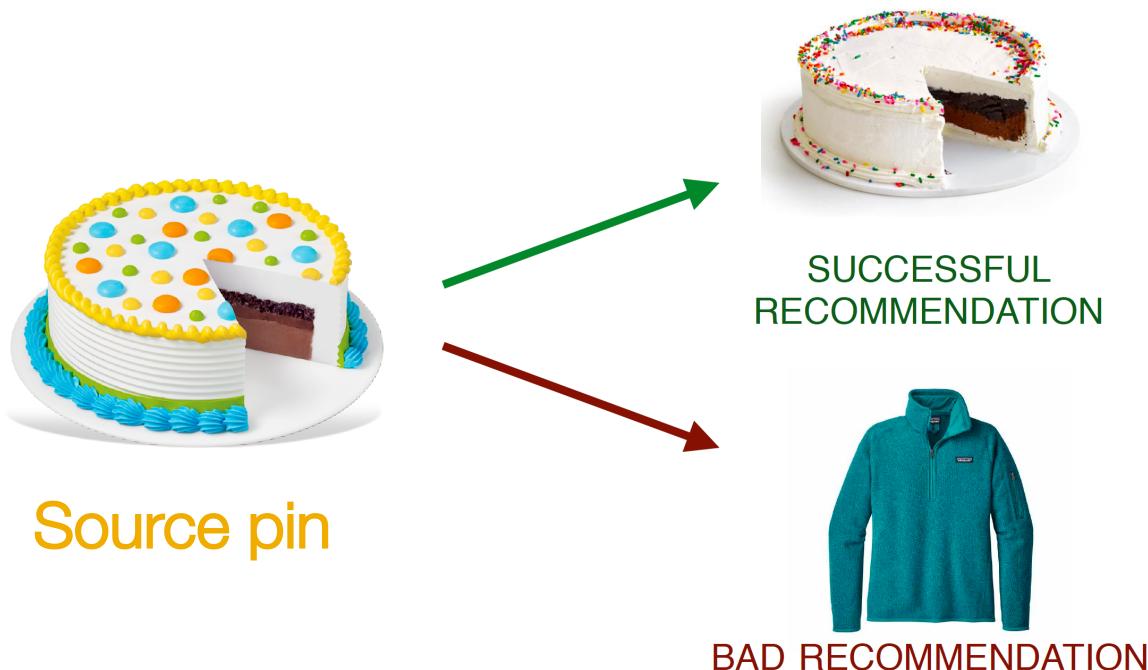
# Embedding Nodes



**Goal:** Map nodes to  $d$ -dimensional embeddings such that **nodes that are related are embedded close together**

# Task Overview

## Task: Recommend related pins to users



**Task:** Learn node embeddings  $z_i$  such that

$$d(z_{\text{cake}1}, z_{\text{cake}2}) < d(z_{\text{cake}1}, z_{\text{sweater}})$$

### Challenges:

- **Massive size:** 3 billion nodes, 20 billion edges
- **Heterogeneous data:** Rich image and text features

# PinSAGE Training

**Goal:** Identify target pin among 3B pins

- **Issue:** Need to learn with resolution of 100 vs. 3B
- **Idea:** Use harder and harder negative samples
- Include more and more hard negative samples for each epoch



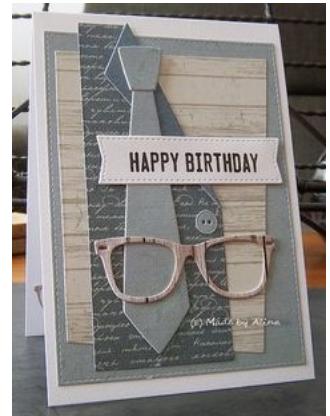
Source pin



Positive



Easy negative



Hard negative

# PinSAGE Efficiency

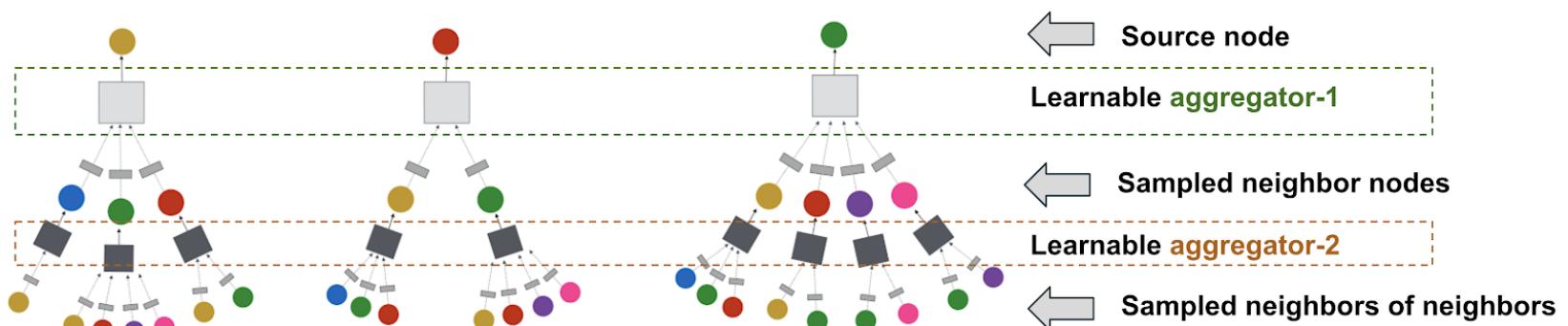
- How to scale the training as well as inference of node embeddings to **graphs with billions of nodes and tens of billions of edges?**
  - 10,000X larger dataset than any previous graph neural network application
- **Key innovations:**
  - Sub-sample neighborhoods for efficient GPU batching
  - Producer-consumer CPU-GPU training pipeline
  - Curriculum learning for negative samples
  - MapReduce for efficient inference

# PinSage: Key Innovations (1)

## ■ Three key innovations:

### 1. On-the-fly graph convolutions

- Sample the neighborhood around a node and dynamically construct **a computation graph**
- Perform a **localized graph convolution** around a particular node
- Does not need the entire graph during training



# PinSage: Key Innovations (2)

- **Three key innovations:**
  1. **On-the-fly graph convolutions**
  2. **Constructing convolutions via random walks**
    - Performing convolutions on full neighborhoods is infeasible:
      - How to select the set of neighbors of a node to convolve over?
    - **Importance pooling:** Define importance-based neighborhoods by simulating random walks and selecting the neighbors with the highest visit counts
  3. **Efficient MapReduce inference**
    - Bottom-up aggregation of node embeddings lends itself to MapReduce
      - Decompose each aggregation step across all nodes into three operations in MapReduce, i.e., *map*, *join*, and *reduce*

# PinSage: Experiments

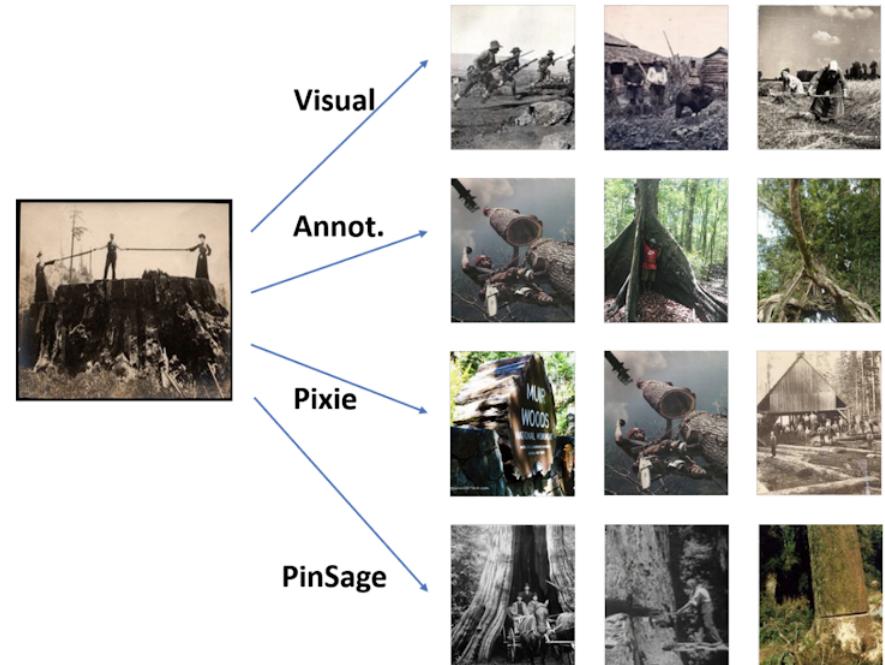
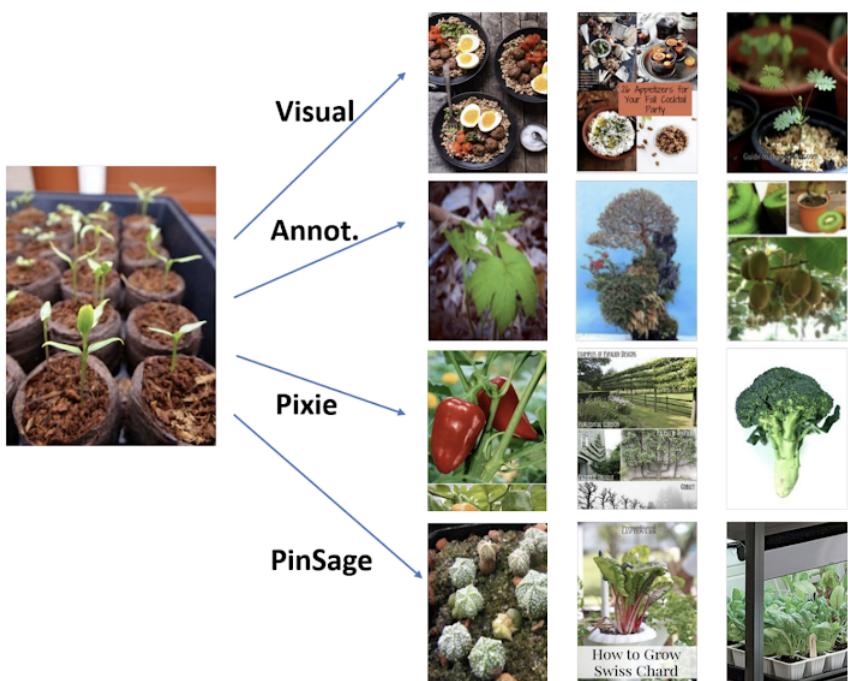
## ■ Baselines:

- **Visual:** Nearest neighbors of CNN visual embeddings for recommendations
- **Annotation:** Nearest neighbors in terms of Word2vec embeddings
- **Combined:** Concatenate embeddings:
  - Uses exact same data and loss function as PinSage

Method	Hit-rate	MRR
Visual	17%	0.23
Annotation	14%	0.19
Combined	27%	0.37
max-pooling	39%	0.37
mean-pooling	41%	0.51
mean-pooling-xent	29%	0.35
mean-pooling-hard	46%	0.56
PinSage	67%	<b>0.59</b>

PinSage gives 150% improvement in hit rate and 60% improvement in MRR over the best baseline

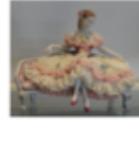
# Example Pin Recommendations



Pixie is a purely graph-based method that uses biased random walks to generate ranking scores by simulating random walks starting at query Pin. Items with top scores are retrieved as recommendations [Eksombatchai et al., 2018]

# PinSAGE Recommendations

Query



PinSAGE



# PinSAGE Recommendations

Query



A Shih Tzu recipe  
Yield: 4 pieces  
Ingredients:  
- a dash of lime, several teaspoons of rabbit, a couple of ounces of dragonfly oil, one part sweet butter, a dash of lavender, a piece of old man, a lot of hogger, a tablespoon of monkey, one part baby food, and a dash of pony hair.

PinSAGE



# General Tips and Practical Demos

# General Tips

- Data preprocessing is important:
  - Use renormalization tricks
  - Variance-scaled initialization
  - Network data whitening
- ADAM optimizer:
  - ADAM naturally takes care of decaying the learning rate
- ReLU activation function often works really well
- No activation function at your output layer:
  - Easy mistake if you build layers with a shared function
- Include bias term in every layer
- GCN layer of size 64 or 128 is already plenty

# Debugging Deep Networks

- **Debug?!**:
  - Loss/accuracy not converging during training
- **Important for model development:**
  - **Overfit on training data:**
    - Accuracy should be essentially 100% or error close to 0
    - If neural network cannot overfit a single data point, something is wrong
  - **Scrutinize your loss function!**
  - **Scrutinize your visualizations!**

# Demo: Human Disease Network

Human Disease Network

snap.stanford.edu/deepnetbio-ismb/ipynb/Human+Disease+Network.html

Marinka

## Embedding the Human Disease Network

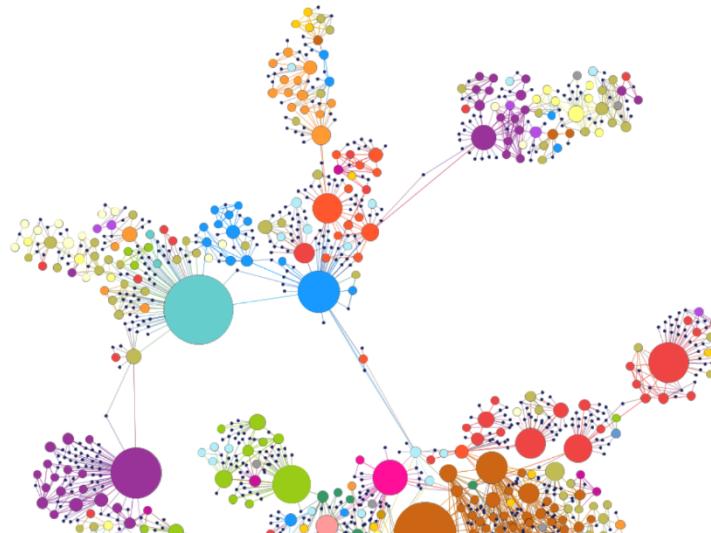
(This demo is a part of [Deep Learning for Network Biology](#) tutorial.)

Human disease network is a network, in which nodes represent diseases and two diseases are connected to each other if they share at least one gene in which mutations are associated with both diseases.

The network is described in Goh et al., [The Human Disease Network](#), PNAS 2007.

The figure below shows the human disease network.

Although the layout of the network was generated independently of any knowledge of disease classes, the resulting network is naturally and visibly clustered according to major disease classes (e.g., bone, cancer, cardiovascular, skeletal, or metabolic diseases; each disease class is represented by a different color). The size of a node is proportional to the number of genes participating in the corresponding disease.



# Demo: Protein Interaction Prediction

Graph Convolutional Prediction x Marinka

snap.stanford.edu/deepnetbio-ismb/ipynb/Graph+Convolutional+Prediction+of+Protein+Interactions+in+Yeast.html

## Graph Convolutional Prediction of Protein Interactions in Yeast

(This demo is a part of [Deep Learning for Network Biology](#) tutorial.)

In this example, we demonstrate the utility of deep learning methods for an important prediction problem on biological graphs. In particular, we consider the problem of predicting [protein-protein interactions](#) (PPIs).

Protein-protein interactions (PPIs) are essential to almost every process in a cell. Understanding PPIs is crucial for understanding cell physiology in normal and disease states. Furthermore, knowledge of PPIs can be used:

- for drug development, since drugs can affect PPIs,
- to assign roles (i.e., protein functions) to uncharacterized proteins,
- to characterize the relationships between proteins that form multi-molecular complexes, such as the proteasome.

We represent the totality of PPIs that happen in a cell, an organism or a specific biological context with a [protein-protein interaction network](#). These networks are mathematical representations of all physical contacts between proteins in the cell.

The development of large-scale PPI screening techniques, especially [high-throughput affinity purification combined with mass-spectrometry](#) and the [yeast two-hybrid assay](#), has caused an explosion in the amount of PPI data and the construction of ever more complex and complete interaction networks. For example, the figure below is a graphical representation of three different types of protein-protein interaction networks in [yeast S. cerevisiae](#). The structure of the binary interaction network is obviously different from the structure of the co-complex interaction network. The network structure of the literature-curated dataset resembles that of the co-complex dataset, even though the literature-curated datasets are reported to contain mostly binary interactions.

However, current knowledge of protein-protein interaction networks is both [incomplete and noisy](#), as PPI screening techniques are limited in how many true interactions they can detect. Furthermore, PPI screening techniques often have high false positive and negative rates. These limitations present a great opportunity for computational methods to predict protein-protein interactions.

The figure displays three network graphs side-by-side. The first graph, labeled 'Binary (Y2H-union)', shows a dense, roughly spherical cluster of red nodes connected by a web of red lines. Below it is a small schematic showing a grid of nodes with dashed lines connecting them. The second graph, labeled 'Co-complex (Combined-AP/MS)', shows a more sparse, branched structure with green nodes and lines, featuring several distinct clusters and some long-range connections. Below it is a small schematic showing a grid of nodes with solid lines connecting them. The third graph, labeled 'Literature (LC-multiple)', shows a complex, highly branched structure with blue nodes and lines, forming a dense web-like pattern. Below it is a small schematic showing a grid of nodes with various line patterns (solid, dashed, dotted) connecting them.

# Outline of Today's Lecture

1. Basics of deep learning for graphs 
2. Graph Convolutional Networks 
3. Graph Attention Networks (GAT) 
4. Practical tips and demos 