

Causal Representation Learning for Medical Image Analysis

Changjie Lu

Summary of the Proposal

Statistical machine learning algorithms have achieved state-of-the-art results on benchmark datasets, outperforming humans in many tasks. However, the out-of-distribution data and confounder, which have an unpredictable causal relationship, significantly degrade the performance of the existing models. Causal Representation Learning (CRL) has recently been a promising direction to address the causal relationship problem in vision understanding. This research proposal summarized recent advances in CRL in vision and analyzed the future research objectives. Firstly, we introduced the basic concept of causal inference. Secondly, we analyzed the CRL theoretical work, especially in invariant risk minimization, and the practical work in feature understanding and transfer learning. Finally, we proposed future research objectives in medical image analysis and expected outcomes.

Background

Correlation does not imply causation [21]. One famous example is the Simpson’s paradox[86] (see Fig. 1). Even if the series of events do have causality, it is hard to distinguish that relationship. One effective way of learning causality is to conduct a randomized controlled trial (RCT), randomly assigning participants into a treatment group or a control group so that people can observe the effect via the outcome variable. However, RCT is inflexible because it targets the sample average, which makes the mechanism unclear. Another widely used information type is observational data, which records every event that could be observed. Nowadays, machine learning algorithms attempt to learn patterns by fitting the observational data, losing sight of the causality. This results in poor performance when generalizing the model to an unseen distribution or learning the wrong causality.

To escape the dilemma mentioned above, Pearl firstly introduced a causality system with the three-layer causal hierarchy, called Pearl Causal Hierarchy (PCH), which contains Association, Intervention, and Counterfactuals[63][76][64]. To support this theory, Pearl developed structural causal model (SCM), which combines structural equation models (SEM), potential outcome framework, and the directed acyclic graphs (DAG)[79][74][61][78][65] for probabilistic reasoning and causal analysis, typically using the do-calculus[62]. With these tools, causal analysis infers probabilities under not only statistical conditions but also the dynamics of probabilities under changing conditions [64]. Currently, causal inference is a popular research direction with comprehensive literature [99][35][20][5][107], and is widely applied in decision evaluation (e.g. healthcare), counterfactual estimation (e.g. representation learning methods), and dealing with selection bias (e.g. advertising, recommendation) [99].

Although most of the causal inference could be applied in low-dimensional data (e.g., tabular data, describable events), research in high-dimensional data is still a struggle. In computer vision, for example, which often suffers from confounder that the model tends to classify a cow in dessert as a camel. The invention of advanced network architecture (i.e. resnet[27], transformer[85], etc.) may even enhance this misunderstanding. Recently, much research introduced the task-driven solution, attempting to discover the fundamental mechanism. (e.g. in image deraining, [71][109][36] introduced the progressive algorithm, in low-light image enhancement, [14] imitate the principle of camera imaging, in point cloud analysis, [68][69] introduced point abstraction.) However, these models still struggle in o.o.d prediction.

Causal representation learning (CRL) is a useful tool to unscramble mechanisms. CRL assumes the data are latent causal variables that are causal related and satisfy conditional SCM, using non-linear mapping. With this assumption, CRL could discover the causal relationship via estimating the distribution after intervention if the causal latent variable and SCM are learnable. Moreover, CRL could even imagine the unseen data according to the counterfactual results, making models

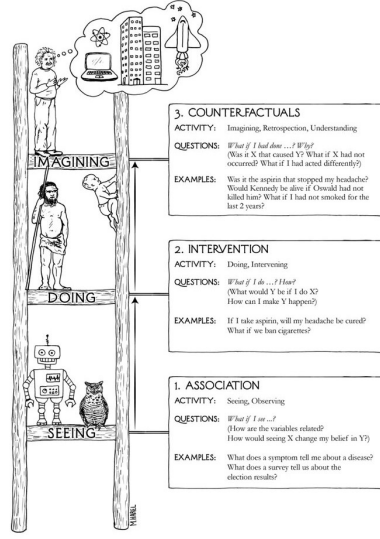


Figure 1: Three levels of Pearl Causal Hierarchy (PCH). (Drawing by [26]) The first level is association $P(y|x)$, (e.g. supervised or unsupervised learning). The second level is intervention $P(y|do(x), c)$, e.g. (feature learning, few-shot learning). The third level is counterfactual $P(y_x|x', y')$, (e.g. zero-shot learning, long-tailed classification.)

robust in the o.o.d prediction. However, distinguishing the confounder and discovering the SCM is very challenging. Therefore, some assumptions like sparsity and independent causal mechanism are introduced as an inductive bias in CRL[75]. Recently, theoretical works with CRL have been developed (e.g. Low-rank[70][6], Generalized Independent Noise condition[95][96], invariant risk minimization[3, 73, 37, 53, 54, 110, 106, 55]) and has shown promising performance in feature understanding (e.g. scene graph generation, pretraining, long-tailed data)[87][84][34, 51, 88, 98, 82, 89, 100, 30, 83, 101, 102] and transfer learning problem (e.g. adversarial methods, generalization, adaptation)[104, 97, 32, 80, 33, 31].

In this proposal, we first introduced the basic concept of causal inference (section 1), theoretical work, practical work (section 2), future research directions and objectives in medical image analysis (section 3), and expected outcomes (section 4).

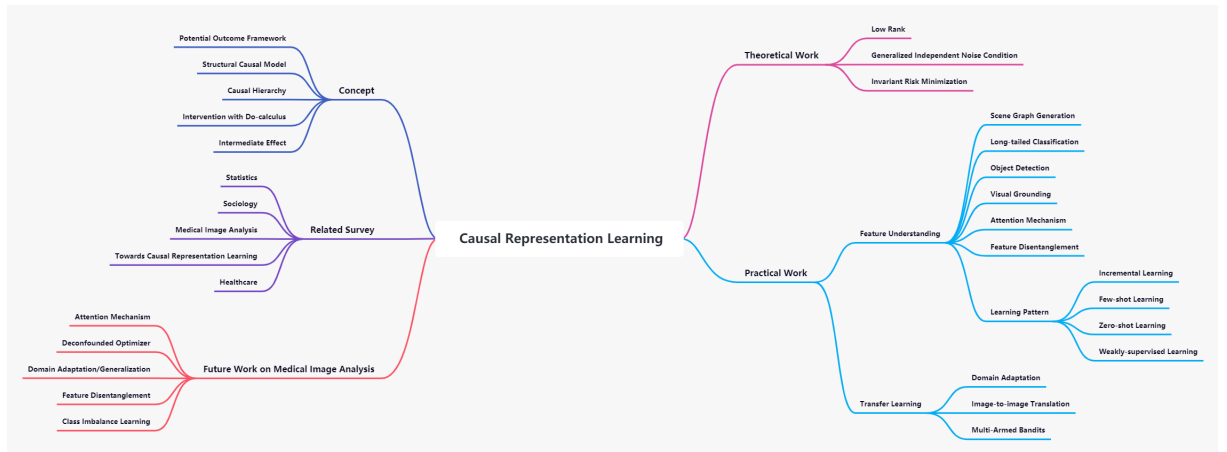


Figure 2: The mindmap of causal representation learning in vision

1 Concept of Causal Inference

In this section, several concepts of causal reasoning are introduced, including potential outcome framework, structural causal model(SCM), and causal graphs via do-calculus.

Treatment	Condition			Causal
	Mild	Severe	Total	
A	15% (210/1400)	30% (30/100)	16% (240/1500)	19.4%
B	10% (5/50)	20% (100/500)	19% (105/550)	12.9%

Table 1: The example of Simpson’s paradox. Although the total deaths of treatment A is less than deaths of treatment B, the treatment B is a worthy choice in terms of causal analysis. In naive method, for example, the treatment effect of A (16%), is given by $\frac{1400}{1500}(0.15) + \frac{100}{1500}(0.30)$. In causal analysis, the treatment effect of A (19.4%) is given by $\frac{1450}{2050}(0.15) + \frac{600}{2050}(0.30)$.

1.1 Potential outcome framework

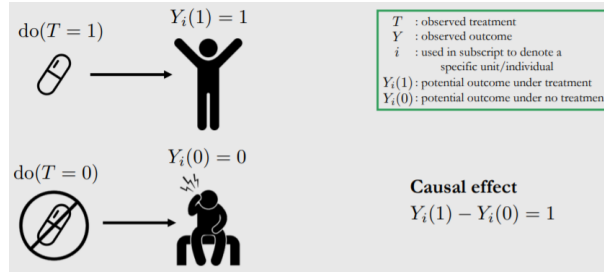


Figure 3: Example of the potential outcomes (Drawn by [60]). The treatment effect are measured on whether to take the drug or not.

An example in Fig.3 shows how the potential outcomes framework works. However, we are in a dilemma that we cannot observe the causal effect on the same person. If one person takes the drug, we will lose the information of the person not taking the drug. This dilemma is known as the "fundamental problem of causal inference"[107].

1.2 Structural Causal Model

Firstly, we define the symbols.

- X , random variable
- $P(Y_x) := P(Y | do(X = x))$
- path (X, Y) : any path from X to Y .
- collider Z , $X \rightarrow Z \leftarrow Y$, $X \perp Y$, $X \not\perp Y | Z$.

The structural causal model[63][64] is a 4-tuple $\langle \mathbf{U}, \mathbf{V}, \mathcal{F}, P(\mathbf{U}) \rangle$, where:

- \mathbf{U} is a set of background variables (exogenous variables), determined by factors outside the model.
- \mathbf{V} is a set $\{V_1, V_2, \dots, V_n\}$ of variables, called endogenous, determined by other variables within the model
- \mathcal{F} is a set of functions $\{f_1, f_2, \dots, f_n\}$, each f_i is a mapping from $U_i \cup PA_i$ (PA: parents) to V_i , where $U_i \subseteq \mathbf{U}$ and PA_i is a set of causes of V_i . The entire set \mathcal{F} forms a mapping from \mathbf{U} to \mathbf{V} . That is, for $i = 1, \dots, n$, $v_i \leftarrow f_i(pa_i, v_i)$.
- $P(\mathbf{U})$ is a probability function defined over the domain of \mathbf{U} .

For example, suppose that there exists a causal relationship between treatment solution X and lung function Y of an asthma patient. Simultaneously, suppose that Y also relies on the level

of air pollution Z . Under this circumstance, X and Y are endogenous variables, Z is exogenous variables. Therefore, the SCM can be instantiated as,

$$\begin{aligned} U &= \{Z, U_x, U_y\}, V = \{X, Y\}, F = \{f_X, f_Y\} \\ f_X &: X \leftarrow f_X(U_x) \\ f_Y &: Y \leftarrow f_Y(X, Z, U_y) \end{aligned} \quad (1)$$

1.3 Causal Hierarchy

1.3.1 Level1 seeing

For any SCM, the formula,

$$P^{\mathcal{M}}(\mathbf{Y} = \mathbf{y}) = \sum_{\{\mathbf{u} | \mathbf{Y}(\mathbf{u}) = \mathbf{y}\}} P(\mathbf{u}) \quad (2)$$

could estimate any joint distribution of $\mathbf{Y} \subset \mathbf{V}$ given by $\mathbf{Y}(U = u)$. Take the image classification task as an example. $\mathbf{V} = \mathbf{X} \cup \mathbf{Y}$, X represents the images, Y represents the labels. The aim is to model $P(\mathbf{Y} | \mathbf{X})$. At this level, we could only build the model by fitting the distribution of observational data.

1.3.2 Level2 doing

In the level of doing, a hypothesis is proposed and then verified. In this condition, a new SCM is built: $\mathcal{M}_{\mathbf{x}} = \langle \mathbf{U}, \mathbf{V}, \mathcal{F}_{\mathbf{x}}, P(\mathbf{U}) \rangle$, where $\mathcal{F}_{\mathbf{x}} = \{f_i : V_i \notin \mathbf{X}\} \cup (\mathbf{X} \leftarrow \mathbf{x})$. Therefore, $P^{\mathcal{M}}$ can be estimated by,

$$P^{\mathcal{M}}(\mathbf{Y}_{\mathbf{x}} = \mathbf{y}_{\mathbf{x}}) = \sum_{\{\mathbf{u} | \mathbf{Y}_{\mathbf{x}}(\mathbf{u}) = \mathbf{y}_{\mathbf{x}}\}} P(\mathbf{u}) \quad (3)$$

where $\mathbf{Y}_{\mathbf{x}}(\mathbf{u})$ represents $\mathcal{F}_{\mathbf{x}}(U = u)$.

1.3.3 Level3 imaging

In the level of imaging, the target is to know the effect whether another decision had been made, which can be formulated by,

$$P(Y'_{x'} | X = x, Y = y) \quad (4)$$

Namely, imaging $do(X = x')$ given $X = x, Y = y$. Based on this, the joint distribution $P^{\mathcal{M}}$ can be estimated by,

$$P^{\mathcal{M}}(\mathbf{y}_{\mathbf{x}}, \dots, \mathbf{z}_{\mathbf{w}}) = \sum_{\{\mathbf{u} | \mathbf{Y}_{\mathbf{x}}(\mathbf{u}) = \mathbf{y}_{\mathbf{x}}, \dots, \mathbf{Z}_{\mathbf{w}}(\mathbf{u}) = \mathbf{z}\}} P(\mathbf{u}) \quad (5)$$

for any $\mathbf{Y}, \mathbf{Z}, \dots, \mathbf{X}, \mathbf{W} \subset \mathbf{V}$.

1.4 Intervention with do-calculus

1.4.1 D-separation

Two sets of nodes X and Y are d-separated by a set of nodes Z if all of the paths between any node in X and any node in Y are blocked by Z . In Fig. 4, we provide an example to illustrate d-separation. If X is the cause, Y is the effect. The other node is confounding. If $W1/W2/W3$ and $M1/M2$ are conditioned on, X and Y are d-separated. If $T2$ is conditioned on, X and Y are not d-separated because the relationship between X and Y could be found by intervening $T2$. If both $T1$ and $T2$ are conditioned on, X and Y are d-separated because $T1$ blocks the information path at the bottom of the figure. This concept explains why $X \rightarrow M1 \rightarrow M2 \rightarrow Y$ is causal association and $X \rightarrow W1 \rightarrow W2 \rightarrow W3 \rightarrow Y$, $X \rightarrow T1 \rightarrow T2 \rightarrow T3 \rightarrow Y$ are non-causal associations.

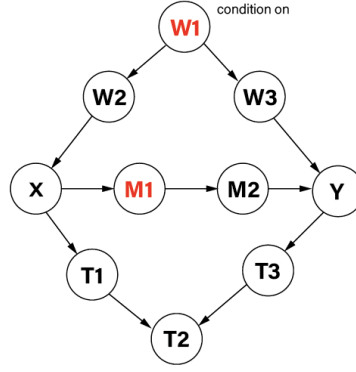


Figure 4: The demonstration for the d-separation. W1 and M1 are conditioned, which block all possible way from $X \rightarrow Y$. Therefore, X,Y are d-separated.

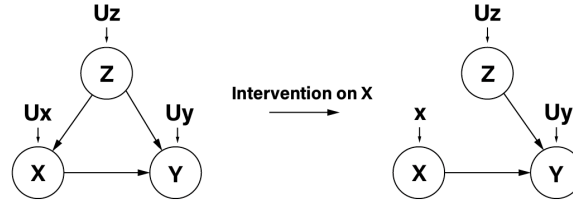


Figure 5: After the intervention on $X = x$, the edge from $Z \rightarrow X$ should be deleted.

1.4.2 Intervention

We use $do(X = x)$ to represent intervention, $P(Y = y|do(X = x))$ represents the probability of $Y = y$ when making $X = x$. In the graphical model, one edge is deleted to represent the intervention on a particular node. From Fig.5, two invariant equations can be formulated by,

$$\begin{aligned} P_m(Y = y|Z = z, X = x) &= P(Y = y|Z = z, X = x) \\ P_m(Z = z) &= P(Z = z) \end{aligned} \quad (6)$$

Z and X are d-separated after the modification, indicating that,

$$P_m(Z = z|X = x) = P_m(Z = z) = P(Z = z) \quad (7)$$

Therefore,

$$\begin{aligned} P(Y = y|do(X = x)) &= P_m(Y = y|X = x) \\ &= \sum_z P_m(Y = y|X = x, Z = z)P_m(Z = z|X = x) \\ &= \sum_z P_m(Y = y|X = x, Z = z)P_m(Z = z) \end{aligned} \quad (8)$$

Finally, the causal effect formula before intervention can be obtained using the relationship of invariance,

$$P(Y = y|do(X = x)) = \sum_z P(Y = y|X = x, Z = z)P(Z = z) \quad (9)$$

This formula is called the adjustment formula, calculating the relationship between X and Y For every value of Z .

Consider the example in Table.1 and the causal graph in Fig.5. $X = A/B$ donates to patients who take the drug A/B. $Z = Mild/Severe$ donates the level of illness. Y donates the death rate. The effect of taking the drug A:

$$E[Y|do(X = A)] = \frac{1450}{2050}(0.15) + \frac{600}{2050}(0.30) \approx 0.194 \quad (10)$$

The effect of taking the drug B:

$$E[Y|do(X = B)] = \frac{1450}{2050}(0.10) + \frac{600}{2050}(0.20) \approx 0.129 \quad (11)$$

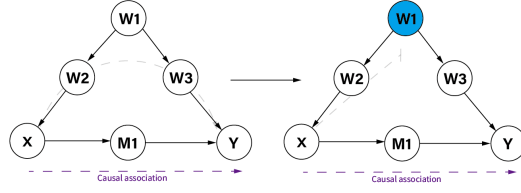


Figure 6: The example of the backdoor adjustment. The second line in Equ.12: W is a sufficient adjustment set, blocking all backdoor paths, only reserving the causation $X \rightarrow Y$. Third line in Equ.12: $do(X)$ blocks all T 's parents.

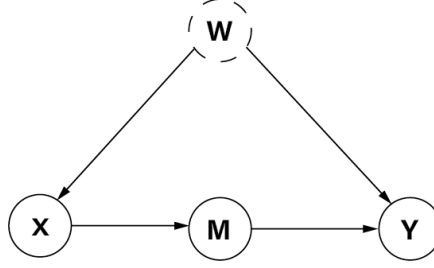


Figure 7: The example of the frontdoor adjustment. Typically, we do twice backdoor adjustments. Firstly, from $X \rightarrow M$, there is no backdoor path. Secondly, from $M \rightarrow Y$, T block the backdoor path $M \leftarrow X \leftarrow W \rightarrow Y$. Therefore, the final equation is defined in step 3.

The result indicates that treatment B has a better effect which is contradictory to the statistical results.

1.4.3 Backdoor criterion

W is said to satisfy the backdoor criterion about (X, Y) , if W :

- blocks all paths between X and Y .
- keeps the same directed paths from $X \rightarrow Y$.
- does not produce a new path.

Backdoor adjustment: if W satisfies the backdoor criterion, then ATE (Average Treatment Effect) is identified.

$$\begin{aligned}
 P(y \mid do(X)) &= \sum_w P(y \mid do(X), w)P(w \mid do(X)) \\
 &= \sum_w P(y \mid X, w)P(w \mid do(X)) \\
 &= \sum_w P(y \mid X, w)P(w)
 \end{aligned} \tag{12}$$

1.4.4 Frontdoor criterion

W is said to satisfy the backdoor criterion about (X, Y) , if:

- W blocks all possible directed path from $X \rightarrow Y$.
- There is no backdoor path from $X \rightarrow W$
- All possible paths from $W \rightarrow Y$ are blocked by X .

Frontdoor adjustment:

- X on M : $P(m \mid do(x)) = P(m \mid x)$

- M on Y: $P(y \mid do(m)) = \sum_x P(y \mid m, x)P(x)$
- X on Y: $P(y \mid do(x)) = \sum_m P(m \mid do(x))P(y \mid do(m)) = \sum_m P(m \mid x) \sum_{x'} P(y \mid m, x') P(x')$

1.4.5 Intermediate Effect

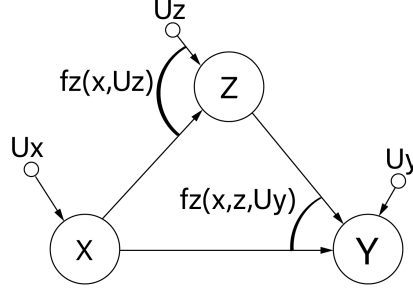


Figure 8: The mediation model without confounding.

In a causal model, a classical intermediate problem can be defined as:

$$x = f_X(U_X), z = f_Z(x, U_Z), y = f_Y(x, z, U_Y) \quad (13)$$

where X donates treatment, Z donates mediator, Y donates outcome. f_X, f_Z, f_Y are any function. U_X, U_Z, U_Y are background variables (see. Fig.8). Based on this, the effect is analyzed by the intervention as follows,

Average Treatment Effect:

$$\begin{aligned} ATE &= E[Y_1] - E[Y_0] \\ &= E[Y \mid do(X = 1)] - E[Y \mid do(X = 0)] \end{aligned} \quad (14)$$

Controlled Direct Effect:

$$\begin{aligned} CDE(z) &= E[Y_{1,Z} - Y_{0,Z}] \\ &= E[Y \mid do(X = 1, Z = z)] - E[Y \mid do(X = 0, Z = z)] \end{aligned} \quad (15)$$

Natural Direct Effect:

$$NDE = E[Y_{1,Z_0} - Y_{0,Z_0}] \quad (16)$$

Natural Indirect Effect:

$$NIE = E[Y_{0,Z_1} - Y_{0,Z_0}] \quad (17)$$

Total Direct Effect:

$$TDE = E[Y_{1,Z_1} - Y_{0,Z_1}] \quad (18)$$

Total Indirect Effect:

$$TIE = E[Y_{1,Z_1} - Y_{1,Z_0}] \quad (19)$$

2 Causal Representation Learning in Vision

In this section, we introduce recent advances of CRL in theoretical works and practical works.

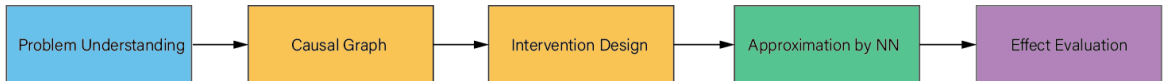


Figure 9: The general workflow for causal representation learning in vision.

2.1 Theoretical Work

Traditional machine learning methods attempt to minimize the empirical risk (ERM). However, these methods can not be generalized to the unseen domain. Moreover, ERM is often over-parameterized, resulting in the discovery of spurious correlations. Based on the concept of causal inference, invariant risk minimization[3] is introduced to solve the problems above. Consider a data representation $\Phi : \mathcal{X} \rightarrow \mathcal{H}$. we desire an invariant predictor across the environment $w \circ \Phi$. If there exists a predictor $w : \mathcal{H} \rightarrow \mathcal{Y}$ achieve optimal performance in every environment \mathcal{E}_{tr} such that $w \in \arg \min_{\bar{w} : \mathcal{H} \rightarrow \mathcal{Y}} R^e(\bar{w} \circ \Phi)$. Therefore, the constrained optimization problem is defined as:

$$\begin{aligned} \min_{\substack{\Phi : \mathcal{X} \rightarrow \mathcal{H} \\ w : \mathcal{H} \rightarrow \mathcal{Y}}} & \sum_{e \in \mathcal{E}_{\text{tr}}} R^e(w \circ \Phi) \\ \text{subject to} & w \in \arg \min_{\bar{w} : \mathcal{H} \rightarrow \mathcal{Y}} R^e(\bar{w} \circ \Phi), \text{ for all } e \in \mathcal{E}_{\text{tr}} \end{aligned} \quad (20)$$

To solve it via gradient descent method, an alternative penalty term is proposed,

$$\min_{\Phi : \mathcal{X} \rightarrow \mathcal{Y}} \sum_{e \in \mathcal{E}_{\text{tr}}} R^e(\Phi) + \lambda \cdot \|\nabla_{w|w=1.0} R^e(w \cdot \Phi)\|^2 \quad (21)$$

where D measures the risk when changing the environment, $\lambda \in [0, \infty)$ is a hyper-parameter balancing the ERM and the IRM. This paper only discussed the linear condition of w . In the colored MNIST synthetic task, the IRM method achieved 66.9% accuracy, whereas the ERM method could only achieve 17.1% accuracy, even lower than random guessing.

Despite the excellent performance on linear conditions, [73] proved that IRM can not find the optimal invariant predictor on most occasions and even suffer from a catastrophic failure in a particular condition. This paper first introduced and analyzed the non-linear scene. [37] deeply discussed the limitation of the IRM and proved that the IRM prefers an invariant predictor with worse o.o.d generalization. Moreover, the invariant loses its effect when using IRM on empirical samples rather than the population distributions. To address the problem that the IRM fails in the non-linear condition, [53] utilized the Bayesian inference to diminish the over-fitting problem and tricks like ELBO and reparameterization to accelerate convergence speed. [110] proposed sparse IRM to prevent the spurious correlation leaking to sub-models. On the dataset side, [54] proposed a Heterogeneous Risk Minimization (HRM) structure to address the mixture of multi-source data without a source label. Based on this work, [55] extended to kernel space, enhancing the ability to deal with more complex data and invariant relationships. [106] thought that the IRM method could not deal with the relationship between input and output in different domains. Therefore, [106] tried to modify the parameters to make the model robust when domain shifting based on mate-learning structure.

IRM method also mutually benefited from another theory. [46] combined information bottleneck with IRM for domain adaptation. The loss is designed in a mutual information expression, whose structure is similar to the IRM.

2.2 Practical Work

Feature understanding and transfer learning are two specific research applications in CRL. Confounders are common in vision datasets, which mislead the machine model into catching the bad relationship. Research in feature understanding in CRL attempts to build the SCM and intervene in the node to discover the causal relationship. For transfer learning, statistical methods suffer from the o.o.d data. Discovering the causal or avoiding the adaptation risk reduce the complexity of training a new transfer learning algorithm and improve performance. This section will present the recent CRL advances in feature understanding and transfer learning.

2.2.1 Feature understanding

In object detection tasks, the highly correlated objects tend to occur in the same image (e.g. the chair and human, because people could sit on the chair instead of commensalism). VC R-CNN[87] thought that observational bias made the model ignore the common causal relationship. They introduced an intervention (confounder dictionary) to measure the true causal effect.

In scene graph generation, [84] compared the counterfactual scene and factual scene, using a Total Direct Effect (TDE) analysis framework to remove the bias in training. [82] proposed Align-RCNN to discover the feature relationship and concatenate those features dynamically.

Task	Performance Improvement over 2nd Model		
Scene Graph Generation[84]	Predicate Classification	Scene Graph Classification	Scene Graph Detection
	51.1%	56.4%	31.3%
	Zero-Shot Relationship Retrieval 25.0%	Sentence-to-Graph Retrieval 33.7%	
Image Captioning[98]	Karpathy Split[38] 5 Captions	Karpathy Split Whole Set	MS-COCO[52]
	0.250%	1.55%	1.02%
Attention Mechanism[89]	CNN-Based NICO[28]	CNN-Based ImageNet-9[15]	CNN-Based ImageNet-A
	8.72%	1.15%	8.33%
	ViT-Based NICO 8.08%	ViT-Based ImageNet-9 2.78%	ViT-Based ImageNet-A 12.7%
Few-shot Learning[101]	miniImageNet	tieredImageNet	
	2.40%	0.94%	
Long-tailed Classification[83]	LVIS V1.0[25] val set	ImageNet-LT	LVIS V0.5 val set
	29.2%	17.3%	19.1%
	LVIS V0.5 eval test server 18.8%		
Incremental Learning[30]	CIFAR-100[42]	ImageNet-Sub	ImageNet-Full
	6.17%	4.76%	3.49%
Image Recognition[87]	MS-COCO	Open Images[44]	VQA2.0 test[2]
	1.41%	1.09%	0.560%
Visual Grounding[34]	RefCOCO[39]	RefCOCO+	RefCOCOg
	2.09%	2.36%	1.77%
	ReferIt Game 1.14%	Flickr30K Entities[67] 0.36%	
Weakly-Supervised Learning[102]	PASCAL VOC 2012[19]	MSCOCO	
	1.84%	2.45%	

Table 2: The performance improvement table for causal representation learning in practical works. Most of the works are plug-and-play and robust to other downstream tasks with only a few costs in total parameters.

In visual grounding (visual language tasks), the location of the target bounding box highly depends on the query instead of causal reasoning. [34] proposed a plug-and-play framework Referring Expression Deconfounder (RED), to make the backdoor adjustment to find the causal relationship between images and sentences.

In image captioning (visual language tasks), [98] thought that the pre-training model contains the confounder. The authors introduced the backdoor adjustment to deconfound the bias. Few-shot learning[91] also requires the pre-training model. Similarly, [101] proposed three solutions, including feature-wise adjustment, class-wise adjustment, and class-wise adjustment. These operations do not need to modify the backbone and could be applied easily in zero-shot learning, meta-learning[29] etc.

In the attention mechanism, most models worked well due to the i.i.d of the data. However, the o.o.d data degrades the performance when using attention. [89] proposed Casual Attention Module (Caam) on original CBAM-based CNN[92] and ViT[16]. This method utilized the IRM and adversarial training[23] with a partitioned dataset to discover confounding factor characteristics.

In feature (disentanglement) representation learning, the classical work simCLR[12] proposed a Self-Supervised Learning (SSL), using a contrastive objective method to recognize similar images. However, the generalization performance and the interpretability are poor. [88] introduced Iterative Partition-based Invariant Risk Minimization (IP-IRM), combining feature disentanglement (data partition) and IRM. This method could discover the critical causal representation for classification tasks. [100] proposed a counterfactual generation framework for zero-shot learning tasks[45] based on counterfactual faithfulness theorem. It designed a two-element classifier, disentangling the feature in class and sample.

In incremental learning[59], [30] used causal-effect theory to explain the forget and anti-forget. It designed models for distilling the causal effect of data, colliding effect, and removing SGD momentum[81] (optimizer) effect of guaranteeing the effectiveness of introducing new data to learn. The interference of the SGD momentum also occurs in long-tailed classification because the update direction of SGD contains the information on data distribution. [83] designed a multi-head normalized classifier in training and made counterfactual TDE inference in testing to remove the excessive tendency for head class.

In weakly-supervised learning[111] for semantic segmentation, the problems often occur in pseudo-masks, for example, object ambiguity, incomplete background, and incomplete foreground. [102] introduced a causal intervention—blocking the connection between context prior and images to remove the fake association between label and images. [102] proposed a context adjustment for the unknown context prior, that is, using a class-specific average mask to approximately constructing a confounder set. [13] proposed a category causality chain and an anatomy-causality chain to solve the ambiguous boundary and co-occurrence problems in medical image segmentation.

2.2.2 Transfer learning

One of the popular research directions in transfer learning is domain adaptation. [80] defined invariance specification: a 2-tuple $\langle \mathcal{P}, \mathbf{M} \rangle$, where \mathcal{P} donates graphical representation, \mathbf{M} donates a group of variable (control the data from different environments). If an invariance spec is found, we could get a stable representation such that the model can be applied to the target domain. Similarly, [104] considered domain adaptation as an inference problem, constructing a DAG and solving it by Bayesian inference.

In image-to-image translation, [97] concluded a DAG and developed an important reweighting-based learning method. This method can automatically select the images and perform translation simultaneously.

In general, transfer learning, [43] introduced the new problem of transfer learning for estimating heterogeneous treatment effects and developed several methods (e.g. Y-Learner). [72] proposed invariant models for transfer learning. [103] utilized a causal approach to Multi-Armed Bandits in reinforcement learning.[58] exploited causal inference to predict invariant conditional distribution in domain adaptation without prior knowledge of the causal graph and the type of interventions.

3 Research Objectives on Medical Image Analysis

Medical image analysis is a high-risk task, significantly requiring an explainable framework so that doctors and patients can rely on the diagnosis. The current works still lack interpretability and

are treated as a black box. Causal representation learning is a promising learning paradigm for medical image analysis. In this section, we mention some prospective research directions.

3.1 Attention mechanism

Attention mechanism is widely applied in medical image analysis, and has shown promising results in lots of datasets and tasks[112][77][90][1][4][47], especially for pure attention model (e.g. ViT[16], Swin-Transformer[57], Swin-UNet[8]). The attention mechanism will bring interpretability to the model. For example, in organ segmentation (e.g. cardiac, brain), attention could highlight the features of a region and suppress other noisy parts. However, the attention mechanism suffers from the data distribution shift (CNN-based attention) and the small scale of datasets (Transformer based). [22] shown that the results are noisy on some datasets, even for the cases of attention mechanisms. The causal attention module (Caam)[84] is a promising method to solve this problem without changing the original framework.

3.2 Deconfounded optimizer

Most of the works have various settings of the optimizer, and the performance will be damaged if we change to another optimizer. As [30][83] mentioned above, the optimizer (e.g. SGD momentum) can be a confounder in incremental learning and long-tailed classification. The design of a multi-head normalized classifier and counterfactual TDE inference could be a solution in the medical field.

3.3 Domain adaptation/generalization

The data distribution shift problem would decrease the performance of the original model due to the various sources from different hospitals. Recently, domain adaptation[93][11][17][18][94][24] and domain generalization[56][105][48][49][50] techniques increased a great deal of accuracy (from $\approx 10\%$ to $\approx 70\%$ in these tasks. However, the model must be trained again when adding a new source for adaptation, and the tricks for domain generalization heavily depend on data pre-processing. Additionally, the performance ($\approx 70\%$) still can not be trusted and applied in clinical application. The series works of IRM [3] provide a new general solution for domain adaptation/generalization problems.

3.4 Feature disentanglement

Feature disentanglement attempts to desperate independent feature to make the model explainable. Specifically, classical feature disentanglement methods utilized the Variational Autoencoder[40] or GAN[23] with a restriction in different channels (e.g. minimize mutual information) to disentangle the high-level semantic representation. In domain adaptation, [66] aimed to find the domain-specific feature and the domain invariant feature to make the model robust. In multi-task learning, [10] proposed a dual-stream network to share the common feature in latent space. [9] utilized a VAE to learn a multi-channel spatial representation of the anatomy. However, the restriction for disentanglement is still loose and lacks interpretability in the current work. We could refer to the concept of IP-IRM[88] to discover the causal representation of medical images.

3.5 Class imbalance learning

The solution to the class imbalance problem traditionally relies on the data pre-processing (e.g. oversampling[41], re-weighting[108] etc.). But, we can not know the data distribution before training. Additionally, the trick, like re-weighting, will lead the head categories under-fitted. We could refer to [83] to invent a multi-head normalized classifier in training and make counterfactual TDE inference in testing to solve the long-tailed problem.

4 Expected Outcomes

The most intuitive result of causal representation learning is to improve the accuracy of tasks (e.g., detection and segmentation). More importantly, causal representation learning has a stronger interpretability of results. Here, we will give an example of outcomes using causal representation

learning in self-supervised anomaly detection in medical image segmentation[7]. In the case of abnormal lung lesions, we tried to capture the invariant characteristics and trained an auto-encoder. This auto-encoder ensured that the reconstruction loss was minimal. If the detected image contained the characteristics of the exception, the reconstruction loss would be significant. Therefore, invariant risk minimization could be applied.

$$L_{\text{IRM}}(\Phi, w) = \sum_{e \in \mathcal{E}_{\text{tr}}} R^e(w \circ \Phi) + \lambda \cdot D(w, \Phi, e) \quad (22)$$

For implementation:

```
scale = torch.Tensor([1.0]).cuda().require_grad_()
loss_rec = rec.err.mean()*scale.mean()
penalty_irm = torch.autograd.grad(loss_rec, scale, create_graph=True)[0]
penalty = torch.sum(penalty_irm**2)*0.2
```

	AE	AE+IRM
Rec	0.669	0.673
Intra	0.694	0.715
Inter	0.815	0.816
Test	0.672	0.680

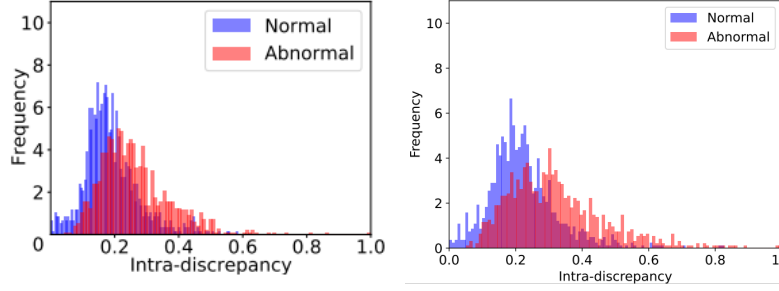


Figure 10: Classification accuracy table (unit: % and classification distribution figures (left:AE, right:AE+IRM)).

The results showed IRM improved classification accuracy. At the same time, the reasons for the improvement can also be explained. As shown in the above analysis, IRM loss captures the domain invariant features, leading to a significant increase in abnormal image reconstruction loss. Therefore, the reconstructed fraction distribution of the abnormal image moves to the right, making it easier to distinguish from the normal image.

5 Conclusion and Prospect

This research proposal reviewed the development of causal representation learning from concept to application. Firstly, we introduced the basic knowledge of causal inference. Secondly, we analyze the theoretical works on IRM and practical works on feature understanding and transfer learning. Finally, we proposed research objectives for medical image analysis and showed an example of the research outcome.

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