Context-Adaptive Statistical Inference: Recent Progress, Open Problems, and Opportunities for Foundation Models

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Abstract

Context-adaptive inference offers a promising way to improve statistical methods, particularly by using foundation models to provide context. Recent progress is reviewed, key challenges are identified, and practical opportunities for integration are explored. By combining these approaches, we can unlock new possibilities for making statistical methods more effective and adaptable in real-world scenarios, driving meaningful advances in both research and practice.

Introduction

Purpose and Scope

TODO: Establishing the framework for our examination of context-adaptive statistical methods and the significance of foundation models.

Conceptual Foundations

Unpacking the core principles and historical impact of adaptive methods within statistical modeling.

A Brief History of Personalized Inference

Personalization aims to solve the problem of *parameter heterogeneity*, where model parameters are *sample-specific*.

$$X_i \sim P(X; heta_i)$$

From N observations, personalized modeling methods aim to recover N parameter estimates $\hat{\theta}_1,\ldots,\hat{\theta}_N$. Without further assumptions this problem is ill-defined, and the estimators have far too much variance to be useful. We can begin to make this problem tractable by imposing assumptions on the topology of θ , or the relationship between θ and contextual variables.

Population Models

The fundamental assumption of most models is that samples are independent and identically distributed. However, if samples are identically distributed they must also have identical parameters. To account for parameter heterogeneity and create more realistic models we must relax this assumption, but the assumption is so fundamental to many methods that alternatives are rarely explored. Additionally, many traditional models may produce a seemingly acceptable fit to their data, even when the underlying model is heterogeneous. Here, we explore the consequences of applying homogeneous modeling approaches to heterogeneous data, and discuss how subtle but meaningful effects are often lost to the strength of the identically distributed assumption.

Failure modes of population models can be identified by their error distributions.

Mode collapse: If one population is much larger than another, the other population will be underrepresented in the model.

Outliers: Small populations of outliers can have an enormous effect on OLS models in the parameter-averaging regime.

Phantom Populations: If several populations are present but equally represented, the optimal traditional model will represent none of these populations.

Lemma: A traditional OLS linear model will be the average of heterogeneous models.

Context-informed models

Conditional and Cluster Models

While conditional and cluster models are not truly personalized models, the spirit is the same. These models make the assumption that models in a single conditional or cluster group are homogeneous. More commonly this is written as a group of observations being generated by a single model. While the assumption results in fewer than N models, it allows the use of generic plug-in estimators. Conditional or cluster estimators take the form

$$\hat{ heta}_0, \dots, \hat{ heta}_C = rg\max_{ heta_0, \dots, heta_C} \sum_{c \in \mathcal{C}} \ell(X_c; heta_c)$$

where $\ell(X;\theta)$ is the log-likelihood of θ on X and c specifies the covariate group that samples are assigned to, usually by specifying a condition or clustering on covariates thought to affect the distribution of observations. Notably, this method produces fewer than N distinct models for N samples and will fail to recover per-sample parameter variation.

Distance-regularized Models

Distance-regularized models assume that models with similar covariates have similar parameters and encode this assumption as a regularization term.

$$(\hat{ heta}_0,\dots,\hat{ heta}_N = rg\max_{ heta_0,\dots, heta_N} \sum_i \left[\ell(x_i; heta_i)
ight] - \sum_{i,j} rac{\| heta_i - heta_j\|}{D(c_i,c_j)}$$

The second term is a regularizer that penalizes divergence of θ 's with similar c.

Parametric Varying-coefficient models

Original paper (based on a smoothing spline function): [1] Markov networks: [2] Linear varying-coefficient models assume that parameters vary linearly with covariates, a much stronger assumption than the classic varying-coefficient model but making a conceptual leap that allows us to define a form for the relationship between the parameters and covariates.

$$\hat{ heta}_0, \dots, \hat{ heta}_N = \widehat{A}C^T \ \widehat{A} = rg\max_{A} \sum_i \ell(x_i; Ac_i)$$

Semi-parametric varying-coefficient Models

Original paper: [3] 2-step estimation with RBF kernels: [4]

Classic varying-coefficient models assume that models with similar covariates have similar parameters, or – more formally – that changes in parameters are smooth over the covariate space. This assumption is encoded as a sample weighting, often using a kernel, where the relevance of a sample to a model is equivalent to its kernel similarity over the covariate space.

$$\hat{ heta}_0, \dots, \hat{ heta}_N = rg\max_{ heta_0, \dots, heta_N} \sum_{i,j} rac{K(c_i, c_j)}{\sum_k K(c_i, c_k)} \ell(x_j; heta_i)$$

This estimator is the simplest to recover N unique parameter estimates. However, the assumption here is contradictory to the partition model estimator. When the relationship between covariates and parameters is discontinuous or abrupt, this estimator will fail.

Contextualized Models

Seminal work [5] Contextualized ML generalization and applications: [6], [7], [8], [9], [10], [11], [12], [13]

Contextualized models make the assumption that parameters are some function of context, but make no assumption on the form of that function. In this regime, we seek to estimate the function often using a deep learner (if we have some differentiable proxy for probability):

$$\widehat{f} = rg \max_{f \in \mathcal{F}} \sum_{i}^{\cdot} \ell(x_i; f(c_i))$$

Latent-structure Models

Partition Models

Markov networks: [14] Partition models also assume that parameters can be partitioned into homogeneous groups over the covariate space, but make no assumption about where these partitions occur. This allows the use of information from different groups in estimating a model for a each covariate. Partition model estimators are most often utilized to infer abrupt model changes over time and take the form

$$\hat{ heta}_0, \dots, \hat{ heta}_N = rg\max_{ heta_0, \dots, heta_N} \sum_i \ell(x_i; heta_i) + \sum_{i=2}^N \mathrm{TV}(heta_i, heta_{i-1})$$

Where the regularizaiton term might take the form

$$ext{TV}(heta_i, heta_{i-1}) = | heta_i - heta_{i-1}|$$

This still fails to recover a unique parameter estimate for each sample, but gets closer to the spirit of personalized modeling by putting the model likelihood and partition regularizer in competition to find the optimal partitions.

Fine-tuned Models and Transfer Learning

Review: [15] Noted in foundational literature for linear varying coefficient models [3]

Estimate a population model, freeze these parameters, and then include a smaller set of personalized parameters to estimate on a smaller subpopulation.

$$egin{aligned} \widehat{\gamma} &= rg \max_{\gamma} = \ell(\gamma; X) \ \widehat{ heta}_c &= rg \max_{ heta_c} = \ell(heta_c; \widehat{\gamma}, X_c) \end{aligned}$$

Context-informed and Latent-structure models

Seminal paper: [16]

Key idea: negative information sharing. Different models should be pushed apart.

$$\hat{ heta}_0, \dots, \hat{ heta}_N = rg\max_{ heta_0, \dots, heta_N, D} \sum_{i=0}^N \prod_{i=0s,t,D(c_i, c_i) < d}^N P(x_j; heta_i) P(heta_i; heta_j)$$

Theoretical Foundations and Advances in Varying-Coefficient Models

Principles of Adaptivity

TODO: Analyzing the core principles that underpin adaptivity in statistical modeling.

Advances in Varying-Coefficient Models

TODO: Outlining key theoretical and methodological breakthroughs.

Integration with State-of-the-Art Machine Learning

TODO: Assessing the enhancement of VC models through modern ML technologies (e.g. deep learning, boosted trees, etc).

Context-Invariant Training

TODO: The converse of VC models, exploring the implications of training context-invariant models. e.g. out-of-distribution generalization, robustness to adversarial attacks.

Context-Adaptive Interpretations of Context-Invariant Models

In the previous section, we discussed the importance of context in model parameters. Such context-adaptive models can be learned by explicitly modeling the impact of contextual variables on model parameters, or learned implicitly in a model containing interaction effects between the context and the input features. In this section, we will focus on recent progress in understanding how context influences interpretations of statistical models, even when the model was not originally designed to incorporate context.

TODO: Discussing the implications of context-adaptive interpretations for traditional models. Related work including LIME/DeepLift/DeepSHAP.

Opportunities for Foundation Models

Expanding Frameworks

TODO: Define foundation models, Explore how foundation models are redefining possibilities within statistical models.

Foundation models as context

TODO: Show recent progress and ongoing directions in using foundation models as context.

Applications, Case Studies, and Evaluations

Implementation Across Sectors

TODO: Detailed examination of context-adaptive models in sectors like healthcare and finance.

Performance Evaluation

TODO: Successes, failures, and comparative analyses of context-adaptive models across applications.

Technological and Software Tools

Survey of Tools

TODO: Reviewing current technological supports for context-adaptive models.

Selection and Usage Guidance

TODO: Offering practical advice on tool selection and use for optimal outcomes.

Future Trends and Predictions

Emerging Technologies

TODO: Identifying upcoming technologies and predicting their impact on context-adaptive learning.

Advances in Methodologies

TODO: Speculating on potential future methodological enhancements.

Open Problems

Theoretical Challenges

TODO: Critically examining unresolved theoretical issues like identifiability, etc.

Ethical and Regulatory Considerations

TODO: Discussing the ethical landscape and regulatory challenges, with focus on benefits of interpretability and regulatability.

Complexity in Implementation

TODO: Addressing obstacles in practical applications and gathering insights from real-world data.

TODO: Other open problems?

Conclusion

Overview of Insights

TODO: Summarizing the main findings and contributions of this review.

Future Directions

TODO: Discussing potential developments and innovations in context-adaptive statistical inference.

References

1. Varying-Coefficient Models

Trevor Hastie, Robert Tibshirani

Journal of the Royal Statistical Society Series B: Statistical Methodology (1993-09-01)

https://doi.org/gmfvmb

DOI: 10.1111/j.2517-6161.1993.tb01939.x

2. Bayesian Edge Regression in Undirected Graphical Models to Characterize Interpatient Heterogeneity in Cancer

Zeya Wang, Veerabhadran Baladandayuthapani, Ahmed O Kaseb, Hesham M Amin, Manal M Hassan, Wenyi Wang, Jeffrey S Morris

Journal of the American Statistical Association (2022-01-05) https://doi.org/gt68hr

DOI: 10.1080/01621459.2021.2000866 · PMID: 36090952 · PMCID: PMC9454401

3. Statistical estimation in varying coefficient models

Jianqing Fan, Wenyang Zhang

The Annals of Statistics (1999-10-01) https://doi.org/dsxd4s

DOI: <u>10.1214/aos/1017939139</u>

4. Time-Varying Coefficient Model Estimation Through Radial Basis Functions

Juan Sosa, Lina Buitrago

arXiv (2021-03-02) https://arxiv.org/abs/2103.00315

5. **Contextual Explanation Networks**

Maruan Al-Shedivat, Avinava Dubey, Eric P Xing

arXiv(2017) https://doi.org/gt68h9

DOI: 10.48550/arxiv.1705.10301

6. **Contextualized Machine Learning**

Benjamin Lengerich, Caleb N Ellington, Andrea Rubbi, Manolis Kellis, Eric P Xing

arXiv(2023) https://doi.org/gt68jg

DOI: 10.48550/arxiv.2310.11340

7. NOTMAD: Estimating Bayesian Networks with Sample-Specific Structures and Parameters

Ben Lengerich, Caleb Ellington, Bryon Aragam, Eric P Xing, Manolis Kellis

arXiv(2021) https://doi.org/gt68jc

DOI: 10.48550/arxiv.2111.01104

8. Contextualized: Heterogeneous Modeling Toolbox

Caleb N Ellington, Benjamin J Lengerich, Wesley Lo, Aaron Alvarez, Andrea Rubbi, Manolis Kellis, Eric P Xing

Journal of Open Source Software (2024-05-08) https://doi.org/gt68h8

DOI: 10.21105/joss.06469

9. Contextualized Policy Recovery: Modeling and Interpreting Medical Decisions with Adaptive Imitation Learning

Jannik Deuschel, Caleb N Ellington, Yingtao Luo, Benjamin J Lengerich, Pascal Friederich, Eric P Xing

arXiv (2023) https://doi.org/gt68jf

DOI: 10.48550/arxiv.2310.07918

10. Automated interpretable discovery of heterogeneous treatment effectiveness: A COVID-19 case study

Benjamin J Lengerich, Mark E Nunnally, Yin Aphinyanaphongs, Caleb Ellington, Rich Caruana *Journal of Biomedical Informatics* (2022-06) https://doi.org/gt68h5

DOI: <u>10.1016/j.jbi.2022.104086</u> · PMID: <u>35504543</u> · PMCID: <u>PMC9055753</u>

11. Discriminative Subtyping of Lung Cancers from Histopathology Images via Contextual Deep Learning

Benjamin J Lengerich, Maruan Al-Shedivat, Amir Alavi, Jennifer Williams, Sami Labbaki, Eric P Xing

Cold Spring Harbor Laboratory (2020-06-26) https://doi.org/gt68h6

DOI: 10.1101/2020.06.25.20140053

12. Contextualized Networks Reveal Heterogeneous Transcriptomic Regulation in Tumors at Sample-Specific Resolution

Caleb N Ellington, Benjamin J Lengerich, Thomas BK Watkins, Jiekun Yang, Hanxi Xiao, Manolis Kellis, Eric P Xing

Cold Spring Harbor Laboratory (2023-12-04) https://doi.org/gt68h7

DOI: <u>10.1101/2023.12.01.569658</u>

13. Contextual Feature Selection with Conditional Stochastic Gates

Ram Dyuthi Sristi, Ofir Lindenbaum, Shira Lifshitz, Maria Lavzin, Jackie Schiller, Gal Mishne, Hadas Benisty

arXiv (2023) https://doi.org/gt68jh
DOI: 10.48550/arxiv.2312.14254

14. Estimating time-varying networks

Mladen Kolar, Le Song, Amr Ahmed, Eric P Xing

The Annals of Applied Statistics (2010-03-01) https://doi.org/b3rn6q

DOI: <u>10.1214/09-aoas308</u>

15. When Personalization Harms: Reconsidering the Use of Group Attributes in Prediction

Vinith M Suriyakumar, Marzyeh Ghassemi, Berk Ustun

arXiv (2022) https://doi.org/gt68jd
DOI: 10.48550/arxiv.2206.02058

16. Learning Sample-Specific Models with Low-Rank Personalized Regression

Benjamin Lengerich, Bryon Aragam, Eric P Xing

arXiv (2019) https://doi.org/gt68jb
DOI: 10.48550/arxiv.1910.06939