

AN INTRODUCTION TO DIFFERENCE-IN-DIFFERENCES AND SYNTHETIC CONTROL METHODS FOR EPIDEMIOLOGISTS

11 November 2022

IAPHS Annual Meeting Workshop

Roch Nianogo and Tarik Benmarhnia





AGENDA FOR TODAY

- Logistics and introductions
- Part I Tarik ~ 80 min
- **A**&Q •
- Break
- Part II Roch ~ 90-120 min
- **A**&Q •
- End



OUTLINE - PART 1

- Causal inference, natural experiments and quasiexperimental methods
- Difference-in-differences methods
- Interrupted Time Series
- Synthetic Control Methods
- More advanced topics



WORKSHOP OBJECTIVES

- Understand the conceptual foundations of quasi-experimental methods based on the timing of a natural experiment
- 2. Understanding how Difference-in-Differences (DID), Interrupted Time Series (ITS) and Synthetic Control Methods (SCM) work and their specific identification assumptions.
- 3. Being able to implement (in R) DID, ITS and SCM in different settings, interpret estimands of interest and check/visualize possible assumptions' violations \rightarrow Part II



RANDOMIZATION AS A SOLUTION TO DEAL WITH CONFOUNDING

- Randomization has been proposed as a solution to deal with confounding
 - Complying with exchangeability between exposed and non-exposed to the policy/treatment
 - For both measured & unmeasured confounding



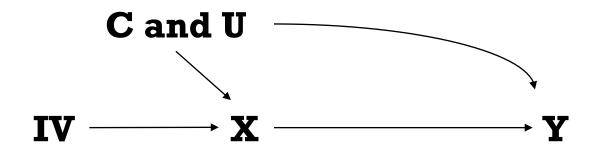
IDEAL RANDOMIZED EXPERIMENTS

- What makes an ideal randomized experiment:
 - No loss to follow-up,
 - Full adherence to the assigned treatment over the duration of the study,
 - A single version of treatment, and double-blind assignment
- Ideal randomized experiments are unrealistic but useful to introduce some key concepts for causal inference



THE IDEA BEHIND RANDOWIZATION

- How do RCT work ...
- Randomization can be analyzed as an "Instrumental Variable"
- The overall aim is to deal with <u>both measured and unmeasured</u> confounding

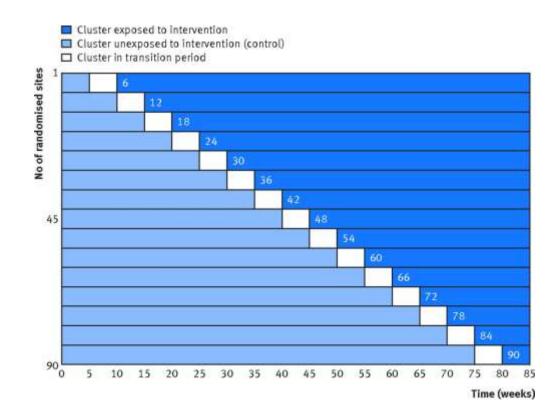


- The problem: in many cases RCT are not feasible due costs, scale or ethical objections
- Or because it is simply too late ..



DIFFERENT TYPES OF RCTS

- Two-arm, parallel design
- Planned cross-over design
- Factorial design
- Cluster randomized trials
- Stepped Wedge Designs
 - Based on the timing of the intervention



Hemming, Karla, et al. "The stepped wedge cluster randomised trial: rationale, design, analysis, and reporting." Bmj 350 (2015): h391.



USING NATURAL EXPERIMENTS

- By capitalizing on natural experiments, Quasi-experimental methods (QEM) can be used as alternatives to experimental methods to provide causal estimates from observational studies.
- The term quasi-experiment refers to:
 - "experiments that have treatments, outcome measures, and experimental units, but do not use random assignment to create the comparisons from which treatment-caused change is inferred" (Cook et al. 1979)
- Different QEM, different configurations, different assumptions to draw causal inference
- To partially deal with both measured and unmeasured confounding



EXAMPLES OF NATURAL EXPERIMENTS NOT ONLY FOR POLICY EVALUATION

- Public policies:
 - Smoking ban
 - Legalization of marijuana
- Clinical Treatments, Vaccination
- Conditional Cash Transfers
- Natural Hazards
 - Earthquakes
 - Wildfires

2 main types of natural experiments

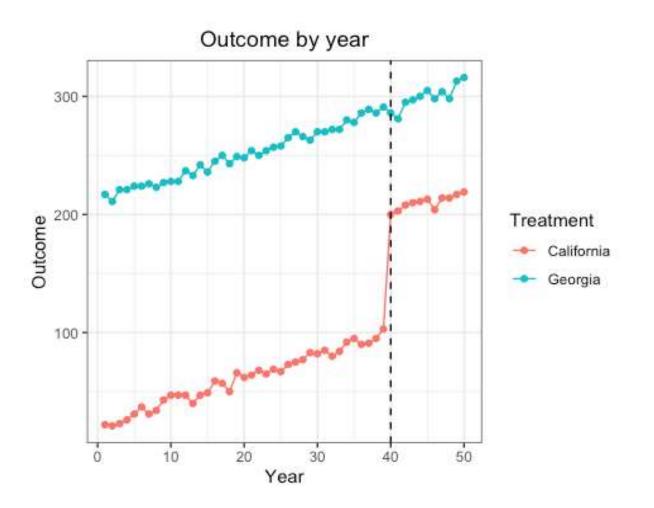
- Timing of the intervention
- Eligibility to a specific policy (e.g. age for vaccination) and other IV strategies

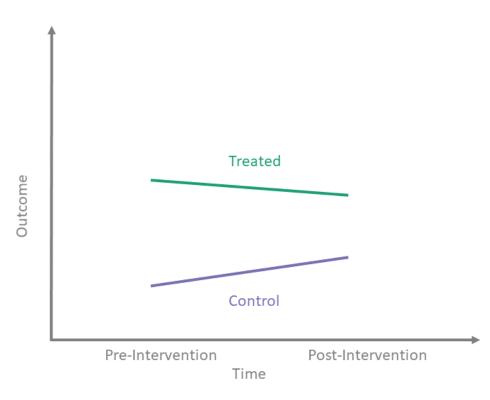


DIFFERENCE-IN-DIFFERENCES



The difference-in-differences idea





By Bret Zeldow and Laura Hatfield



DIFFERENCE-IN-DIFFERENCES THE STANDARD APPROACH

To estimate the effect of interest

$$(\mu_{11} - \mu_{10}) - (\mu_{01} - \mu_{00})$$

 $i = 0$ is control group, $i = 1$ is treatment.
 $t = 0$ is pre-period, $t = 1$ is post-period.

- Simple Regression modeling approach (with 2 groups)
 - $E[Y \mid a, t, did] = \beta_0 + \beta_1 a + \beta_2 t + \beta_3 did$
 - Where:
 - a represents the group with the policy
 - t represents the period after the policy implementation
 - DID is the interaction between a and t

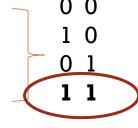


HOW DOES IT WORK?



$$E[Y \mid a, t, did] = \beta_0 + \beta_1 t + \beta_2 a + \beta_3 [a*t]$$

Like a double fixed effect



You can also include more complex time trends



DID ASSUMPTIONS

- •The key assumptions of the DiD analysis are:
 - 1. The trend in the control group represents a good approximation for the counterfactual trend of the treated group in the absence of the treatment.
 - 2. Common Shock Assumption
 - 3. No spillover



MANY APPLICATIONS

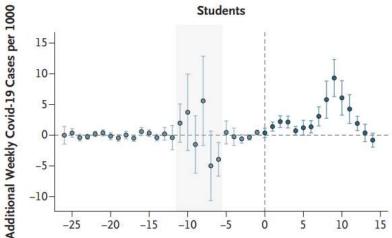
The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Lifting Universal Masking in Schools — Covid-19 Incidence among Students and Staff

Tori L. Cowger, Ph.D., M.P.H., Eleanor J. Murray, Sc.D., M.P.H., Jaylen Clarke, M.Sc., Mary T. Bassett, M.D., M.P.H., Bisola O. Ojikutu, M.D., M.P.H., Sarimer M. Sánchez, M.D., M.P.H., Natalia Linos, Sc.D., and Kathryn T. Hall, Ph.D., M.P.H.

Weekly Difference between Districts That Lifted Masking and Districts That Sustained Masking



Removing user fees for facility-based delivery services: a difference-in-differences evaluation from ten sub-Saharan African countries

Britt McKinnon,1* Sam Harper,1 Jay S Kaufman1 and Yves Bergevin2

JAMA Pediatrics | Original Investigation

Difference-in-Differences Analysis of the Association Between State Same-Sex Marriage Policies and Adolescent Suicide Attempts

Texting Bans and Fatal Accidents on Roadways: Do They Work? Or Do Drivers Just React to Announcements of Bans?[†]

By Rahi Abouk and Scott Adams*

Evaluating the effect of hierarchical medical system on health seeking behavior: A difference-in-differences analysis in China

Zhongliang Zhou a, Yaxin Zhao b, , Chi Shen , Sha Lai , Rashed Nawaz , Jianmin Gao

Gotta catch'em all! Pokémon GO and physical activity among young adults: difference in differences study

Katherine B Howe, 1.2 Christian Suharlim, 3 Peter Ueda, 4.5 Daniel Howe, Ichiro Kawachi, 2 Eric B Rimm 1.6.7

DID COUPLED WITH PROPENSITY SCORE METHODS

- When <u>multiple control groups are available</u>, it is possible to use available information on time-varying and time-fixed confounders
- We can use propensity score matching and IPTW for example
- By doing so, we aim at identifying similar observations at each time point in the control groups

Using propensity scores in difference-in-differences models to estimate the effects of a policy change

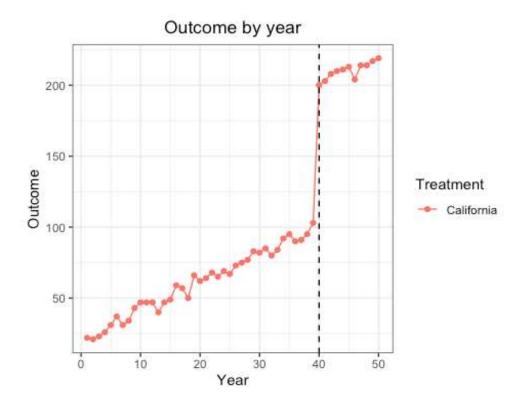
Elizabeth A. Stuart · Haiden A. Huskamp · Kenneth Duckworth · Jeffrey Simmons · Zirui Song · Michael E. Chernew · Colleen L. Barry

LETTER

Quantifying the impact of changing the threshold of New York City heat emergency plan in reducing heat-related illnesses



INTERRUPTED TIME SERIES: DID WITHOUT CONTROL GROUPS



Estimation

The traditional approach:

The following model can be used to estimate the effect of the policy on the outcome y $y = \alpha + \beta_1(year) + \beta_2(post) + \beta_3(xi) + \beta_4(xt) + \beta_5(xit) + \beta_6(year \times post) + \epsilon$

Can be also done through a 2-stage approach:

- 1. Building and optimizing a predictive model for Y in the pre-treatment period
- Predicting Y in the post-treatment period using the model developed in stage 1 and compare with observed outcomes

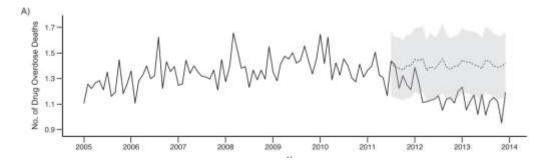
Various approaches can be used for stage 1 (ARIMA, random forest, and other ML algorithms)



ITS EXAMPLES

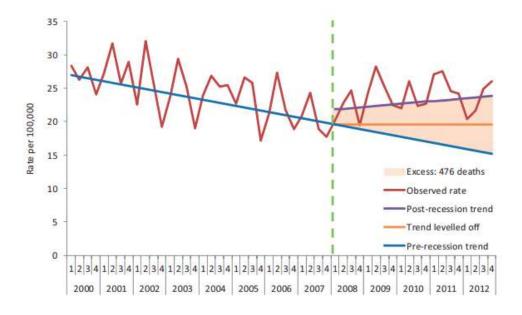
Florida's Opioid Crackdown and Mortality From Drug Overdose, Motor Vehicle Crashes, and Suicide: A Bayesian Interrupted Time-Series Analysis

Kenneth A. Feder, Ramin Mojtabai, Elizabeth A. Stuart*, Rashelle Musci, and Elizabeth J. Letourneau



Impact of the economic recession and subsequent austerity on suicide and self-harm in Ireland: An interrupted time series analysis

Paul Corcoran, 1,2* Eve Griffin, 1 Ella Arensman, 1,2 Anthony P Fitzgerald, 2 and Ivan J Perry 2





CONTROLLED ITS OR DID?

- When using one (or more) control group.s, there is no fundamental distinction between Controlled ITS and DID models
- They yield the same counterfactuals and identify the same treatment effects.
- The terminology simply varies across disciplines



ALTERNATIVE OPTIONS FOR SELECTING CONTROL GROUPS

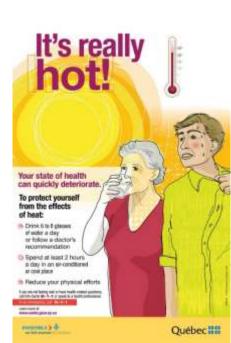
- Besides geographical units that did not receive the treatment/policy, it is possible to consider different types of control groups
- It is possible to use different outcomes or population subgroups to contrast the change in the outcome over time
 - For example, if a given policy only targets individuals above 65 years, it will be possible to use the 64 years of age and below subgroup as a reference group



A Difference-in-Differences Approach to Assess the Effect of a Heat Action Plan on Heat-Related Mortality, and Differences in Effectiveness According to Sex, Age, and Socioeconomic Status (Montreal, Quebec)

Tarik Benmarhnia, 1 Zinzi Bailey, 1 David Kaiser, 2 Nathalie Auger, 3 Nicholas King, 4,5 and Jay S. Kaufman 1,5

- •The Montreal heat warning system
 - Implemented in 2004
 - 'active watch' alert level, when daily max temperatures exceed 30°C
 - Focus on vulnerable populations
 - Age
 - SES
 - Gender as a placebo



THE ANALYTICAL APPROACH:

- Assigning days (the unit of analysis) to two groups: an "intervention" group that meets an eligibility criterion and a "non-intervention" group that does not.
- the "intervention" group: if 'active watch' alert level is present (heat wave days)
- The non-intervention group: non heat wave days
- The **counterfactual quantity** being estimated is:
 - The difference in the daily number of deaths between intervention (heat wave) and non-intervention (non-heat wave) days in the post-2004 period, had the HAP not been implemented



ANALYSES

- Quasi-Poisson Model to estimate a number of 'prevented' daily deaths during heat waves after the policy implementation.
- $\log(E(Y_{ct})) = \beta_0 + \beta_1 E_{ct} + \beta_2 I_t + \beta_3 E_{ct} I_t + f(confounders_{ct}) + offset_{ct}$.
 - E_{ct} to be an indicator variable taking the value of 1 if day t in community c (here Montreal) is an eligible day (i.e., exceeds the community's threshold for activating its HAP)
 - I_t to be an indicator variable taking the value of 1 if day t is post-HAP implementation and the value 0 otherwise
 - β_3 represents our coefficient of interest (DID estimate), capturing whether the HAP affected daily mortality after its implementation
- Cumulative heat effect (lag 0-5) and harvesting effect
- Sensitivity analysis: Defining the policy implementation at arbitrary ('fake') policy implementation periods (2000 and 2002)
- Equity in the causal effect
 - To assess heterogeneity in the policy causal effect, we calculated differences-indifferences-in-differences (DIDID) estimates



DID ASSUMPTIONS

- No time trends in daily mortality among non-eligible days
- Among non-eligible days (less than 30°C), daily mortality did not change before and after 2004
- Short interval of time (4 years before and 4 years after the initiation of the HAP intervention), to limit confounding due to population adaptation and urban changes



RESULTS

- Main effect: 2.52 deaths per day (95% CI: -0.34, 5.38)
- Represents ~50% of deaths attributable to HWs (using the same definition)

Table 2. Estimated effect of the heat action plan program on equity.

| Potential modifiers of the program benefits | Heterogeneity in the program effect ^a estimate | 95% CI | <i>p</i> -Value ^b |
|---|---|---------------|------------------------------|
| Sex (men vs. women) | 1.38 | (-1.60, 4.36) | 0.36 |
| Age (≥ 65 vs. < 65 years) | 2.44 | (0.27, 4.59) | 0.03 |
| Neighborhood SES (lowest SES tercile vs. highest SES tercile) | 2.48 | (0.69, 4.27) | < 0.01 |

^aFrom DIDID (differences-in-differences-in-differences) estimates (Poisson model adjusted for temporal trends); 95% CIs were obtained by bootstrapping (1,000 samples).



 $^{{}^{}b}p$ -Values are obtained from a Wald test on the interaction term (i.e., DID estimate considering as health outcome the daily difference between two groups).

SENSITIVITY ANALYSES

Table 3. Sensitivity analyses for the estimated effects of the heat action plan program.

| Sensitivity analyses | DID estimate | 95% CI | <i>p</i> -Value ^a |
|--|--------------|---------------|------------------------------|
| Arbitrary programs | | | |
| Program implemented in 2000 ^b | 0.94 | (-2.08, 3.96) | 0.54 |
| Program implemented in 2002 ^c | 0.42 | (-3.62, 2.77) | 0.80 |
| Other hot days definitions | | | |
| When maximum temperature is above 28°C | 0.58 | (-1.77, 2.93) | 0.63 |
| When maximum temperature is above 32°C | 2.79 | (-2.65, 8.23) | 0.32 |
| Cumulative heat ^d | 4.87 | (0.67, 8.20) | 0.03 |
| Accounting for displacement ratio ^e | 1.87 | (0.29, 3.47) | 0.02 |
| Restriction to non-eligible days above 25°C | 2.23 | (-0.80, 5.27) | 0.15 |

^ap-Values are obtained from a Wald test on the interaction term (i.e., DID estimate).



^bUsing mortality and temperature data for periods 1996–1999 vs. 2000–2003.

^cUsing mortality and temperature data for periods 1998–2001 vs. 2002–2005.

^dConsidering a cumulative heat effect up to 5 consecutive hot days (lag 0-5).

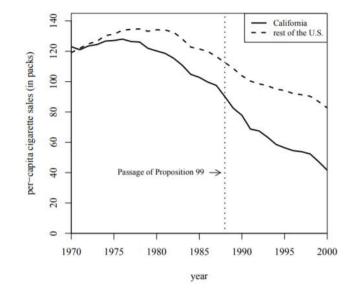
^eThe displacement ratio (Saha et al. 2014) was 0.65.

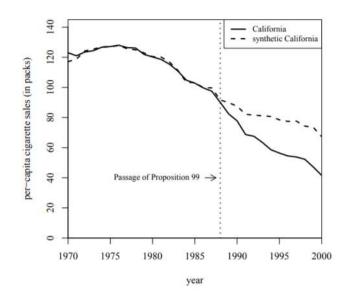
SYNTHETIC CONTROL METHODS



THE INTUITION

- When using a DID, it is 'sometimes' difficult to establish whether the parallel trends assumption is met and whether the control group is a sufficiently accurate representation of what would have happened in the treated area without the intervention
- Synthetic control methodology (SCM) allows the construction of a counterfactual by selecting a <u>weighted average</u> of the outcome variable from a group of units similar to the treated unit
- The intuition behind this method using the original paper by Abadie et al. 2010







HOW DOES IT WORK?

- The synthetic control is based on the vector of weights **W** that minimizes the imbalance between the treated unit and a weighted average of the controls across a set of variables **X** (e.g. pre-intervention outcomes and/or covariates), $(X_1 X_0 W)'V(X_1 X_0 W)$
 - X₁ and X₀ contain the pre-treatment outcomes and covariates for the treated unit and control units respectively, and V captures the relative importance of these variables as predictors of the outcome of interest.
- In this setting, we assume the weights W to be positive and summing to 1 to avoid extrapolations issues (Abadie et al 2010). More recent approaches relaxed this assumption
- The treatment effect for the treated unit (i = 1), τ_{1t} , can then be estimated by $(Y_{1t} \hat{Y}_{1t}^0)$ for each post-intervention period separately, and these can be averaged over time to obtain an ATT over the post-intervention period
- This is a non-parametric approach, but statistical inference can be obtained via permutation tests
- What happens when we have multiple treated units?
 - Discussed later ...



STEPS IN CONDUCTING A SYNTHETIC CONTROL STUDY

- 1. Ensure the theory behind the intervention is well understood. Develop or present a conceptual model to make the theory transparent.
 - To ensure areas that have also been exposed to a similar intervention are excluded from the pool of potential controls
- Identify potential control units that are plausibly eligible
- 3. Develop the synthetic control.
 - An optimization procedure using the outcome variables from the potential control areas to select the best weighting of units from the donor pool to create a synthetic control
- 4. Run outcome analysis and present results
- 5. Run robustness checks (discussed later)



KEY ASSUMPTIONS

Synthetic control methodology as a tool for evaluating population-level health interventions

Janet Bouttell, Peter Craig, James Lewsey, Mark Robinson, Frank Popham²

| Table 2 Key assumptions of synthetic control methodology | | | | | |
|---|---|--|--|--|--|
| Assumption | Assessment | | | | |
| Treated units and potential control units in the donor pool are similar. | Similar levels in variables known to influence outcome variable (see box 1 for objective and subjective elements of this assessment). | | | | |
| 2. There is no contamination — spillover of effects of intervention into potential control units. | Based on background knowledge of researchers. | | | | |
| 3. No external shocks in potential control units. | Based on background knowledge of researchers informed by review of trends in outcome variable. | | | | |



EXAMPLES

SEX WORK REGULATION AND SEXUALLY TRANSMITTED INFECTIONS IN TIJUANA, MEXICO

TROY QUASTa.* and FIDEL GONZALEZb

^aHealth Policy and Management, College of Public Health, University of South Florida, Tampa, FL, USA
^bEconomics and International Business, Sam Houston State University, Huntsville, TX, USA

A New Tool for Case Studies in Epidemiology—the Synthetic Control Method

David H. Rehkopf, and Sanjay Basub

Do medical marijuana laws reduce addictions and deaths related to pain killers?*

David Powell a.*, Rosalie Liccardo Pacula a.b, Mireille Jacobson b.c

- 2 RAND, Santa Monica, United States
- b NBER, Cambridge, MA, United States
- CUniversity of California, Irvine, United States

Original Contribution

Health Behaviors, Mental Health, and Health Care Utilization Among Single Mothers After Welfare Reforms in the 1990s

Sanjay Basu*, David H. Rehkopf, Arjumand Siddiqi, M. Maria Glymour, and Ichiro Kawachi

Effects of changes in permit-to-purchase handgun laws in Connecticut and Missouri on suicide rates



Cassandra K. Crifasi *, John Speed Meyers, Jon S. Vernick, Daniel W. Webster

Johns Hopkins Center for Gun Policy and Research, Department of Health Policy and Management, Johns Hopkins Bloomberg School of Public Health, Baltimore, MD, United States



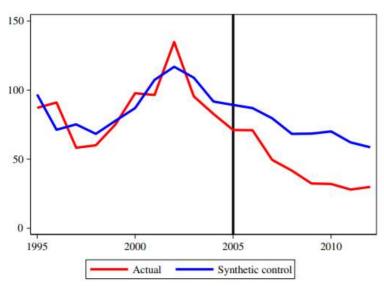
SEX WORK REGULATION AND SEXUALLY TRANSMITTED INFECTIONS IN TIJUANA, MEXICO

TROY QUAST^{a,*} and FIDEL GONZALEZ^b

*Health Policy and Management, College of Public Health, University of South Florida, Tampa, FL, USA
*Economics and International Business, Sam Houston State University, Huntsville, TX, USA

ABSTRACT

While reducing the transmission of sexually transmitted infections is a common argument for regulating sex work, relatively little empirical evidence is available regarding the effectiveness of these policies. We investigate the effects of highly publicized sex work regulations introduced in 2005 in Tijuana, Mexico on the incidence of trichomoniasis. State-level, annual data for the 1995–2012 period are employed that include the incidence rates of trichomoniasis by age group and predictor variables. We find that the regulations led to a decrease in the incidence rate of trichomoniasis. Specifically, while our estimates are somewhat noisy, the all-ages incidence rate in the 2005–2012 period is roughly 37% lower than what is predicted by our synthetic control estimates and corresponds to approximately 800 fewer reported cases of trichomoniasis per year. We find that the decreases are especially pronounced for 15–24 and 25–44 age cohorts. Copyright © 2016 John Wiley & Sons, Ltd.

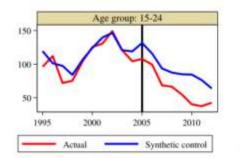


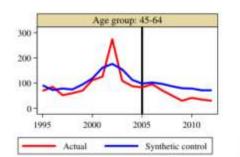
Note: the synthetic control includes Chihuahua (67.0%), Distrito Federal (18.3%), and México (13.6%).

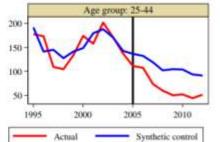
Synthetic control Weights

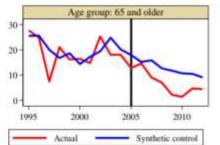
Table II. Synthetic control weights

| | All | 15-24 years old | 25-44 years old | 45-64 years old | 65+ years old |
|------------------|-------|--------------------|--------------------|--------------------|------------------|
| State | | | | | |
| Campeche | 0.000 | 0.000 | 0.000 | 0.000 | 0.000 |
| Chihoahua | 0.670 | 0.197 | 0.587 | 0.866 | 0.608 |
| Distrito Federal | 0.183 | 0.000 | 0.397 | 0.000 | 0.000 |
| Jalisco | 0.000 | 0.654 | 0.000 | 0.000 | 0.000 |
| México | 0.136 | 0.149 | 0.000 | 0.020 | 0.371 |
| Puebla | 0.000 | 0.000 | 0.016 | 0.000 | 0.000 |
| Sonora | 0.000 | 0.000 | 0.000 | 0.000 | 0.000 |
| Tabasco | 0.000 | 0.000 | 0.000 | 0.000 | 0.022 |
| Tamuulipas | 0.000 | 0.000 | 0.000 | 0.114 | 0.000 |
| Tlaxcala | 0.000 | 0.000 | 0.000 | 0.000 | 0.000 |
| Veracruz | 0.000 | 0.000 | 0.000 | 0.000 | 0.000 |
| | | | | | |











OVERVIEW:

COMPARING DIFFERENT APPROACHES TO EVALUATE NATURAL EXPERIMENTS BASED ON THE TIMING OF THE INTERVENTION

- Uncontrolled before and after designs
 - Single or multiple measurements
- Interrupted time series design
- Controlled Before and after with single measurements
- Difference-in-Differences
- Augmented DID approaches
 - With propensity score methods
- Synthetic Control Methods

No Control Group

With one Control Group

With >1 Control Groups



MULTIPLE TREATED GROUPS

- When dealing with multiple treated units, there are two distinct settings:
 - All treated units received the intervention of interest at the same time
 - Covered today
 - Treated units received the intervention at different times
 - Not covered today
- When multiple units receive the intervention at the same time:
 - A simple approach is to estimate a separate ATT for each treated unit and then conduct a meta-analysis to get a pooled estimate (and also information about heterogeneity across units)
 - Or aggregate the treated units and form a synthetic control for the aggregated treated unit (Acemoglu et al. 2013; Dube and Zipperer, 2015)



EXTENSIONS OF TRADITIONAL SC: GENERALIZED SYNTHETIC CONTROL (GSC)

Xu et al. 2017 proposed generalized synthetic control (**GSC**): estimates the average treatment effect on the treated using time-series cross sectional data

Improves efficiency and interpretability from SC, and can be used with multiple treated units and time varying confounders

GSC overcomes limitations in SC including:

- Only handles one treated unit at a time
- No formal measures of variance in traditional SC



HOW DOES GSC WORK?

- Generalized synthetic control (GSC) methods estimate the average treatment effect on the treated (ATT) using time-series cross sectional data
- Well suited for:
 - Time varying confounding: temperature, other weather events
 - Widespread exposure multiple exposed units
- The intuition behind this approach

$$Y_{it} = \delta_{it} D_{it} + x'_{it} \beta + \lambda'_{i} f_{t} + e_{it}$$

- ■D_{it} treatment indicator
- ■x_{it} observed covariates
- ■f, latent factors
- λ_i factor loadings

$$Y_{it}^{0} = x_{it}'\beta + \lambda_{i}'f_{t} + e_{it}$$

$$Y_{it}^{1} = \delta_{it} + x_{it}'\beta + \lambda_{i}'f_{t} + e_{it}$$

$$\widehat{\Delta}_{it} = Y_{it}^1 - \widehat{Y}_{it}^0$$



CASE STUDY: 2007 SOUTHERN CALIFORNIA WILDFIRES ON RESPIRATORY HOSPITALIZATIONS [SHERIDAN ET AL.]

As the climate changes, wildfires are expected to increase in frequency, intensity and duration (especially in California)

In this case study we use:

- Satellite based smoke plume data and burn area data to classify wildfire exposure
- OSHPD respiratory hospitalization data by zip code for outcome
- We capitalize on the timing of the wildfire event



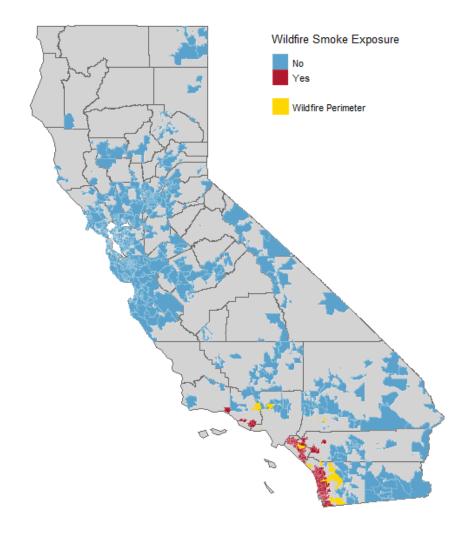


RESULTS

Out of 1779 zip code tabulation areas in California

685 were included:

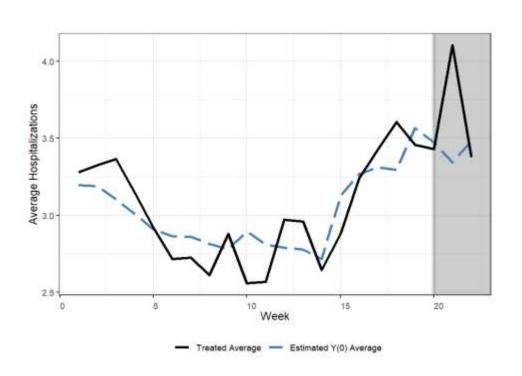
- 130 exposed
- 555 unexposed

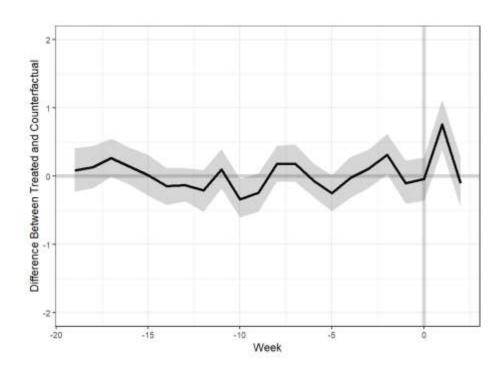


Map of California zip codes exposed to wildfire smoke week of October 20th 2007.



RESULTS





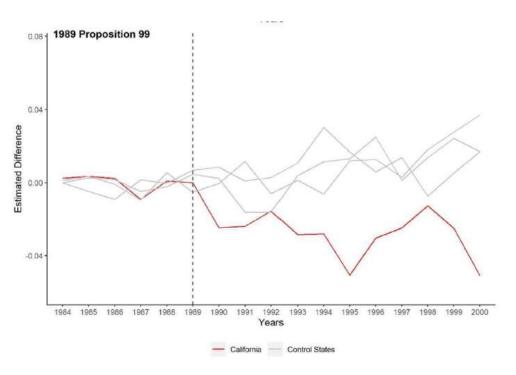
We found an 18% (95%CI: 10%-29%) average increase in respiratory hospitalizations as a result of the wildfire storm



THE IMPORTANCE OF FALSIFICATION TESTS

Permutation tests

- Many assumptions cannot be checked empirically
 - It is therefore important to design a set of falsification/placebo tests to improve the inference of interest
- Negative control approaches
 - Lipsitch M et al. (2012): Negative controls: a tool for detecting confounding and bias in observational studies.



Sheridan et al. "Evaluating the impact of the California 1995 smoke-free workplace law on population smoking prevalence using a synthetic control method." Preventive medicine reports 19 (2020): 101164.



TOPICS NOT DISCUSSED TODAY

- Alternative approaches
 - E.g. Augmented Synthetic Control
 - Ben-Michael, E., Feller, A., & Rothstein, J. (2021). The augmented synthetic control method. Journal of the American Statistical Association, 116(536), 1789-1803.
- Multiple treated units at different times (staggered DID/SCM)
 - A very active area of research
 - Goodman-Bacon (2018) proposed a solution based on a weighted average of all possible DID estimators (using some groups multiple times) in the sample of interest. This approach requires an additional identifying assumption of time-invariant treatment effects.
 - Callaway and Sant'Anna (2020) proposed an analytical solution for such case where there are more than two time periods and units that can become treated at different points in time while relaxing the time-invariant treatment effects assumption
- Multi-stage interventions
- The Scale of interest
 - Important to think about the scale of the interaction term
 - When Is Parallel Trends Sensitive to Functional Form? (Roth and Sant'Anna, 2021)
- Lagged/delayed effects
 - Many available tools



IN SUMMARY DESIGNING A STUDY CAPITALIZING ON THE TIMING OF AN INTERVENTION

When designing a study in which the timing of an intervention is capitalized on, we recommend to undertake the three following steps

- 1. Define a clear research question, ideally with a well-defined intervention with specific implementation time and targeted population;
 - √ The target trial framework which considers the design of an ideal randomized experiment, can be helpful in framing the study and the target causal estimand of interest.
- 2. Propose an **identification strategy** about how we can exploit the setting of the intervention implementation of interest to estimate the targeted causal quantity of interest (estimand) and identify possible falsifications tests
- 3. Propose an **estimation strategy** which often relates to the statistical or modelling approach (i.e. estimator) that we will use to obtain a given estimate (e.g. regression coefficient) to quantitatively represent the targeted estimand of interest (i.e. ATT).



REVISITING THE WORKSHOP OBJECTIVES

- Understand the conceptual foundations of quasi-experimental methods based on the timing of a natural experiment
- 2. Understanding how Difference-in-Differences (DID), Interrupted Time Series (ITS) and Synthetic Control Methods (SCM) work and their specific identification assumptions.
- 3. Being able to implement (in R) DID, ITS and SCM in different settings, interpret estimands of interest and check/visualize possible assumptions' violations
 - √ Coming next



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THANK YOU

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