

What Causes Cholera? Hugely Important in 1850s London



Horrendous way to die - dehydration, convulsions, blue skin, die within hours

Scourge of mid-1800s London – 1831-32 6,526 dead; 1849 14,137; 1853-54 10,738 Massive uncertainty as to cause

• Bad air (miasma); bad breeding (poverty); bad ground (plague pits)

Huge public health & policy question – and one man knew the answer:

• John Snow & bad water – effort to prove contaminated water as causal agent

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Why John Snow and 1850s Cholera?

This work (joint with Peter Vinten-Johansen) is to re-examine and reconsider Snow's analysis of South London

Why Snow? Three reasons:

- **1** Rollicking Good Tale full of heroism, death, and statistics
- 2 **Causal Inference** template for how to marshal evidence in support of a causal explanation
- **3 Statistics & Instruction** The data are simple but the analysis demonstrates multiple data analytic tools we use today
 - Snow cited as first instance of both Difference-in-Differences (Angrist & Pischke) and IV / randomization (Green 2018)
 - combining maps and data (GIS or geographic information systems)
 - regression and error analysis

Snow's cholera work is also a humbling reminder of the sometimes meandering path towards truth: even with overwhelming evidence and strong analysis Snow failed to convince the medical establishment, the public, or the authorities

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Broadly, Three Strands or "Blocks" of Evidence

Snow marshaled evidence in 1855 & 1856 to convince skeptics

- 1 Isolated events, e.g. Albion Terrace
 - 1849, Discovery of waterborne theory
 - single event, 17 houses
- 2 Broad Street Outbreak
 - Aug-Sep 1854, 700 deaths over roughly 2 weeks, 10 square blocks
- 3 South London "Grand Experiment"
 - Summer & Fall 1854, customers supplied by two water companies
 - large scale, 400k mixed (quasi-random) subjects

Data or Evidence Blocks



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Preview: Hypotheses, Snow's Analysis, Modern Re-Analysis

Hypothesis	Challenge & Alternatives	Snow's Analysis	Problems	Modern Approach
Water Causal	Challenge – Confound Alternatives – Elevation – Crowding – Social class	1855 Table IX – Direct comparison of mixed populations (quasi-randomized trial)	 1855: all subdistricts together (no detailed population-by- supplier) 1856: unbalanced 	 Randomization limited to jointly-supplied (16) subdistricts Regression & subdistrict FE control for balance Count regression
	– Miasma – others	1855 p. 89 (Table XII) – Difference-in- Differences	– 1855: marginal significance (substantial heterogeneity)	Combine DiD & Prediction – DiD with continuous treatment effect
Water Most Im- portant	Challenges – Confound – "How important?" Alternatives – As above	1856 Table VI – Prediction from supplier population fractions	 Only 2 suppliers No way to measure quality of fit Ignores change 1849 to 1854 	 (population fractions) 3 suppliers DiD controls for confounders Residual Deviance & <i>R</i>² measures % explained Count regression

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Snow Reconsidered

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Outline

- Overview: John Snow and the Story of Cholera Cholera, John Snow, and Waterborne Theory Data, Timeline, and Locations
- 2 John Snow's Evidence & Causal Inference
- 3 Albion Terrace "Discovery" of Theory
- 4 Broad Street Pump Famous for "The Map"
- 5 South London "Grand Experiment"

Overview

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Cholera – Disease of Poor Sanitation

What is Cholera?

- Vibrio Cholerae bacterium that infects the small intestine of humans
- Causes severe diarrhea (& vomiting) that drains fluids
- Death from dehydration & organ failure
- Oral Rehydration Therapy highly succesfull (roughly 1960s)
 - In case you ever need it, here's the recipe 1 liter boiled water, 1/2 teaspoon salt, 6 teaspoons sugar, mashed banana (potassium)

Cholera thrives in crowded cities with poor sanitation

- Transmitted through recycling (drinking) sewage
- When cholera exits one victim, needs to find a way into gut of others
- Victorian London was an ideal playground for cholera to thrive

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Cholera Loved Victorian London

Victorian London was an ideal playground for cholera

- Mid-1800s London was dirty, smelly place with no organized sewage treatment
- Efforts to improve sanitation made things worse
 - cesspools relatively safe did not provide access to thousands of guts
- Public Health Act of 1848 accelerated the connection of houses to sewage lines
 - helped clean up streets, flushed filth to Thames
- By mid-1800s, cholera had easy access from the gut of one to thousands of victims

Contemporaries were aware of dirty water (Punch 1849)

• But water not recognized as vector for cholera



Solution - Construction of Bazalgette "Outfall Sewers"

Sewers that sloped towards outfalls (discharge points) lower on the Thames

- Construction started (under Bazalgette) 1859, response to 1858 "Great Stink"
- Embankments along Thames what we see today
 - Embedded discharge pipes still used today (?)
 - Decreased width, increased flow scouring effect
- Moved sewage downstream, below London & water in-take



One final outbreak, 1866, limited to east London, last area unserved by sewers $_{\mathcal{OQ}}$

John Snow's Research & Publications

Doctor - pioneer in anesthesia & medical hygiene

• Provided Queen Victoria with anesthesia during childbirth

Research and writing on Cholera

- 1849: "On the Mode of Communication of Cholera"
 - Laid out theory and evidence for waterborne transmission
- 1855: "On the Mode of Communication of Cholera"
 - Substantially expanded, additional evidence and argument
- 1856: "Cholera and the water supply in the south district of London in 1854"
 - Refined randomized analysis

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John Snow's 1849 Theory & 1855 Evidence

1849: Snow developed theory of infection & transmission

- Based on medical knowledge and study of single events
 - Horsleydown & Albion Terrace

Fully-developed & modern theory of disease

- Infects & reproduces in the small intestine
- Exits from victim, into water supply
- Infects new victims through drinking dirty water

Implications for patterns of infection, across scales

• "from the membrane of the small intestine all the way up to the city itself" (Johnson)

Snow's work grounded by theory

Snow had a good idea – a causal theory about how the disease spread – that guided the gathering and assessment of evidence. (Tufte)

1855: evidence & argument to convince skeptics



Alternative Theories

Miasma (Smells & Airborne)

- Cholera infectious & transmitted through the air
- Generally accepted in mid-1800s

Elevation, Crowding & Class, Others

- \bullet Elevation: lower elevation \rightarrow more infection
- Crowding & Class: lower class & crowding \rightarrow more infection

None of these absolutely crazy - correlated with cholera (and dirty water)

- Raw sewage associated with bad smells & dirty drinking water
- Lower class associated with crowding & poor sanitation

Other non-infectious theories (I won't seriously consider)

- Emanations from the ground
- Plague burying-pit near Broad Street pump

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1 Overview: John Snow and the Story of Cholera

Cholera, John Snow, and Waterborne Theory

Data, Timeline, and Locations

- 2 John Snow's Evidence & Causal Inference
- 3 Albion Terrace "Discovery" of Theory
- 4 Broad Street Pump Famous for "The Map"
- 5 South London "Grand Experiment"
 - Overview
 - Snow's Analysis
 - Modern Analysis
- 6 Conclusion

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Overview: John Snow and the Story of Cholera Cholera, John Snow, and Waterborne Theory Data, Timeline, and Locations

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Modify Katz & Singer as "Causal Assessment Procedure"

Still tentative, based on Katz & Singer's analysis of possible Chemical & Biological Weapons attacks, 1970s-80s, "Can an Attribution Assessment Be Made for Yellow Rain?"

- 1 Divide evidence into blocks or types of evidence
- 2 Assign to each block a veritas rating quality of data
- 3 Develop groups of hypotheses
- 4 Assess each evidence block for strength of rejection for each hypothesis
 - Consider *rejection* of hypotheses (refute, neutral, consistent) rather than strength of association (support of hypotheses)
- 5 Organize evidence blocks by hypothesis into matrix
- 6 Choose hypothesis not contradicted
- 7 Strongest hypothesis checked

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Theory, Data, Hypothesis Testing

Data or Evidence Blocks Albion Terr Broad St South London summer/fall 1854 17 houses ~10 sa blocks ~400k subjects mixed single outbreak 2wks, 700 deaths treated & untreated Theory & Hypotheses theory: water & small intestine H2: water primary H1: water a causal factor causal factor

alternatives: elevation, crowding, class, ...

Hypothesis or Testing Blocks



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"Grand Experiment" – Water Supply Changes

Two water companies served south London – Southwark & Vauxhall Co and Lambeth Co. - 486,936 customers, 300,000 intimately mixed

In 1830s & 1840s companies competed for customers, often on same street

In many cases a single house has a supply different from that on either side. Each company supplies both rich and poor, both large houses and small; there is no difference in the condition or occupation of the persons receiving the water of the different companies. (Snow 1855 p 75)

1849 epidemic

- Both companies drew water from low in the Thames near Vauxhall bridge 1852
 - Lambeth Company moved source to Thames Ditton (upstream of London)
 - In response to Act of Parliament, requiring move (by 1855)

1854 epidemic

- Southwark & Vauxhall Co supplied dirty water
- Lambeth Co supplied cleaner water

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32 Subdistricts, 12 S&V only, 16 joint, 4 Lambeth

Registration Districts & Sub-Districts – Need to keep straight

- Deaths collected weekly by Registrar-General, by District & Subdistrict
- In this region of South London, 32 sub-districts
- Snow viewed "mixed" as crucial experiment



- "First 12" Southwark & Vauxhall Water Co only dirty water 1849 & 1854
- "Next 16" Mixed or Joint Southwark & Vauxhall Co and Lambeth Water Co 1849 dirty water, 1854 part dirty (S&V) & part clean (Lambeth)
- "Final 4" Lambeth Water Co only not relevant, not supplied in 1849

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Deaths: Combined (All Suppliers) vs Direct (By Supplier)

Data available in 1855

- Deaths (combined all suppliers) 1849 & 1854, full epidemic
- Population (combined all suppliers)
- Deaths by supplier, first 7 weeks of epidemic (collected by Snow)

Data available in 1856 (originally published by Simon)

• Population by supplier (only S&V shown here)

						1854, first 7 wks		
	subdistricts	Deaths 1849	Deaths 1854	Supplier	Population 1851	Deaths S&V	Deaths Lam	Pop S&V
1	St. Saviour	283	371	sv	19,709	115	0	16,337
2	St. Olave	157	161	sv	8,015	43	0	8,745
13	Christchurch	256	113	SV & Lambeth	16,022	11	13	2,915
14	Kent Road	267	174	SV & Lambeth	18,126	52	5	12,630
29	Norwood	2	10	Lambeth	3,977	0	2	0
	TOTAL	6,328	5,042		486,936	1,263	98	266,516

Combined (all suppliers)Direct (by supplier) $D_{subdist} = D_{S\&V} + D_{Lam} + D_{Other}$ $\{D_{S\&V}, D_{Lam}, D_{Other}\}$

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Snow's Analysis – 3 Approaches

Mixed or quasi-random direct comparison

- Snow determined supplier by bill or chloride test
- Visited all houses (deaths) for 7 weeks ending Aug 26



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Diff-in-Diffs comparison of combined (all suppliers) mortality rates

- For each subdistrict, observe combined deaths all suppliers
- Compare 1849 vs 1854 and Treated (clean) vs untreated (dirty) subdistricts

Actual v Predicted across subdistricts for 1854 only

• Predicting mortality based on fraction of S&V vs Lambeth customers

Snow's Analysis – 3 Approaches

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Water Most Im- portant	Challenges 1 – Confound – – "How s important?" f Alternatives – As above	1856 Table VI – Prediction from supplier population fractions	 Only 2 suppliers No way to measure quality of fit Ignores change 1849 to 1854 	 (population fractions) – 3 suppliers – DiD controls for confounders – Residual Deviance & R² measures % explained – Count regression

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Snow Modern in View of Mixing (Randomization)

Recognized that mixing (randomization) would average out differences

As there is no difference whatever, either in the houses or the people receiving the supply of the two Water Companies, or in any of the physical conditions with which they are surrounded, it is obvious that no experiment could have been devised which would' more thoroughly test the effect of water supply on the progress of cholera than this. (1855 p. 75)

Cited as first instance of Randomization and Instrumental Variables (Greene 2018, also Deaton, others)

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Direct Comparison of Mixed or Randomized Population

Table: Houses, Deaths, and Mortality Rates per 10,000 Households, First Seven Weeks of 1854 Cholera Epidemic – Table IX

Water Supplier	Number of houses	Deaths from Cholera	Deaths in each 10,000 houses
Southwark & Vauxhall Co supply	40,046	1,263	315.4
Lambeth Co supply	26,107	98	37.5
Rest of London	256,423	1,422	59
Ratio Effect: Southwark & Vauxhall vs Lambeth			8.40

Note that this corrects a rounding error in the "Deaths in each 10,000 houses" for Lambeth in Snow's original table

- Found LARGE Lambeth effect
- But suffered from potential confounding includes all subdistricts

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Problem: Direct Comparison Uses All Subdistricts



- Snow wanted to use Joint (Mixed) subdistricts
- Population (houses) by supplier for overall region only
- Potential for confounding (bias if S&V-only subdistricts different than joint)

Second Approach – Diff-in-Diffs

Hypothesis	Challenge & Alternatives	Snow's Analysis	Problems	Modern Approach	
Water Causal	Challenge – Confound Alternatives – Elevation – Crowding – Social class	1855 Table IX – Direct comparison of mixed populations (quasi-randomized trial)	– 1855: all subdistricts together (no detailed population-by- supplier) – 1856: unbalanced	 Randomization limited to jointly-supplied (16) subdistricts Regression & subdistrict FE control for balance Count regression 	
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Simple Diff-in-Diffs – Before vs After, Treated vs Control

Table: Mortality Rates 1849 & 1854, Summary Snow 1855 Table XII

	1849	1854	Ratio
	Deaths	Deaths	1849 -
	per 10,000	per 10,000	1854
Always Dirty – Southwark & Vauxhall Water	134.9	146.6	0.92
Company Only ("First 12" subdistricts)	only	only	time
Dirty / Clean – Joint Southwark & Vauxhall and Lambeth Companies ("Next 16" subdistricts)	130.1 dirty, joint	84.9 (partial) clean	1.53 diff in time & treatment
Ratio: Next 16 less First 12	0.96 diff in region	1.73 diff in region & treatment	1.67 (partial) treatment

Comparing the S&V-only subdistricts vs the Jointly-supplied subdistricts

- Interestingly, Snow did not convert deaths to rates missed an opportunity
- Large treatment effect, but need to evaluate statistical significance

Problem: treatment effect only marginally significant ...

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Snow's Analysis

DiD as Regression

$$\begin{aligned} &\ln\left(\textit{Rate}_{\textit{subdist},yr}\right) = \ln\left(\textit{count}_{\textit{subdist},yr}/\textit{population}_{\textit{subdist},yr}\right) = \hat{\mu} + \hat{\delta}_{54} \cdot I_{yr=1854} \\ &+ \hat{\gamma}_J \cdot I_{\textit{subdist}=joint} + \hat{\beta} \cdot I_{\textit{subdist}=joint} \cdot I_{yr=1854} + \varepsilon_{s,y} \end{aligned}$$

Region or Sub-Districts – Supplied by	1849 Death Rate (log)	1854 Death Rate (log)	Diff 1854 less 1849
First 12 – Southwark Only	μ	$\mu + \delta_{\bf 54}$	δ_{54}
Next 16 – Joint Southwark and Lambeth	$\mu + \gamma_J$	$\mu + \delta_{54} + \beta + \gamma_J$	$\delta_{54}+eta$
Diff Joint less Southwark	γ_J	$\beta + \gamma$ J	β

Regression framework allows us to

- Use subdistrict detail, and additional regressors (if available)
- Test for statistical significance (both for finite population and "within-sample" variation)
- Extend the DiD framework to continuous treatment and actual-vs-predicted

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Snow Highlighted Difference in "Lambeth Degree"



• Four subdistricts where "the supply of the Lambeth Water Company is more general than elsewhere"

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Snow Highlighted Difference in "Lambeth Degree"

	1849	1854	Ratio
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	per 10,000	per 10,000	1854
Always Dirty - Southwark & Vauxball Water	134.9	146.6	0.92
Company Only ("First 12" subdistricts)	dirty, S&V	dirty, S&V	diff in
company only (That 12 Subdistricts)	only	only	time
Dirty / Clean - "More Lambeth" in Joint (4	138.8	47.2	2.94
cubdictricte)	dirty more	more	time &
subdistricts)	unity, more	clean	more
Dirty / Clean - "Less Lambeth" in Joint (12	127.6	05.6	1.34
subdistricts)	dirty loss	less clean	time &
subdistricts	unity, less	less clean	less
	0.97	3.11	3.20
Ratio: "More Lambeth" vs Dirty	diff in	region &	more
	region	more	treatment
	1.06	1.53	1.45
Ratio: "Less Lambeth" vs Dirty	diff in	region &	less
	region	less	treatment

Larger effect for "More Lambeth"

• Now, treatment effect is highly significant (see below)

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Third Approach - in 1856, Actual-vs-Predicted

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Actual-vs-Predicted by Subdistrict

	1	2	3	4	6	10
Subdistrict	Рор 1851	Popul Estin by Su South- wark	ation nates pplier Lambet	h Both Cos	Actual Rate	Predicted Rate
Christchurch, Southwark	16.022	2.915	13.234	16,149	70.5	51.0
St. Saviour, Southwark	19,709	16,337	898	17,235	191.8	153.1
St. Olave, Southwark	8,015	8,745	0	8,745	200.9	160.0
St. John, Horsleydown	11,360	9,360	0	9,360	133.8	160.0
St. James, Bermondsey	18,899	23,173	693	23,866	191.5	156.1

$$\hat{R}_{subdis, both} = \frac{N_{subdis, Southwark} \cdot \hat{R}_{Southwark} + N_{subdis, Lambeth} \cdot \hat{R}_{Lambeth}}{N_{subdis, Southwark} + N_{subdis, Lambeth}} = F_S \cdot \hat{R}_S + F_L \cdot \hat{R}_L$$

 $\hat{R}_{Southwark}$, $\hat{R}_{Lambeth}$ Overall rate for supplier (average across all subdistricts) $\hat{R}_{Southwark} = 160$ and $\hat{R}_{Lambeth} = 27$

- Correlation is 0.75 ($R^2 = 56\%$), but Snow had no statistics, could only argue individual cases
- Problems: a) Ignores "Other" (pumps, wells); b) Population not accurate (see St. James)

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Conditional Map – Highlights High S&V Mortality

Vertical: Fraction of population Lambeth

Horizontal: Fraction of population Southwark & Vauxhall

- Fraction Lambeth Low & High: Low mortality (along vertical)
- Fraction S&V Low: Low mortality; Fraction S&V high: High mortality (along horizontal)



Conditional Map – Highlights High S&V Mortality

Vertical: Fraction of population Lambeth

Horizontal: Fraction of population Southwark & Vauxhall

- Fraction Lambeth Low & High: Low mortality (along vertical)
- Fraction S&V Low: Low mortality; Fraction S&V high: High mortality (along horizontal)



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6 Conclusion

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Overview - Problems and Modern Re-Analysis

Hypothesis	Challenge & Alternatives	Snow's Analysis	Problems	Modern Approach
Water Causal	Challenge – Confound Alternatives – Elevation – Crowding – Social class	1855 Table IX – Direct comparison of mixed populations (quasi-randomized trial)	– 1855: all subdistricts together (no detailed population-by- supplier) – 1856: unbalanced	 Randomization limited to jointly-supplied (16) subdistricts Regression & subdistrict FE control for balance Count regression
	– Miasma – others	1855 p. 89 (Table XII) – Difference-in- Differences	– 1855: marginal significance (substantial heterogeneity)	Combine DiD & Prediction – DiD with continuous treatment effect
Water Most Im- portant	Challenges – Confound – "How important?" Alternatives – As above	1856 Table VI – Prediction from supplier population fractions	 Only 2 suppliers No way to measure quality of fit Ignores change 1849 to 1854 	 (population fractions) – 3 suppliers – DiD controls for confounders – Residual Deviance & R² measures % explained – Count regression

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Population-by-Supplier & Direct Comparison (Mixed, Randomized)

Publication (in 1856) of population-by-supplier allowed direct comparison of mixed subdistricts

- Snow published the Actual-vs-Predicted, but not re-analysis of direct comparison (quasi-randomized) not sure why
- Re-analysis shows same result large Lambeth effect

Water Supplier	Number of houses	Deaths from Cholera	Deaths in each 10,000 houses	Approx Deaths per 10,000 persons		Population by Sup- plier	Deaths	Deaths per 10,000 persons
Southwark & Vauxhall Co supply	40,046	1,263	315.4	100.9	S&V only supply, "First 12"	119,603	738	132.2
					S&V only supply, "Next 16" joint	118,888	525	94.6
Lambeth Co supply	26,107	98	37.54	12.01	Lambeth supply, ''Next 16'' joint	147,961	94	13.6
Ratio Effect: S&V vs Lambeth			8.40	8.40	Ratio Effect, joint subdistricts			6.95
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Randomization & Subdistrict Confounding

Still does not rule out confounding, because subdistricts vary and randomization is within but not across subdistricts

- Hypothetical Example: subregion 2 has low mortality and high proportion of Lambeth customers, produces overall low Lambeth mortality
- But the true effect of Lambeth (vs S&V) is small

	Southv Pop	vark & Va Deaths	u×hall Mortality Rate	/ Рор	Lambeth Deaths	Mortalit <u>.</u> Rate	y Ratio
Subregion 1 Subregion 2 Overall (total)	106,999 11,889 118,888	514 11 525	102.90 19.83 94.63	14,796 133,165 147,961	35 59 94	50.69 9.49 13.61	2.03 2.09 6.95
Ratio of Subregions			5.19			5.34	

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Randomization as Averaging Over Confounders

Treatment effect is $\hat{\beta}_l$:

$$\ln\left(\textit{Rate}_{\textit{supplier}}\right) = \hat{\rho}_{\textit{S}} + \hat{\beta}_{\textit{L}} \cdot \textit{I}_{\textit{supplier}=\textit{Lam}} + \varepsilon_{\textit{supplier}}$$

Confounding: different unobservables for S&V vs Lambeth (correlated): $E(\varepsilon_s \mid I_{s=L} = 0) \neq E(\varepsilon_s \mid I_{s=L} = 1) \neq 0$

- Violates standard regression conditions
- Randomization forces $E(\varepsilon_s \mid I_{s=L} = 0) = E(\varepsilon_s \mid I_{s=L} = 1) (\neq 0)$

Now, confounders drop out (the same for both S&V and Lambeth):

$$\mathsf{E}\left(\mathsf{ln}\left(\mathsf{Rate}_{\mathsf{supplier}}
ight) \mid \mathit{I}_{\mathsf{s}=\mathsf{L}}=1
ight) - \mathsf{E}\left(\mathsf{ln}\left(\mathsf{Rate}_{\mathsf{supplier}}
ight) \mid \mathit{I}_{\mathsf{s}=\mathsf{L}}=0
ight) = \hat{eta}_{\mathsf{L}}$$

BUT, averaging is within subdistricts, not across - still potential confounding

Solution? Subdistrict Fixed Effects

Note in passing, randomization can also be viewed as Instrumental Variable (IV) that introduces variation orthogonal to observables and unobservables

See Heckman (1996), Deaton (2018) for randomization as IV

Snow credited (by Greene, others) with first use of randomization as $IV_{(a)} \rightarrow (a) \rightarrow (a$

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Negative Binomial vs Poisson Model

We think about rates, but random variable is *count* (deaths)

 $R_{subdist,supplier} = count_{subdist,supplier} / population_{subdist,supplier}$

First inclination to use Poisson model (counts Poisson)

• But does not fit the data

Explanation is simple

- Each subdistrict is Poisson (must be approximation to sum of Bernoulli alive / dead)
- But random seeding of outbreak and propagation means that actual Poisson rate varies from subdistrict-to-subdistrict

My conclusion?

- Poisson is not the natural starting point
- Subdistrict is Poisson, but drawn from a distribution with some variance (distribution, not fixed number)
- Overall is mixture of Poissons e.g. Negative Binomial as Gamma mixture of Poissons

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Joint 1854 Poisson Act vs Pred, Southwark Supplied

Overdispersion – Fixed Poisson Does Not Fit



Joint 1854 Poisson Act vs Pred, Lambeth Supplied

Mortality per 10,000 Separately for S&V and Lambeth Customers (for the First Seven Weeks, ending 26th August 1854). Actual and Predicted (with 95% confidence bands)

Note how many observations outside 95% bands: 5/32 (15%)

- Formal test: Residual Deviance (if low Resid Dev, high p-value, fail to reject)
- Use Negative Binomial count regression

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Modern Analysis

Summary of Results

Table: Various Fitted Models

	1 Direct Southwark v Lambeth Joint Region	2 Actual v Predicted DiD	3 DiD Single	4 DiD More Lambeth	5 DiD, Actual vs Predicted, Direct Southwark vs Lambeth
	1855 Table VIII	1856 Table VI 1855 Table XII	1855 Ta	able XII	
Raw Treatment	6.95		1.67	3.20	
Estimated Treat.	6.70	3.49	1.65	3.10	5.31
z-ratio	-12.04	-5.23	-2.03	-3.20	-7.08
p-value	2.3E-33	1.7E-07	4.21%	0.14%	1.5E-12
ResDev p-value	15.83%	14.58%	21.45%	15.74%	5.79%
Pseudo RSg	80.2%	69.3%	16.8%	25.1%	85.5%
Rate RSq	70.7%	72.1%	19.9%	24.5%	83.9%
No "Obs"	32	56	56	56	100
No Parameters	3	6	5	7	8
Notes	FE Poisson,				Treat Eff lower
	p-value 2.5%				because 1849
	FE $\hat{\beta}_{I} = 6.83$,				Lam (1.50x);
	z=16				Late TE 3.1x

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Conclusion from Mixed (Quasi-Randomized)

Strongly supports "Water as a causal factor"

- Strong Lambeth (treatment) effect (Lambeth 6.7-times lower, z-ratio 12.0)
- Evidence against confounding
 - Presumption that randomization averages over confounders within
 - Subdistrict Fixed Effects do not change result reject confounding across

Randomization valuable, but randomization alone cannot answer all questions:

- Not strong evidence on "water as primary causal factor"
- No evidence on non-water differences between S&V vs Lambeth customers

For these, turn to DiD and comparison across subdistricts and across time

- Arguably "more difficult" test than mixed (randomized) direct comparison
- Example of the complementary nature of randomized and observational data

DiD Combined – Continuous Treatment

Simple (discrete treatment) DiD:

 $\ln\left(\textit{Rate}_{\textit{subdist},\textit{yr}}\right) = \hat{\mu} + \hat{\delta}_{54} \cdot \textit{I}_{\textit{yr}=1854} + \hat{\gamma}_J \cdot \textit{I}_{\textit{subdist}=\textit{joint}} + \hat{\beta} \cdot \textit{I}_{\textit{subdist}=\textit{joint}} \cdot \textit{I}_{\textit{yr}=1854} + \varepsilon_{\textit{s},\textit{y}}$

Region or Sub-Districts – Supplied by	1849 Death Rate (log)	1854 Death Rate (log)	Diff 1854 less 1849
First 12 – Southwark Only	μ	$\mu + \delta_{\bf 54}$	δ54
Next 16 – Joint Southwark and Lambeth	$\mu + \gamma_J$	$\mu + \delta_{54} + \beta + \gamma_J$	$\delta_{\bf 54}+\beta$
Diff Joint less Southwark	γ_J	$eta+\gamma_J$	β

But the treatment effect can be written as continuous based on fraction Lambeth:

$$\ln\left(\textit{Rate}_{\textit{subdist},\textit{yr}}\right) = \hat{\rho}_{\textit{S}} + \hat{\delta}_{\textit{54}} \cdot \textit{I}_{\textit{yr}=1854} + \hat{\rho}_{\textit{L}} \cdot \textit{F}_{\textit{L}} + \hat{\rho}_{\textit{O}} \cdot \textit{F}_{\textit{O}} + \hat{\beta}_{\textit{L}} \cdot \textit{F}_{\textit{L}} \cdot \textit{I}_{\textit{yr}=1854} + \varepsilon_{\textit{s},\textit{yr}}$$

This now combines DiD and an extended Actual-vs-Predicted

- Act-v-Pred incorporates 3 suppliers: S&V, Lambeth, "Other"
- Underlying mortality rates $(\hat{\rho}_S, ...)$ estimated from data to minimize SSQ
- Compares Act-v-Pred across subdistricts and time, comparing S&V vs Lambeth in 1849

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DiD Continuous Treatment Tests Confounding & Importance

Continuous DiD allows testing for importance of treatment, as well as testing (and controlling for) confounding

$$\ln\left(\textit{Rate}_{\textit{subdist},\textit{yr}}\right) = \hat{\rho}_{\textit{S}} + \hat{\delta}_{\textit{54}} \cdot \textit{I}_{\textit{yr}=1854} + \hat{\rho}_{\textit{L}} \cdot \textit{F}_{\textit{L}} + \hat{\rho}_{\textit{O}} \cdot \textit{F}_{\textit{O}} + \hat{\beta}_{\textit{L}} \cdot \textit{F}_{\textit{L}} \cdot \textit{I}_{\textit{yr}=1854} + \varepsilon_{\textit{s},\textit{y}}$$

- $\hat{\rho}_L$ controls for non-water differences between S&V and Lambeth in 1849 (confounding)
- $\hat{\beta}_L$ measures treatment effect, purged of non-water differences ($\hat{\rho}_L$)
- R^2 (coefficient of determination for rates) measures how well prediction explains observations

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Examining Hypothesis 2: Water as Primary Factor

$$\ln\left(\textit{Rate}_{\textit{subdist},\textit{yr}}\right) = \hat{\rho}_{\textit{S}} + \hat{\delta}_{\textit{54}} \cdot \textit{I}_{\textit{yr}=\textit{1854}} + \hat{\rho}_{\textit{L}} \cdot \textit{F}_{\textit{L}} + \hat{\rho}_{\textit{O}} \cdot \textit{F}_{\textit{O}} + \hat{\beta}_{\textit{L}} \cdot \textit{F}_{\textit{L}} \cdot \textit{I}_{\textit{yr}=\textit{1854}} + \varepsilon_{\textit{s},\textit{y}}$$

Hypothesis:	Base	Alt 1	Alt 2	Alt 3	Alt 4a	Alt 4b
	Water + Treatment Effect	Water No Treatment Effect	No Water, Subdistricts Only	Water + Treatment + Subdistricts	Water + Treatment + Housing Density	Housing Density Alone
Treat ratio	3.49	-	-	3.76	3.49	-
Res Dev p	14.6%	22.4%	0.08%	0.09%	12.3%	25.4%
Rate R ²	72.1%	56.9%	76.1%	89.7%	72.7%	24.5%
BIC	614.5	631.9	692.8	635.0	617.6	665.0
		remove $\hat{\beta}_L$	remove $\hat{\beta}_L, \ \hat{\rho}_S,$ include FE	include FE	include density	density only

Conclusion: water matters, explains large proportion of rate variance, other factors (housing density, all subdistrict fixed characteristics) do not matter

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Review so Far: Support Water, Reject Others

Hypothesis	Challenge & Alternatives	Snow's Analysis	Problems	Modern Approach	
Challenge – Confound Water Alternatives Causal – Elevation – Crowding – Social class		1855 Table IX – Direct comparison of mixed populations (quasi-randomized trial)	 1855: all subdistricts together (no detailed population-by- supplier) 1856: unbalanced 	 Randomization limited to jointly-supplied (16) subdistricts Regression & subdistrict FE control for balance Count regression 	
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Water Most Im- portant	Challenges – Confound – "How important?" Alternatives – As above	1856 Table VI – Prediction from supplier population fractions	 Only 2 suppliers No way to measure quality of fit Ignores change 1849 to 1854 	 (population fractions) 3 suppliers DiD controls for confounders Residual Deviance & R² measures % explained Count regression 	

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Modern Analysis

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ResDev p-value	15.83%	14.58%	21.45%	15.74%	5.79%
Pseudo RSg	80.2%	69.3%	16.8%	25.1%	85.5%
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No "Obs"	32	56	56	56	100
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Notes	FE Poisson,				Treat Eff lower
	p-value 2.5%				because 1849
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Summary

Direct (by supplier) Comparison using $\{D_{S\&V}, D_{Lam}, D_{Other}\}$

- Limited to joint subdistricts, same finding as Snow Table IX
- re Randomization, reject subdistrict confounding
- Strongly support water as a causal factor (not ruling out others)

Actual v Predicted DiD using $D_{subdist} = D_{S\&V} + D_{Lam} + D_{Other}$

- Combines Snow's idea of Actual vs Predicted and DiD
- Arguably tougher test than direct (compare 1849 vs 1854)
- Strongly supports water as *a* causal factor and water as *primary* causal factor Simple DiD

• Using only data from 1855, still strongly supports water as a causal factor

• Statistics: shows importance of *correctly* measuring SEs, and NegBinom over Poisson

Combined & Direct using $D_{subdist} = D_{S\&V} + D_{Lam} + D_{Other}$ and $\{D_{S\&V}, D_{Lam}, D_{Other}\}$

- Reinforces other results

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Overview: John Snow and the Story of Cholera Cholera, John Snow, and Waterborne Theory Data, Timeline, and Locations

2 John Snow's Evidence & Causal Inference

3 Albion Terrace - "Discovery" of Theory

4 Broad Street Pump - Famous for "The Map"

5 South London "Grand Experiment"

Overview

Snow's Analysis

Modern Analysis

6 Conclusion

Supporting and Extending David Freedman's Comments

This detailed analysis of Snow's work supports Freedman's (1991) comments about Snow:

Snow's work is ... a success story for scientific reasoning based on nonexperimental data statistical technique can seldom be an adequate substitute for good design, relevant data, and testing predictions against reality in a variety of settings,

But it modifies Freedman's skepticism about statistical arguments

I do not think that regression can carry much of the burden in a causal argument, [and] Arguments based on statistical significance of coefficients seem generally suspect.

to a more nuanced view: Snow's work proves the importance of marrying good design with good statistical analysis

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Conclusion: Theory, Data, Hypothesis Testing

Data or Evidence Blocks

Albion Terr	Broad St	South London
17 houses single outbreak	~10 sq blocks 2wks, 700 deaths	summer/fall 1854 ~400k subjects mixed treated & untreated

Theory & Hypotheses

water & small	miasma	elevation,
intestine	(airborne)	class,

Hypothesis or Testing Blocks



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Still Much to Learn From John Snow

This work (joint with Peter Vinten-Johansen) is to re-examine and reconsider Snow's analysis of South London

Why Snow? Three reasons:

- **1** Rollicking Good Tale full of heroism, death, and statistics
- 2 **Causal Inference** template for how to marshal evidence in support of a causal explanation
- **3 Statistics & Instruction** The data are simple but the analysis demonstrates multiple data analytic tools we use today
 - Snow cited as first instance of both Difference-in-Differences (Angrist & Pischke) and IV / randomization (Green 2018)
 - combining maps and data (GIS or geographic information systems)
 - regression and error analysis

Snow's cholera work is also a humbling reminder of the sometimes meandering path towards truth: even with overwhelming evidence and strong analysis Snow failed to convince the medical establishment, the public, or the authorities

7 Appendix Tables & Figures

Quantitative Analysis of Maps – Walking Neighborhoods

More Detail for Difference-in-Differences

Raw Mortality Rates for 1849 & 1854

Mortality Rates from Snow Table XII

	Sub Districts	1849 per	1854 per	Water
	Sub-Districts	10,000	10,000	Supplier
1	St. Saviour,	144	188	SouthwarkVauxhall
	Southwark			
2	St. Olave, Southwark	196	201	SouthwarkVauxhall
3	St. John, Horsleydown	169	130	SouthwarkVauxhall
4	St. James,	132	192	SouthwarkVauxhall
	Bermondsey			
5	St. Mary Magdalen	186	175	SouthwarkVauxhall
6	Leather Market	148	155	SouthwarkVauxhall
7	Rotherhithe	198	158	SouthwarkVauxhall
8	Battersea	92	56	SouthwarkVauxhall
9	Wandsworth	115	178	SouthwarkVauxhall
10	Putney	15	17	SouthwarkVauxhall
11	Camberwell	132	135	SouthwarkVauxhall
12	Peckham	47	89	SouthwarkVauxhall
13	Christchurch,	160	71	Southwark&Lambeth
	Southwark			
14	Kent Road	147	96	Southwark&Lambeth
15	Borough Road	197	170	Southwark&Lambeth
16	London Road	144	52	Southwark&Lambeth
17	Trinity, Newington	152	-100 - < ₽	Southwark&Lambeth
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Mortality Rates from Snow Table XII

	First 12 sub-districts	135	147	first12	
32	Sydenham	105	27	Lambeth	4
31	Duiwich	0	0	Lambeth	-
30	Streatham	171	17	Lambeth	4
29	Norwood	5	25	Lambeth	
	Camberwell		25	T 1 1	-
28	St. George,	111	83	Southwark&La	nbeth
27	Clapham	70	101	Southwark&La	nbeth
26	Brixton	55	33	Southwark&La	nbeth
25	Kennington (2nd)	81	75	Southwark&La	nbeth
24	Kennington (1st)	77	125	Southwark&La	nbeth
23	Lambeth Church (2nd)	203	72	Southwark&La	mbeth
22	Lambeth Church (1st)	117	27	Southwark&La	nbeth
21	Waterloo Road (2nd)	132	64	Southwark&La	nbeth
20	Waterloo Road (1st)	137	41	Southwark&La	nbeth
19	St. Mary, Newington	102	66	Southwark&La	mbeth
18	St. Peter, Walworth	149	130	Southwark&La	nbeth
	Sub-Districts	10,000	10,000	Supplier	
		1849 per	1854 per	Water]