

A Gentle Introduction to Causal Mediation Analysis

Justin Armanini^{1, 2, *}

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About me

Justin Armanini is a PhD candidate at the University of Milano-Bicocca, funded by Fondazione IRCCS Istituto Nazionale dei Tumori, Milan, Italy.

His research focuses on **merging Causality and Natural Language Processing (NLP)** methods to support clinical decisions.



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DEI TUMORI

- ① Causal Mediation Analysis theory
- ② An application of Causal Mediation Analysis for Fairness Assessment

Causal Mediation Analysis theory

What is (Causal) Mediation Analysis?

- Mediation analysis is the study of how a treatment X influences an outcome Y **through one or more intermediate variables**, called mediators M
- Mediation analysis enables us to understand **the pathways by which a cause produces an effect**
- Causal mediation analysis performs such analysis through the lenses of **Causal Inference framework**
- While traditional mediation is prevalent in the literature, causal mediation has been gaining in popularity since the early 2000s as an alternative method for assessing mediation [1]

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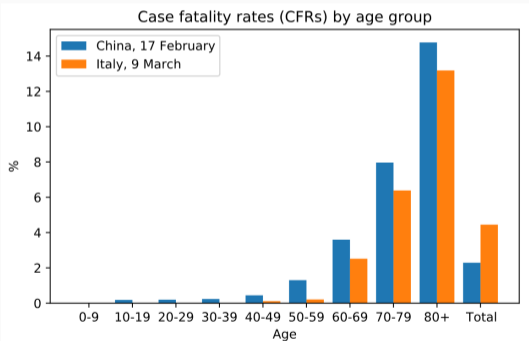
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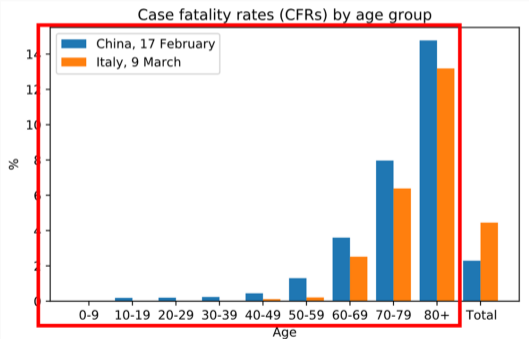
Case Fatality Rate (CFR): proportion of confirmed cases that end fatally



- Problem:
 - for all age groups CFRs in Italy are lower than those in China,
 - but the total CFR in Italy is higher than that in China
- Which country was “better” at managing the pandemic?
- Simpson’s Paradox[2]: data tells different stories when analyzed overall vs. by age group

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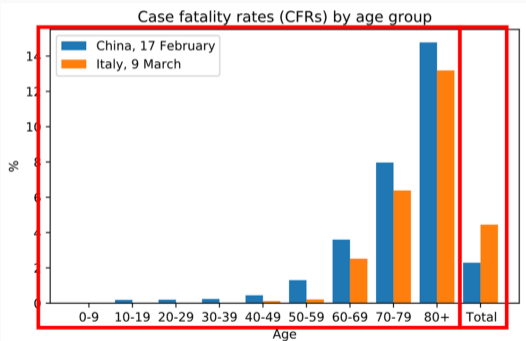
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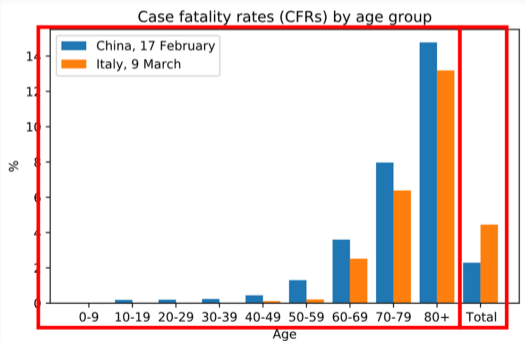
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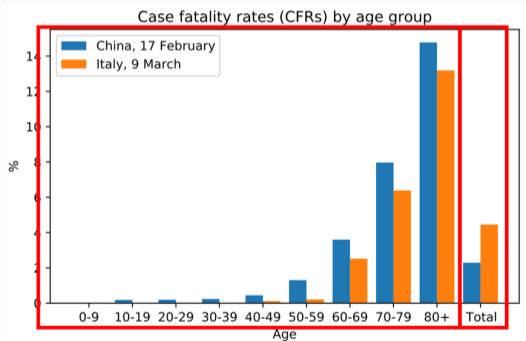
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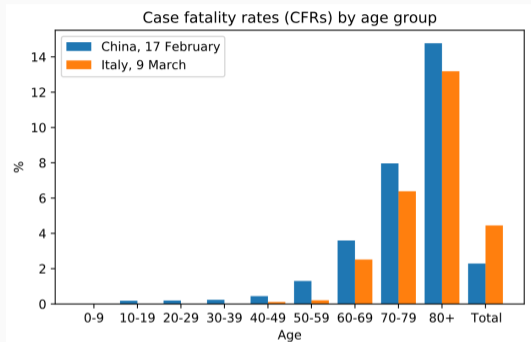
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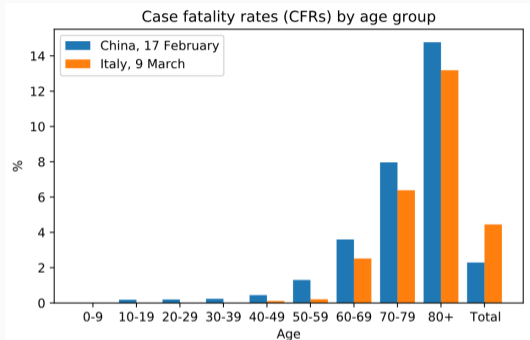
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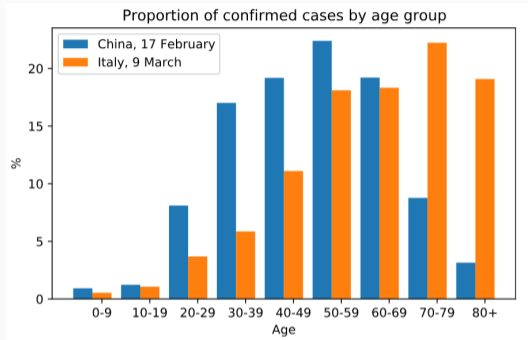
How can such a pattern be explained?

CFRs are relative frequencies!

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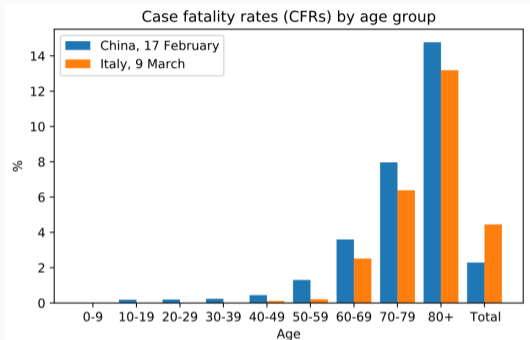
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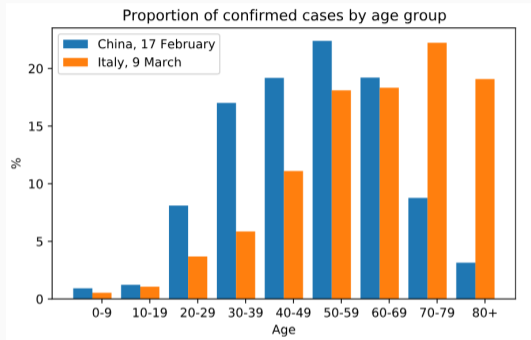
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- Case age distribution differed by country
- Italy: majority aged 60+ (higher mortality risk)
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Conclusion

1. The Italian population is older than the Chinese one
2. Italy had a larger share of confirmed cases among elderly people
3. The elderly are generally at higher risk when contracting COVID-19 effect
4. This explains the Simpson's paradox in the data

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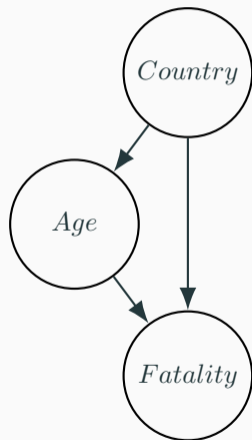
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Causal Mediation Analysis

Causal Graph

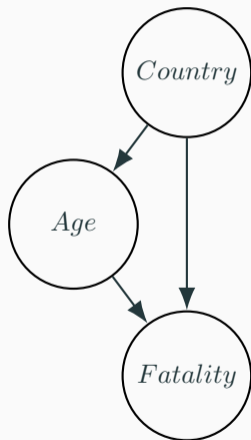


This is a simple and coarse-grained view of a complex underlying phenomenon:

- (*Country* → *Age*) the case demographic is country-dependent
- (*Age* → *Fatality*) COVID-19 is more dangerous for the elderly
- (*Country* → *Fatality*) summarizes country-specific influences on case fatality **other than age** (medical infrastructure, availability of hospital beds and ventilators, local expertise, pandemic-preparedness, air pollution levels, etc.)

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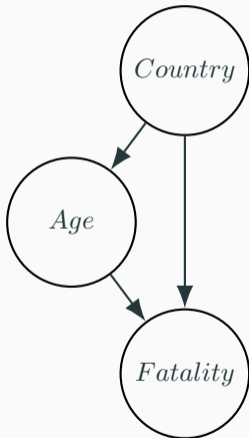


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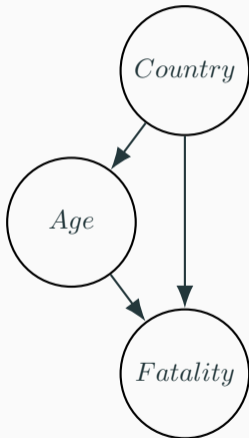


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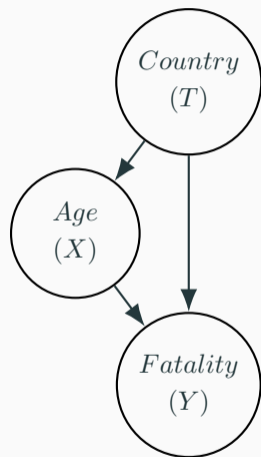
Indeed in our example we are

- **not** concerned with how **controlling X** may bring about change in Y,
- **but** in understanding how **natural variations of X** affect the outcome Y
- we cannot force a change from country X to country Y!

We can compute four types of effects:

- **Total Causal Effect**
- **Controlled Direct Effect**
- **Natural Direct Effect**
- **Natural Indirect Effect**

We will see their causal estimands, Pearl provides theorems for identification in [\[3\]](#)



Total Causal Effect (TCE)

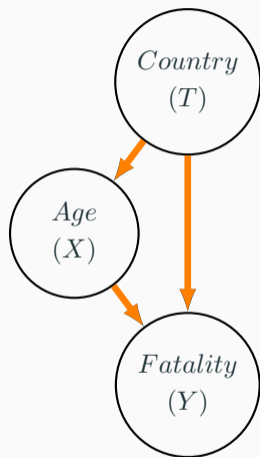
Question:

"What would be the (**overall**) effect on fatality of changing country from China to Italy?"

Causal estimand:

$$\begin{aligned} \text{TCE}_{0 \rightarrow 1} = & \mathbb{E}[Y | do(T = 1)] \\ & - \mathbb{E}[Y | do(T = 0)] \end{aligned}$$

0:China; 1:Italy



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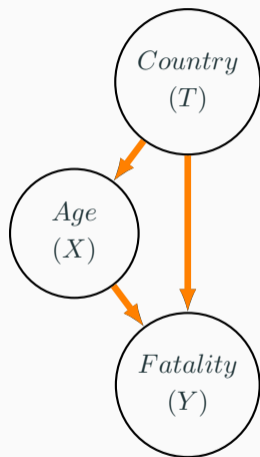
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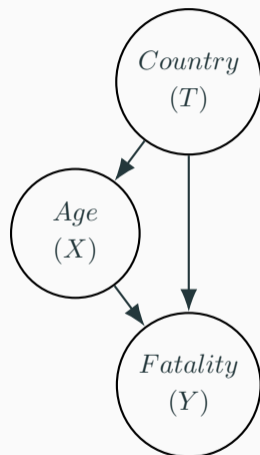
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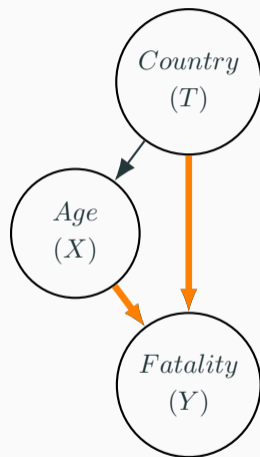
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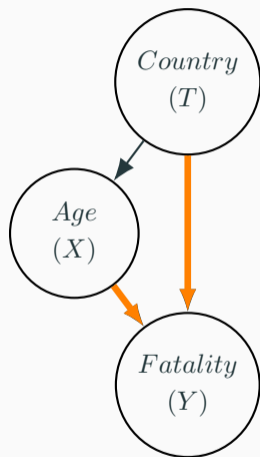
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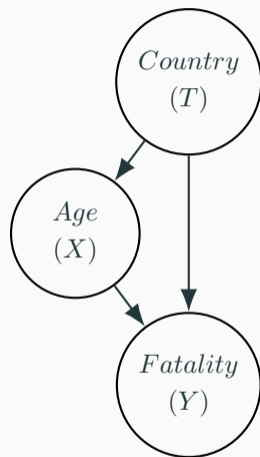
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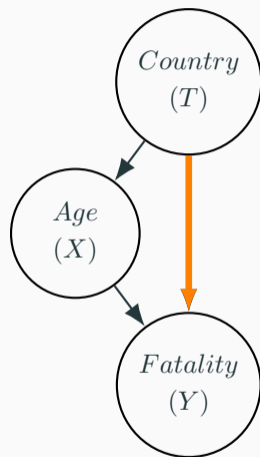
Question:

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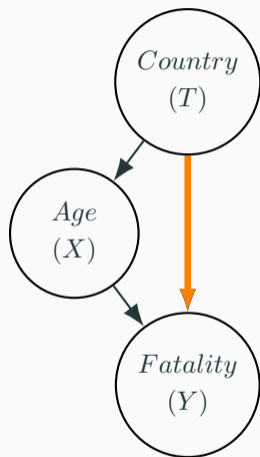
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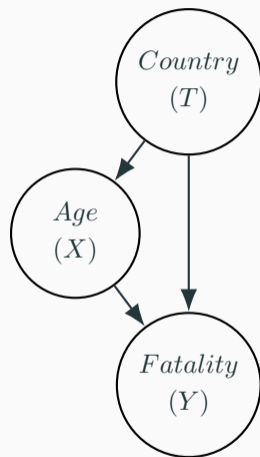
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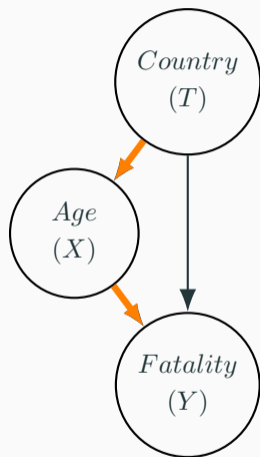
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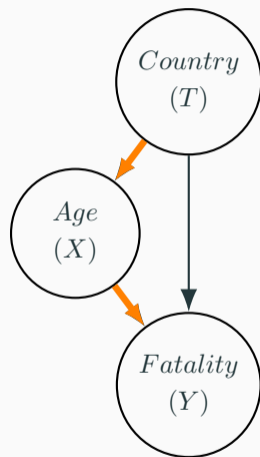
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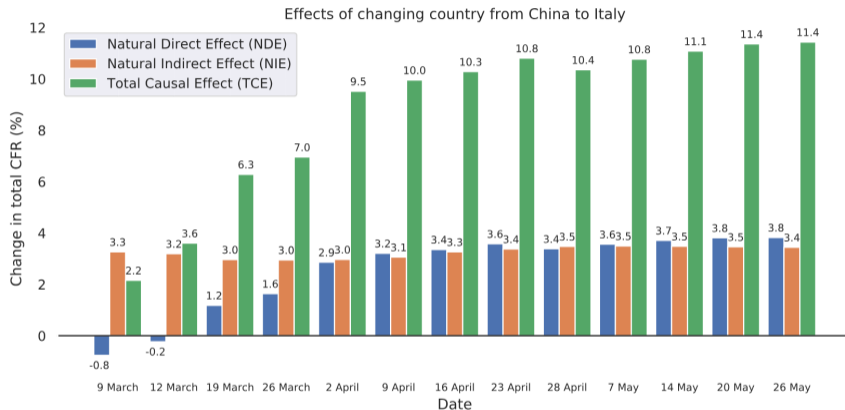
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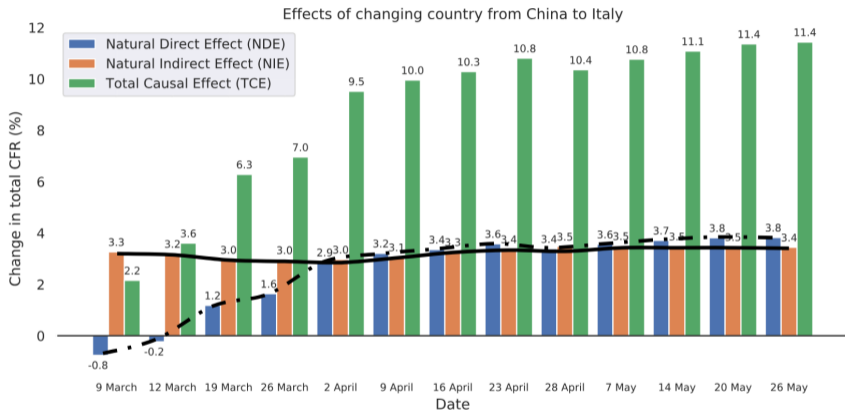
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Simpson's Paradox through Causal Mediation Queries



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- Over time, it is mainly the NDE that drives the observed changes over time

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An application of Causal Mediation Analysis for Fairness Assessment

Under algorithmic fairness [4]:

- we aim to uncover discriminatory biases of models
- we frame **discrimination** as a causal influence of a protected attribute X (such as age, sex, ethnicity, etc.) on an outcome Y of interest along **paths that are considered unfair** in the specific context
- (again) we study how natural variations of X affect the outcome Y

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Triage Discrimination: Myth or Reality?

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³Bicocca Bioinformatics Biostatistics and Bioimaging Centre – B4, Via Follereau 3, 20854, Veduggio al Lambro (MB), Italy

⁴Department of Epidemiology and Data Science, Fondazione IRCCS Istituto Nazionale dei Tumori, Via Giacomo Venezian 1, 20133, Milan, Italy

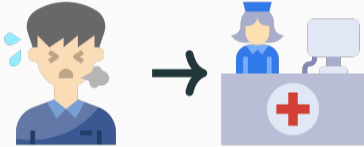
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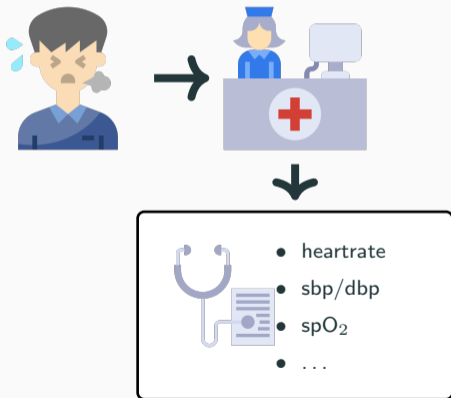
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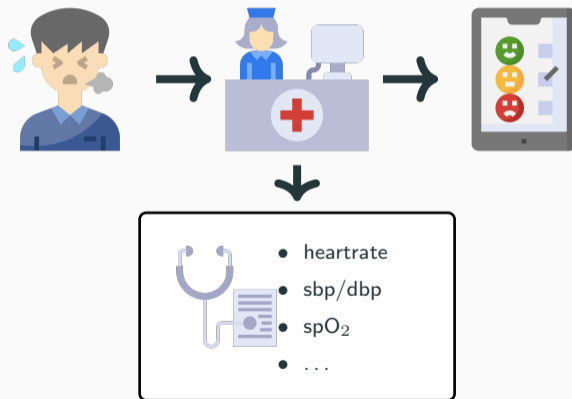
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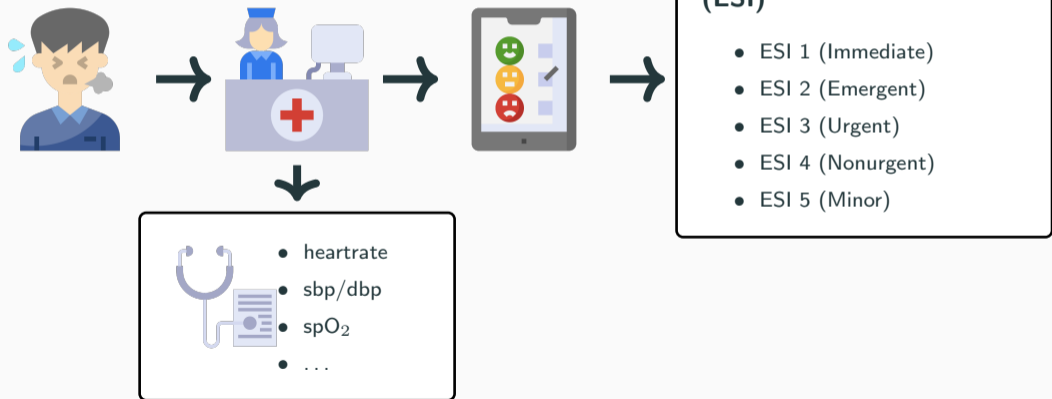
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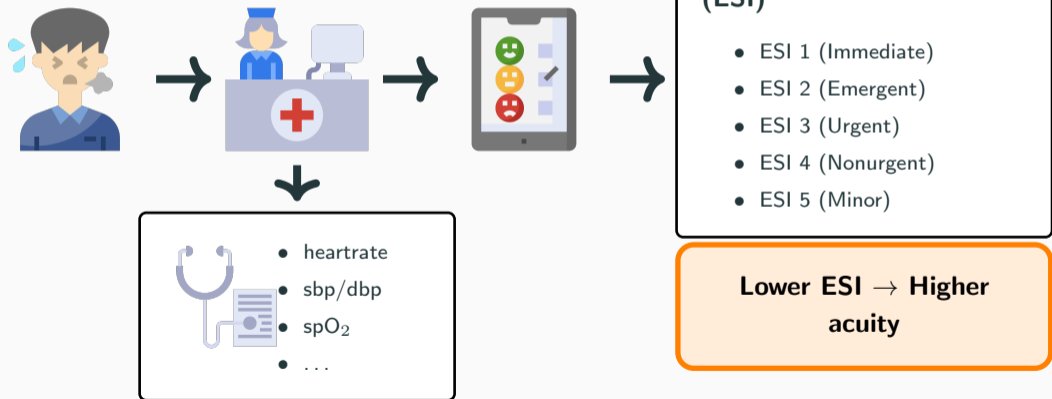
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By Rikki Klaus, CNN

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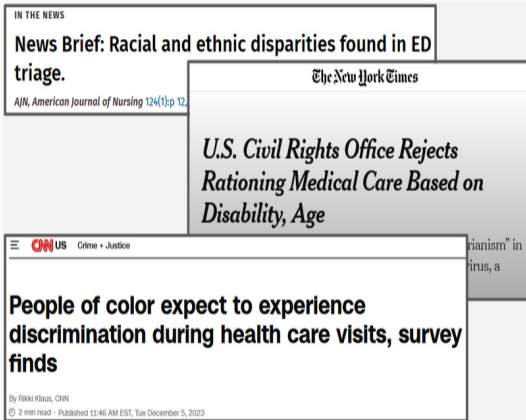
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We use the term 'race' as it is used in the relevant literature—as a socially constructed category—and only in a neutral, descriptive sense!



Why is Discrimination a Concern?



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- Inadequate pain management
- Increased mortality rates
- Decreased trust in medical institutions
- Violations of fundamental principles of medical ethics and social justice

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- Is there disparity in triage based on demographics such as race, gender, and age?

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Our original contribution: first application of causal mediation analysis to triage discrimination

MIMIC-IV-ED (v2.2)

- **Source:** Beth Israel Deaconess Medical Center, Boston (US)
- **Time period:** 2011-2019
- **Sample size:** 160k patients

Variables

- **Demographics:** Age, Gender, Race
- **Clinical:** temperature, heart rate, respiratory rate, spO2, sbp, dbp, pain level
- **Outcome:** ESI Acuity Score (1-5)

Preprocessing

- **Discretized continuous variables** according to the healthcare literature
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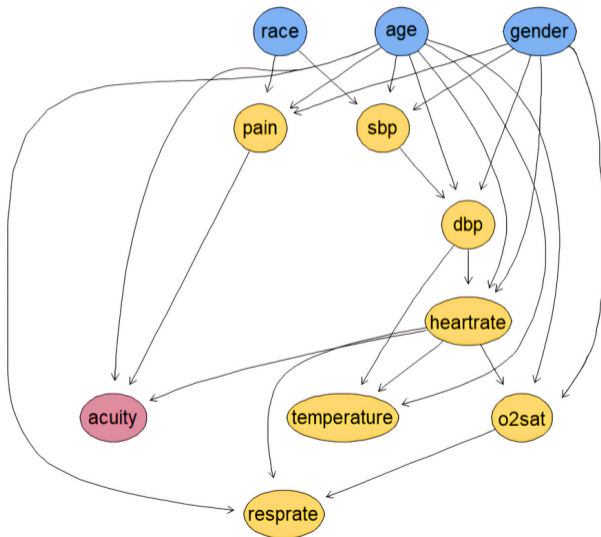
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Learned Causal Network



Causal Network

- Directed Acyclic Graph (DAG)
- Global probability distribution over variables

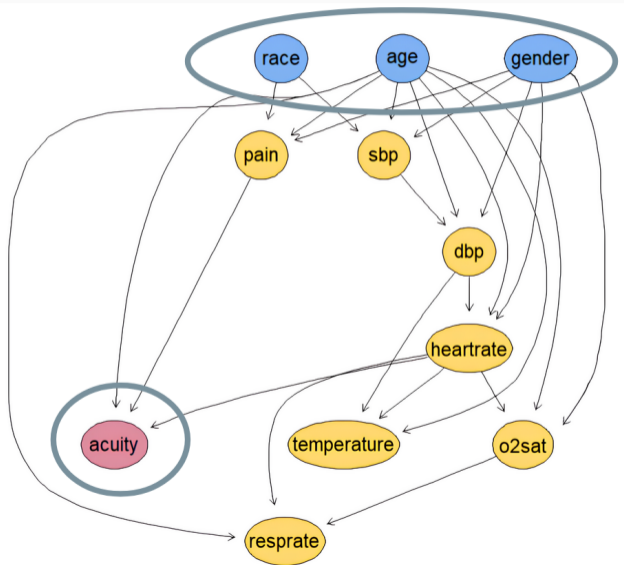
Interpretation

- Edges represent cause-effect relationships
- Distinguishes direct vs. indirect effects

Causal Learning

- Data
- Domain knowledge

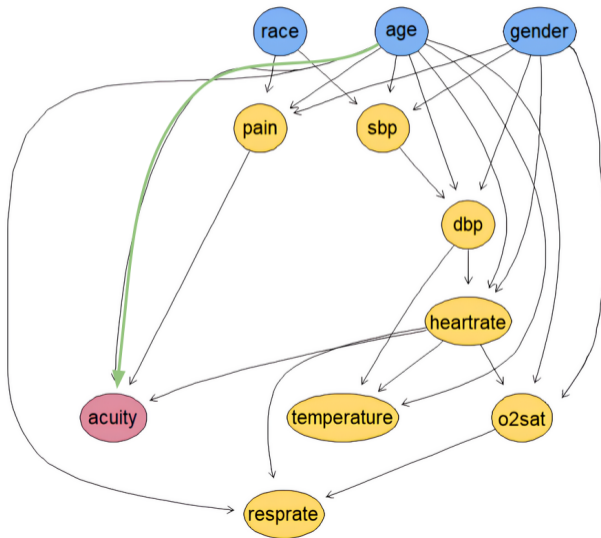
Mapping Mediation Queries



Total Causal Effect

What is the overall causal effect of a demographic variable on the triage outcome?

Mapping Mediation Queries



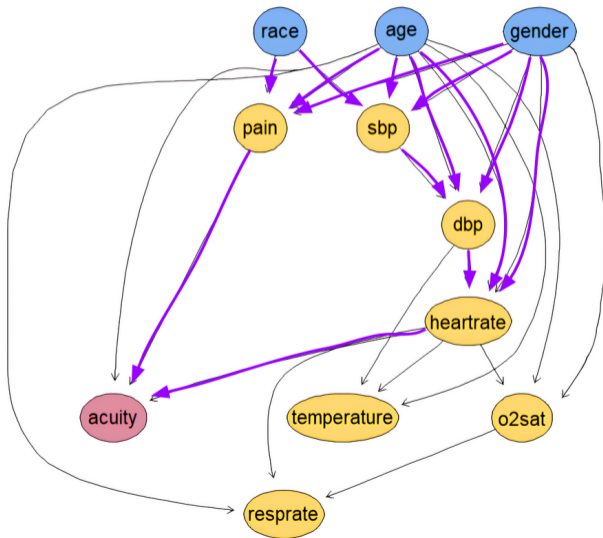
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("Potentially unfair disparity" or "Potential discrimination")

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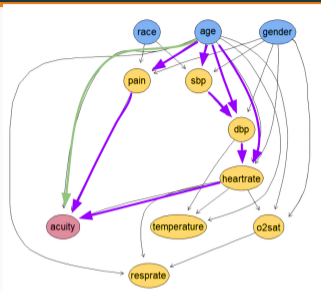
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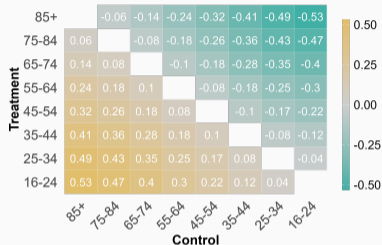
What is the causal effect mediated by clinical factors?
("Clinically explained disparity")

Causal Mediation Analysis: Does Discrimination Occur by Age? Why?



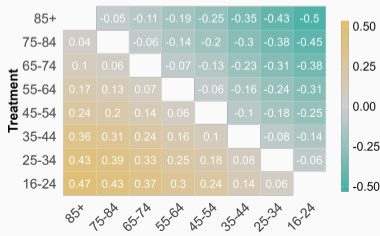
Age as treatment

TCE



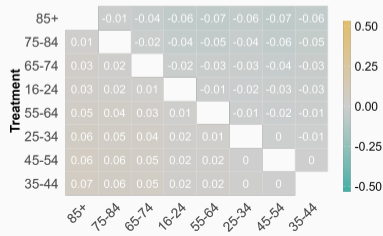
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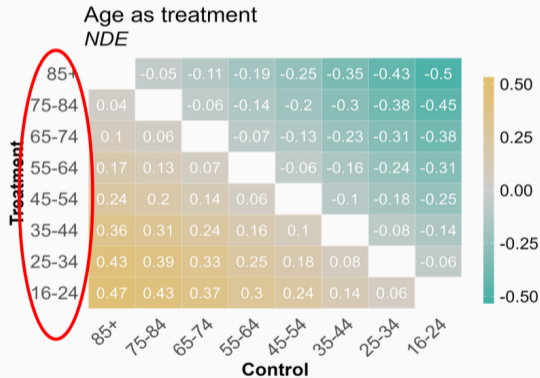


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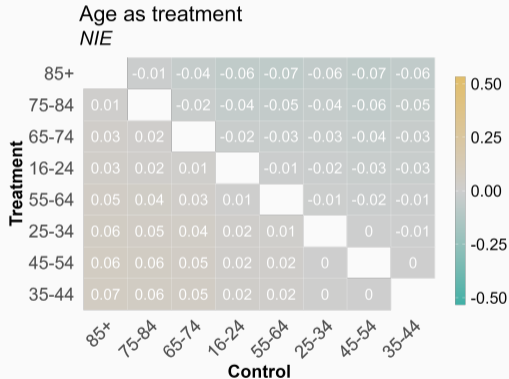
NIE



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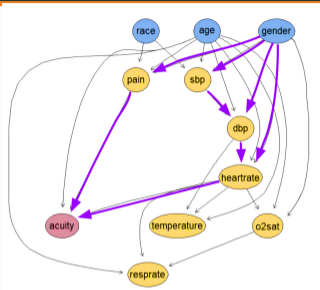


"Potentially unfair disparity"

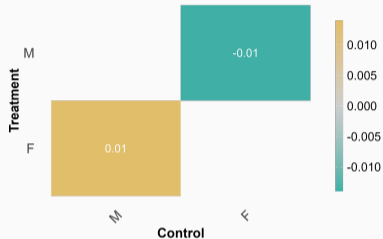


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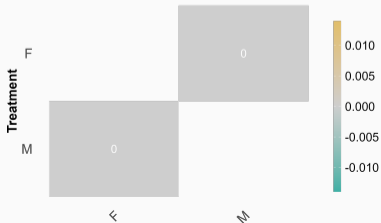
Causal Mediation Analysis: Does Discrimination Occur by Gender? Why?



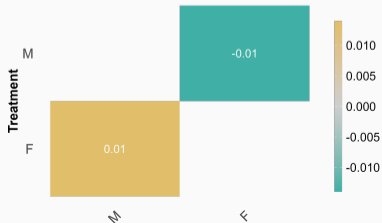
Gender as treatment
TCE



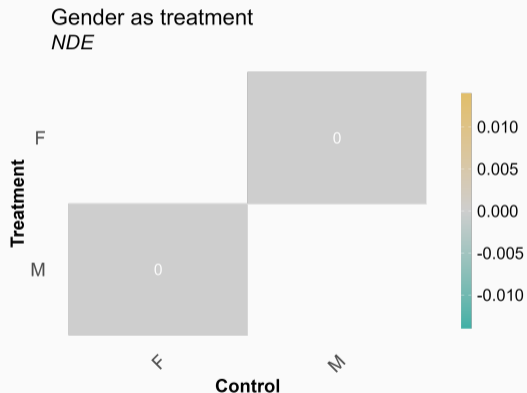
Gender as treatment
NDE



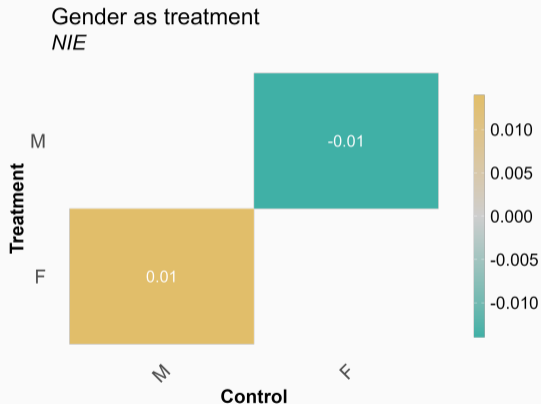
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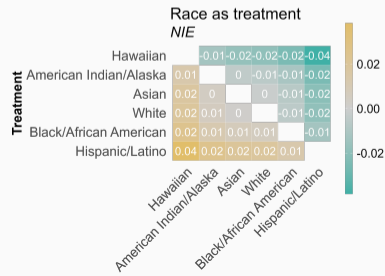
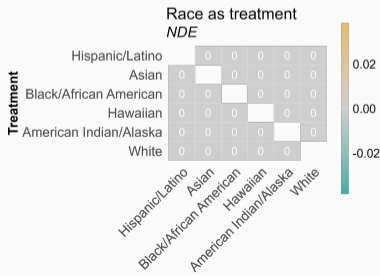
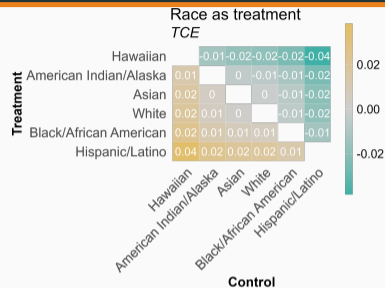
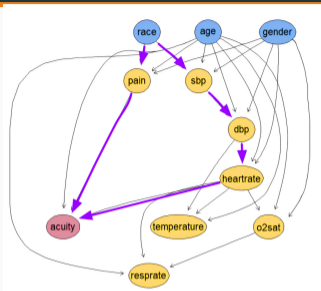


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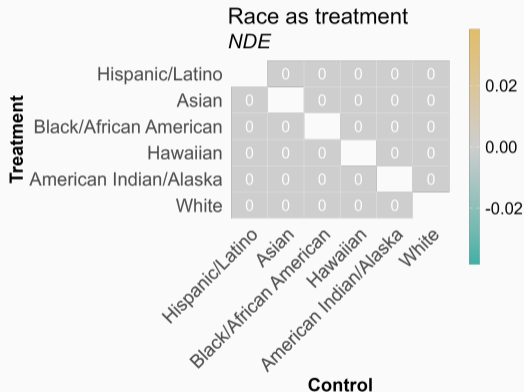


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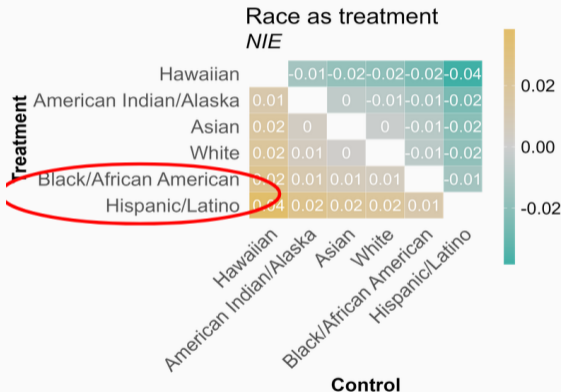
Causal Mediation Analysis: Does Discrimination Occur by Race? Why?



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"Potentially unfair disparity"



"Clinically explained disparity"

Statistical Methods

Data

Causal Mediation

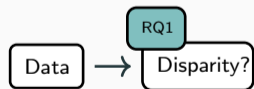
Data

Conclusions

Statistical Methods

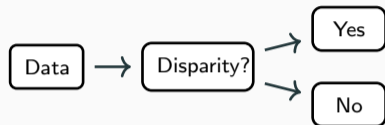


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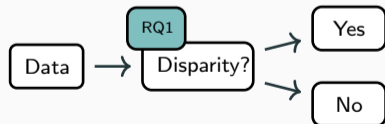


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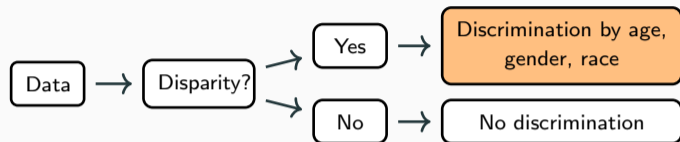


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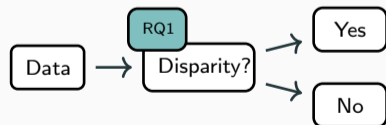


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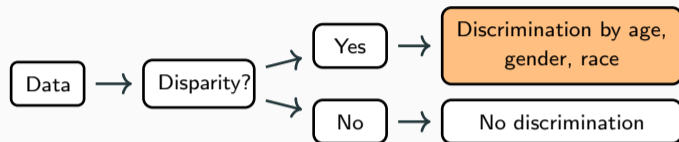


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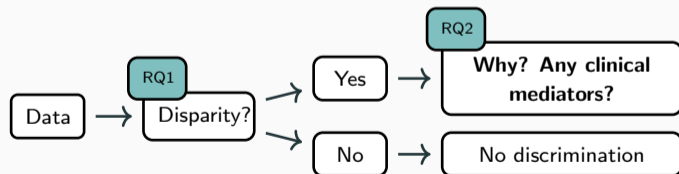


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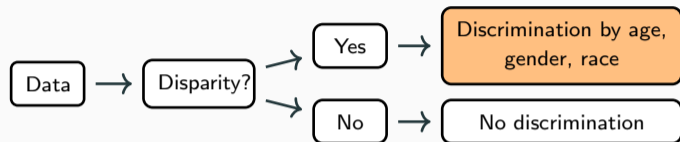


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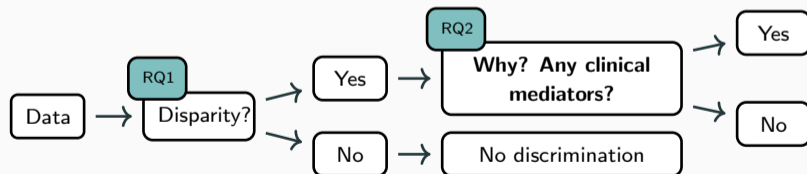


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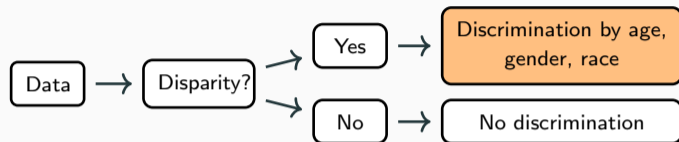


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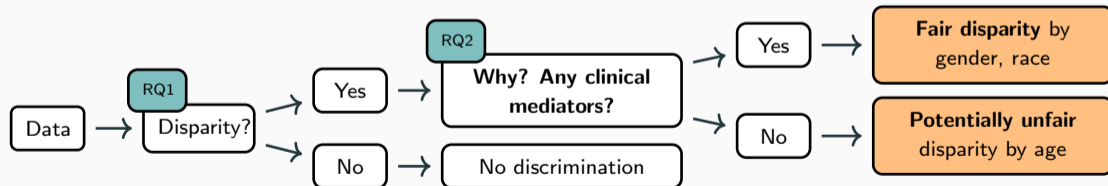


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Causal Mediation



Key Takeaways

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Thank You for your Attention!

MADLab

The Models and Algorithms for Data & Text Mining Laboratory is a research lab at University of Milano-Bicocca focused on Causal Networks, Bayesian Networks and Continuous-Time Bayesian Networks applied to Healthcare and Medicine.

More at: <https://mad.disco.unimib.it/>



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