Improving prediction of adolescent suicide attempts with electronic health records:

Leveraging external data sources of similar patients

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UCONN

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Suicide is rising in adolescence

- According to the CDC, suicide rates of 10 to 24 year olds in the US increased 57% between 2007 and 2018^{1,2}
- Almost 80% of individuals under 19 years old who died by suicide had contact with healthcare in year prior³
- With this, federal mandates have been established to identify and prevent suicide, especially in healthcare^{4,5}



Suicide prediction

- One method to identify at-risk patient is creating statistical algorithms
- To date, several algorithms have been published using electronic health records (EHRs)⁶⁻⁸
- However algorithms only identify <u>around half</u> of true positive pediatric attempters when setting specificity to 90% (similar to declaring the 10% of patients as high risk)



Suicide prediction

| Study | Cohort | Model | AUC | Sensitivity at 90% Specificity |
|--|--------------------------------------|---------------------------|-----------|--------------------------------------|
| Su et al (2020) ⁶ | 10-18 patients in pediatric hospital | Logistic regression | 0.84-0.86 | 0.53-0.65 |
| Xu et al (2021) ⁷ | 10-24 inpatients in CT | Logistic regression | 0.82 | 0.43 |
| Barak-Corren et al (2020) ⁸ | Patients in pediatric hospital | naïve Bayes classifier | 0.72 | 0.37 |

Improving prediction

Since we can only detect ~50% of cases, we must continue to improve algorithms, but how?

- Machine learning models?
 - E.g, random forest
 - difficult to interpret for clinicians and does not increase information from EHR
- Use clinical notes?
 - E.g., Natural Language
 Processing
 - Very costly, time-consuming, and complex

- Collect more data like social determinants of health surveys?
 - I.e., patient-reported outcomes
 - Difficult in real-world context, time-consuming, costly, lack of staff compliance
- Leverage data from other patients or individuals?
 - Only if we have enough information to match patients

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Leveraging data with fusion

- Using a principle in transfer learning, we may be able to generate additional patient features
- Specifically, using *data fusion⁹*, we may be able to match patients on their known features (i.e., demographics and diagnosis codes) with patients or individuals in other datasets on the same and generate new features
- Other datasets (*external datasets*) may have information about risk not found in the EHR used for prediction (*target datasets*)

Leveraging data with fusion

What kind of risk information could external datasets have?

- More data. May contain more data to create better "informed" suicide risk scores when modelling
- Unique features. May contain unique risk factors like social support not well-measured in the target data
- Unique cases. May contain attempters with diagnosis profiles not found in the target data to learn more about target attempters

The general framework

Now that you have a basic idea of the concept, let's describe the general framework:

- 1) Identify an external dataset(s) and feature(s) of interest
- 2) Match each patient in both datasets using a similarity metric like Pearson's *r*
- 3) Generate new features in the target data by aggregating values from top similar patients in the external data

- They sought to improve suicide risk prediction in CT 10-24 pediatric inpatients (the Hospital Inpatient Discharge Database)
- 485 cases and 38806 controls from 2012-2017
- Transferred risk scores from a model using inpatient and outpatient medical claims data (All Claims Paid Database)



- Ran a logistic model in the external data and extracted patient risk scores
- Matched patients using Pearson's r
- Generated new target features by creating a weighted sum of top 1, 10, 20, 50, and 100 most similar patients
- Ran logistic regression in target data including local EHR and fused features





| _ | HIDD (target cohort) | | APCD (exte | rnal cohort) |
|------------------|----------------------|----------------|--------------|----------------|
| | Case | Control | Case | Control |
| No. of patients | 485 | 38 806 | 2053 | 153 433 |
| Sex, N (%) | | | | |
| Female | 308 (63.51) | 22 937 (59.11) | 1281 (62.40) | 76 533 (49.88) |
| Male | 177 (36.49) | 15 869 (40.89) | 772 (37.60) | 76 900 (50.12) |
| Age group, N (%) | | | | |
| 10–14 | 72 (14.85) | 6266 (16.15) | 368 (17.92) | 46 374 (30.22) |
| 15–19 | 253 (52.16) | 12 798 (32.98) | 931 (45.35) | 53 184 (34.66) |
| 20–24 | 160 (32.99) | 19 742 (50.87) | 754 (36.73) | 53 875 (35.11) |

| Models AUC (9 | AUC (95% CI) | Sensitivity (SD) | | |
|---------------|-------------------|------------------|-----------------|--|
| | | 95% specificity | 90% specificity | |
| Conventional | 0.82 (0.81, 0.84) | 0.24 (0.07) | 0.42 (0.07) | |
| Fusion | 0.86 (0.84, 0.89) | 0.65 (0.02) | 0.70 (0.03) | |

- They found a substantial improvement in prediction
 - Likely provided outpatient information about target patients
 - Risk information about many other patients
- Coefficients of model predictors refined by fused risk scores
- Model of fused risk scores only provided "at-chance" performance

Many more avenues of fusion

- As mentioned earlier, we may leverage different types of external features and data
- In the next slides, I will present case studies including:
 - Some examples of these different data types
 - Various methods for similarity matching

- In case study 1, I present my recent publication, using the same target dataset in Xu et al⁷, HIDD
- However, instead of risk scores, we fuse social determinants of health (SDOH) from a large survey-based study
- SDOH represent a wide number of biopsychosocial factors that describe the circumstances in which we are born and live¹⁰



- SDOH includes concepts such as social support, housing, finances, education, religion, etc.
- Literature suggests SDOH are unique risk factors and protective factors related to suicide risk¹¹⁻¹⁴
- In EHR, SDOH are not welldocumented and if present, may only approximate many of these concepts¹⁵



- We extracted 23 SDOH from The National Longitudinal Study of Adolescent to Adult Health (Add Health)¹⁶
 - N cases = 230; N controls = 6271
- For example:
 - Degree to which mother cares and father cares
 - Frequency of hanging out with friends
 - Living in a one-family house
 - Religious orientation
 - Body image
 - Having physical altercations while intoxicated
 - Perception of being killed by age 21

- We followed the same framework as Xu et al (2021)⁷ but expanded it in a number of ways:
- Fusion variables were generated using Pearson's *r* and Manhattan *d*
- A second set of fusion variables were generated by calculating weighted versions of similarity scores
 - Weights for each diagnosis code in matching derived from a logistic model of suicide risk in Add Health

| Models | AUC (95% CI) | Sensitivity (95% CI) | | |
|--------------|--------------------|----------------------|--------------------|--|
| | | 95% specificity | 90% specificity | |
| Conventional | 0.82 (0.81, 0.83) | 0.28 (0.25, 0.31) | 0.44 (0.39, 0.49) | |
| Fusion | 0.83 (0.82, 0.84)* | 0.31 (0.27, 0.35)* | 0.48 (0.43, 0.52)* | |

*Note. *improvements significant at 95% confidence level*

- Including fused SDOH variables improved prediction modestly
 - Totally unique patients
 - Used mock diagnosis codes to match patients as Add Health did not contain these codes directly
 - SDOH may work in-tandem/interact with each other, complex relationships
- Various SDOH with different matching methods appeared in all models and highly predictive in subsequent analyses
 - Provided unique prediction of attempts
 - Mother caring
 - Having no religious orientation

Fusing Suicide risk screening results

- In case study 2, I present preliminary work predicting suicide attempts in 10-18 patients that were hospitalized and/or seen in the emergency department of a large urban medical center in CT
- Here, I acknowledge the importance of suicide risk screening and the potential to fuse screening results from a pediatric hospital which universally screened patients seen in their ED
- As a component of federal mandates^{4,5}, suicide risk screening is becoming an essential task in healthcare facilities
- Screeners show validity in predicting ideation & suicide attempts^{17,18}

- A large pediatric hospital in CT used a pipeline of two screeners:
 - The Ask Suicide-Screening Questions (ASQ) survey¹⁸
 - 5 items result in negative, non-acute positive, acute positive
 - The Columbia Suicide-Severity Rating Scale (C-SSRS) screen¹⁹
 - 6 items results in no, low, moderate, high risk

| Suicide Risk Screel | | O | |
|--|---------------|------|----|
| — Ask the patient: ———————————————————————————————————— | | | 1) |
| 1. In the past few weeks, have you wished you were dead? | O Yes | Q No | 2) |
| 2. In the past few weeks, have you felt that you or your family would be better off if you were dead? | O Yes | Q No | |
| 3. In the past week, have you been having thoughts about killing yourself? | O Yes | Q No | |
| 4. Have you ever tried to kill yourself? | O Yes | O No | |
| If yes, how? | | | - |
| When? | | | |
| | | | 6) |
| If the patient answers Yes to any of the above, ask the following acu | ity question: | | |
| 5. Are you having thoughts of killing yourself right now? | O Yes | Q No | |
| If yes, please describe: | | | |

| SUICIDE IDEATION DEFINITIONS AND PROMPTS | | Past month | |
|--|--------|---------------|--|
| Ask questions that are bolded and <u>underlined</u> . | YES | NC | |
| Ask Questions 1 and 2 | | | |
| Have you wished you were dead or wished you could go to sleep and not wake up | ø2 | | |
| 2) Have you actually had any thoughts of killing yourself? | | | |
| If YES to 2, ask questions 3, 4, 5, and 6. If NO to 2, go directly to question 6. | | | |
| 3) Have you been thinking about how you might do this? | | | |
| E.g. "I thought about taking an overdose but I never made a specific plan as to when where or how I would actually do itand I would never go through with it." | | | |
| 4) Have you had these thoughts and had some intention of acting on them? | | | |
| As opposed to "I have the thoughts but I definitely will not do anything about them." | | | |
| 5) Have you started to work out or worked out the details of how to kill yourself Do you intend to carry out this plan? | 2 | | |
| | | _ | |
| 6) Have you ever done anything, started to do anything, or prepared to do anything end your life? | to YES | NC | |
| Examples: Collected pills, obtained a gun, gave away valuables, wrote a will or suicide note, | | | |
| took out pills but didn't swallow any, held a gun but changed your mind or it was grabbed fr your hand, went to the roof but didn't jump; or actually took pills, tried to shoot yourself, cu | | | |
| yourself, tried to hang yourself, etc. | | | |
| If YES, ask: <u>Was this within the past three months?</u> | | | |

- Target data included 338 cases and 7533 controls between 2012-2017
- External N = 17366 patients screened in the first two years (2019-2021)
 - 2799 Positive screens
- We use similarity matching as in case study 1 but also expand methods to include:
 - Jaccard's distance
 - cosine similarity

| | Target | | Exte | ernal |
|---------------------------|-------------|--------------|-----------------|-----------------|
| Variable | Cases | Controls | Positive Screen | Negative Screen |
| No of patients, N (%) | 338 (4.29) | 7533 (95.71) | 2799 (16.12) | 14567 (83.88) |
| Gender, N (%) | | | | |
| Male | 104 (30.77) | 3564 (47.31) | 845 (30.19) | 7483 (51.37) |
| Female | 234 (69.23) | 3969 (52.69) | 1954 (69.81) | 7084 (48.63) |
| Age group, N (%) | | | | |
| 10-13 | 95 (28.11) | 2877 (38.19) | 991 (35.41) | 6801 (46.69) |
| 14-18 | 243 (71.89) | 4656 (61.81) | 1808 (64.59) | 7766 (53.31) |
| Race/ethnicity, N (%) | | | | |
| White | 160 (47.34) | 2672 (35.47) | 1423 (50.84) | 6201 (42.57) |
| Black or African American | 70 (20.71) | 2118 (28.12) | 421 (15.04) | 2635 (18.09) |
| Hispanic or Latino | 81 (23.96) | 2297 (30.49) | 646 (23.08) | 4251 (29.18) |
| Other | 27 (7.99) | 446 (5.92) | 309 (11.04) | 1480 (10.16) |

| Models | AUC (95% CI) | Sensitivity (95% Cl) | | |
|--------------|-------------------|----------------------|-------------------|--|
| WOUCIS | AUC (95% CI) | 95% specificity | 90% specificity | |
| Conventional | 0.83 (0.79, 0.87) | 0.49 (0.42, 0.57) | 0.63 (0.55, 0.72) | |
| Fusion | 0.84 (0.80, 0.87) | 0.54 (0.46, 0.62)* | 0.64 (0.56, 0.72) | |

Note. *improvements significant at 95% confidence level

- Fusion improved prediction in a similar magnitude as case study 1, but only when setting specificity to 95%
- Inclusion of fused suicide risk screening variables provided both unique prediction and adjusted coefficient weights of local diagnosis codes
- Particular suicide risk scores were important to prediction:
 - Cosine: top 1000 similar patients
 - Weighted Jaccard's distance: Top 10 similar patients
 - Weighted Manhattan distance: Top 10 similar patients
 - Methods generate variables with differing importance

Fusing Similarity with cases



- In the final case study, I again present preliminary results predicting attempts in 10-24 patients that were hospitalized or seen in the ED, but within a *small suburban* medical center
- Here, another avenue of similarity is explored, which is to skip feature generation and include similarity scores directly

- With this, following the same framework, various similarity scores were fused to the target cohort (N cases = 173; N controls = 4322)
 - Similarity with cases in a large multicenter dataset of hospitalized and emergency patients in CT (N cases = 2828; N controls = 92752)
 - Expand the framework by not only calculating average scores but also median scores
 - Aggregating top k similar patients is not required

| | Tai | rget | External |
|---------------------------|-------------|--------------|--------------|
| Variable | Cases | Controls | Cases |
| No of patients, N (%) | 174 (3.87) | 4322 (96.13) | 2828 (2.96) |
| Gender, N (%) | | | |
| Male | 58 (33.33) | 1910 (44.19) | 1023 (36.17) |
| Female | 116 (66.67) | 2412 (55.81) | 1802 (63.72) |
| Age group, N (%) | | | |
| 10-14 | 34 (19.54) | 1063 (24.60) | 548 (19.38) |
| 15-19 | 91 (52.30) | 1455 (33.66) | 1253 (44.31) |
| 20-24 | 49 (28.16) | 1804 (41.74) | 1027 (36.32) |
| Race/ethnicity, N (%) | | | |
| White | 129 (74.14) | 2505 (57.96) | 1491 (52.72) |
| Black or African American | 6 (3.45) | 359 (8.31) | 363 (12.84) |
| Hispanic or Latino | 22 (12.64) | 1171 (27.09) | 752 (26.59) |
| Other | 17 (9.77) | 287 (6.64) | 222 (7.85) |

| Models | AUC (95% CI) | Sensitivity (95% CI) | | |
|--------------|-------------------|----------------------|-------------------|--|
| WOUCIS | | 95% specificity | 90% specificity | |
| Conventional | 0.81 (0.77, 0.85) | 0.41 (0.36, 0.46) | 0.59 (0.53, 0.66) | |
| Fusion | 0.84 (0.78, 0.89) | 0.51 (0.43, 0.60)* | 0.61 (0.52, 0.72) | |

- Alike case study 2, fusion only improved predictions when setting specificity to 95%, however, AUC appeared to trend toward improvement
- <u>INCLUDING DIAGNOSIS CODES</u>, the strongest predictor in models (by magnitude) was case similarity using standard Pearson's *r*
- Inclusion of case similarity lessened importance of key local diagnosis codes:
 - Prior attempts
 - Suicidal ideation
 - Depression
Concluding statements

- All of these studies are generally **proof of concept**, however, improvements in prediction appear hopeful
- Identifying as many attempters as possible before an event occurs is critical, especially in pediatric populations
- Applied and methodological work is needed to better understand the mechanics underlying differences in improvement, similarity metrics and top patients included, and generated fusion variables
- All of these studies have limitations and full-scale grants dedicated to fusion may lead to larger improvements, better understanding

Concluding statements

- Importantly, obtaining information such as SDOH or suicide risk screening results directly from patients is ideal
- However data collection, especially in healthcare settings, is timeconsuming, costly, and subject to noncompliance²⁰
- "Estimating" these important features via data fusion is a free, time efficient method
- If such fusion algorithms are deployed in real-world settings, we may be able to help prevent additional suicide attempts in pediatric populations, as well as, in general populations

Thank you!

Thank you all for attending my talk!

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Similarity equations

- Below is Pearson's *r* and Manhattan *d*
- For i in n_t = total sample of target, For j in n_e = total sample of external, For q in p features

$$r_{ij} = \frac{\sum_{q}^{p} W \left(x_{iq}^{t} - \frac{\sum_{q}^{p} W x_{iq}^{t}}{\sum_{q}^{p} W_{q}} \right) \left(x_{jq}^{e} - \frac{\sum_{q}^{p} W x_{jq}^{e}}{\sum_{q}^{p} W_{q}} \right)}{\sum_{q}^{p} W_{q}} \left(\frac{\sum_{q}^{p} W \left(x_{iq}^{t} - \frac{\sum_{q}^{p} W x_{iq}^{t}}{\sum_{q}^{p} W_{q}} \right)^{2}}{\sum_{q}^{p} W_{q}} \right) \left(\frac{\sum_{q}^{p} W \left(x_{jq}^{e} - \frac{\sum_{q}^{p} W x_{jq}^{e}}{\sum_{q}^{p} W_{q}} \right)^{2}}{\sum_{q}^{p} W_{q}} \right)$$

$$d_{ij} = 1 - \frac{\sum_{q}^{p} W_{q} \left| x_{iq}^{t} - x_{jq}^{e} \right|}{p}$$

Time for an example

First study in context of suicide risk: Xu et al (2021)⁷

| Models | PPV (<i>SD</i>) | |
|--------------|-------------------|-----------------|
| | 95% specificity | 90% specificity |
| Conventional | 0.057 (0.018) | 0.050 (0.009) |
| Fusion | 0.134 (0.015) | 0.082 (0.008) |

Social determinants of health

PPV (95% CI)

| Models | | | |
|--------------|----------------------|----------------------|--|
| | 95% specificity | 90% specificity | |
| Conventional | 0.068 (0.059, 0.077) | 0.054 (0.046, 0.063) | |
| Fusion | 0.074 (0.064, 0.085) | 0.058 (0.050, 0.067) | |

Suicide risk screening

| Models | | |
|--------------|-------------------|-------------------|
| | 95% specificity | 90% specificity |
| Conventional | 0.30 (0.27, 0.33) | 0.22 (0.20, 0.24) |
| Fusion | 0.32 (0.29, 0.35) | 0.22 (0.20, 0.24) |
| | | |

Case similarity

PPV (95% CI)

| Models | | |
|--------------|-------------------|-------------------|
| | 95% specificity | 90% specificity |
| Conventional | 0.24 (0.22, 0.26) | 0.19 (0.18, 0.21) |
| Fusion | 0.28 (0.25, 0.32) | 0.20 (0.17, 0.22) |