

Department of Medicine 5841 South Maryland Avenue Chicago, IL 60637 ☎: 773 7021234

Ishanu Chattopadhyay

Assistant Professor
Section of Hospital Medicine
Department of Medicine
900 E 57th Street
KCBD 10152
Chicago IL 60637
a: 814 5315312
Sishanu@uchicago.edu
zed.uchicago.edu

Editor Science Advances

Dear Editor

Please find enclosed the manuscript entitled "Reduced False Positives in Autism Screening Via Digital Bio-markers Inferred from Deep Co-morbidity Patterns" for your consideration for publication in Science Advances as an original article.

Autism spectral disorder (ASD) is currently screened for in toddlers using standard questionnaires, .e.g. M-CHAT/F with a high number of false positives (85%) and a low sensitivity (38%) – translating to long wait-times for confirmatory diagnosis (≈ 1 year). Children lose crucial time within which interventions are the most effective. Additionally, questionnaires are prone to interpretive biases leading to systematic under-diagnosis in diverse communities.

Here we report an orthogonal methodology that requires **no questionnaires or blood-work**. Using novel stochastic learning on individual diagnostic codes already recorded during past doctor's visits, we design a risk estimator significantly out-performing the state-of-the-art. Without pre-selecting comorbid disorders, we learn hidden patterns in medical histories to enable cutting down false positives by upto half without losing specificity. Additionally, we get new insights into co-morbid risk that might shed light into ASD pathobiology.

Author contact details:

- Dmyto Onishchenko onishchenko@uchicago.edu 3123588479
- Yi Huang yhuang10@uchicago.edu 3124040499
- Jim van Horne jvanhorne@chicagobooth.edu 3125027754
- Peter J. Smith—psmith2@peds.bsd.uchicago.edu 7737023095
- Michael Msall mmsall@peds.bsd.uchicago.edu 7737040885
- Ishanu Chattopadhyay (correponding author)— ishanu@uchicago.edu 8145315312

We are led by a multi-disciplinary team of machine learning experts (Chattopadhyay PH. 8145315312) and clinical practitioners with substantial scholarly work in pediatric developmental psychology and autism: 1) Peter J. Smith, Professor of Pediatrics and Executive Committee Chair, American Academy of Pediatrics' Section on Developmental & Behavioral Pediatrics, and 2) Prof. Michael Msall, Section Chief of Developmental & Behavioral Pediatrics, UChicago, and Fellow of Joseph P. Kennedy Research Center on Intellectual & Neurodevelopmental Disabilities, UChicago.

We look forward to, and humbly hope for, your positive response.

Sincerely,

Ishanu Chattopadhyay Chicago, IL Thursday 22nd October, 2020