acute disease presentation are linked to poorer adherence. Recognizing the characteristics of adherence patterns within specific diagnostic categories is crucial for designing precise interventions to enhance patient outcomes and optimize the overall effectiveness of treatment.

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Linking Digital Traits from Facial Expression, Voice, and Head Motion to Montgomery–Åsberg Depression Rating Scale Subscales

Z. Zhu¹, Y. Wu¹, J. Seidel², D. Roy¹ and E. Salzmann^{2*}

¹Boehringer Ingelheim Pharmaceuticals, Inc., CT, United States and ²Boehringer Ingelheim International GmbH, Ingelheim am Rhein, Germany

*Corresponding author. doi: 10.1192/j.eurpsy.2024.576

Introduction: The 10-item Montgomery–Åsberg Depression *Rating* Scale (*MADRS*) measures different dimensions of depression symptomatology. Digital traits may generate deeper understanding of the MADRS subscales and provide insights about depression symptomatology.

Objectives: To identify digital traits that predict specific MADRS subscales and ascertain which digital traits are important for which MADRS subscales.

Methods: During a Phase II decentralised clinical trial in major depressive disorder (MDD), patients completed the MADRS and used AiCure (LLC, New York, NY, USA), a smartphone application, to complete image description tasks at baseline. Digital measurements identified from the literature as relevant to MDD symptomatology were conducted using audio and video data derived from the image description tasks. Digital measurements included speech (rate, sentiment and first-person singular pronouns), vocal acoustics (intensity, pause fraction and fundamental frequency), facial expressivity (regional facial movement) and head pose (Euclidean and angular head movement). Digital traits analysis involved data pre-processing followed by machine learning (ML) using Elastic Net, Decision Tree, and Random Forest models; model performance was evaluated using 5-fold crossvalidation and mean absolute error (MAE). Important digital traits were calculated by percentage change in MAE after permuting a specific variable. Important digital traits for the MADRS Apparent Sadness subscale score were mapped to defined, interpretable domains.

Results: The ML model predictions varied for different MADRS subscales (**Table**). Overall, Elastic Net and Random Forest models

outperformed Decision Tree across all subscales scores other than suicidal thoughts. Half of the literature-based digital traits contributed to the prediction of \geq 1 MADRS sadness sub-scale score. The important digital traits for the Apparent Sadness subscale score could be mapped to 4 domains (**Figure**); this aligned with findings from the literature. **Image:**

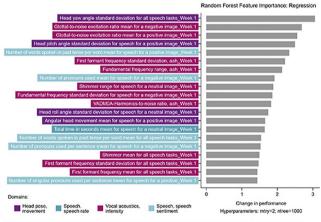
Table. Machine learning model performance on MADRS Total Score and subscale scores, MAE (standard error)

MADRS Scales (baseline)	Elastic Net	Decision Tree	Random Forest
Apparent Sadness	0.80 (0.10)	1.07 (0.20)	0.83 (0.10)
Concentration Difficulties	0.80 (0.26)	1.01 (0.13)	0.84 (0.10)
Inability to Feel	0.79 (0.15)	1.04 (0.22)	0.81 (0.18)
Inner Tension	0.82 (0.33)	1.05 (0.31)	0.82 (0.33)
Lassitude	0.78 (0.20)	0.92 (0.30)	0.77 (0.15)
Pessimistic Thoughts	0.87 (0.16)	1.03 (0.21)	0.85 (0.22)
Reduced Appetite	0.85 (0.32)	0.75 (0.08)	0.75 (0.26)
Reduced Sleep	0.82 (0.11)	0.81 (0.24)	0.81 (0.13)
Reported Sadness	0.61 (0.24)	0.72 (0.25)	0.60 (0.27)
Suicidal Thoughts	0.97 (0.13)	0.88 (0.22)	0.98 (0.08)
MADRS Total	0.83 (0.14)	0.96 (0.20)	0.85 (0.17)

Serien text indicates best performance (MAE (DE)) across the three machine learning methods for each subscale; highlighted text indicates so ap three lowest total (MAE (3E). MORDS: Montperenv-Alabre Decreasion Rating Scale; MAE: mean absolute error: SE: standard error

Image 2:

Figure. Important traits for MADRS Apparent Sadness subscale score at baseline



ADRS, Montgomery-Asberg Depression Rating Scale; miry, miry is a hyper-parameter in Random Forest that specifies the size of the variable subset that is andomy picked for each random forest iteration; ntree, number of trees used in aggregation; VADMOA, Vocal Acoustic Digital Measure Assessment, 0A

Conclusions: Digital traits collected from patients with MDD were able to predict certain MADRS subscales better than others. **Funding:** Boehringer Ingelheim.

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