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- 2 3 Time in treatment: Examining mental illness trajectories across inpatient 4 psychiatric treatment 5 Hyuntaek Oh^{a,b*}, Jaehoon Lee^c, Seungman Kim^c, Katrina A. Rufino^{b,d}, Peter Fonagy^e, 6 7 John M. Oldham^{a,b}, Bella Schanzer^{a,b}, Michelle A. Patriguin^{a,b} 8 9 ^a Menninger Department of Psychiatry and Behavioral Sciences, Baylor College of 10 Medicine, 1977 Butler Blvd., Houston TX, 77030 USA 11 ^b The Menninger Clinic, 12301 S Main St, Houston TX, 77035 USA 12 ^c Department of Educational Psychology and Leadership, College of Education, Texas 13 Tech University, 3002 18th Street, Lubbock TX, 79409 USA 14 ^d Department of Social Sciences, University of Houston – Downtown, One Main St, 15 Houston TX, 77002 USA ^e Research Department of Clinical, Educational and Health Psychology, University 16 17 College London, UK 18 19 * Corresponding Author: Hyuntaek Oh, PhD. 20 12301 S Main St, Houston TX 77030 USA 21 Email: hoh@menninger.edu
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24 Abstract

25 Early discharge or reduced length of stay for inpatient psychiatric patients is related to 26 increased readmission rates and worse clinical outcomes including increased risk for 27 suicide. Trajectories of mental illness outcomes have been identified as an important 28 method for predicting the optimal length of stay but the distinguishing factors that 29 separate trajectories remain unclear. We sought to identify the distinct classes of 30 patients who demonstrated similar trajectories of mental illness over the course of 31 inpatient treatment, and we explore the patient characteristics associated with these 32 mental illness trajectories. We used data (N = 3,406) from an inpatient psychiatric 33 hospital with intermediate lengths of stay. Using growth mixture modeling, latent mental 34 illness scores were derived from six mental illness indicators: psychological flexibility, 35 emotion regulation problems, anxiety, depression, suicidal ideation, and disability. The 36 patients were grouped into three distinct trajectory classes: (1) High-Risk, Rapid 37 Improvement (HR-RI); (2) Low-Risk, Gradual Improvement (LR-GI); and (3) High-Risk, 38 Gradual Improvement (HR-GI). The HR-GI was significantly younger than the other two 39 classes. The HR-GI had significantly more female patients than males, while the LR-GI 40 had more male patients than females. Our findings indicated that younger females had 41 more severe mental illness at admission and only gradual improvement during the 42 inpatient treatment period, and they remained in treatment for longer lengths of stay, 43 than older males.

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45 Keywords: length of stay, growth mixture modeling, trajectory, anxiety, depression46

47 Introduction

48 Inpatient psychiatry lengths of stay (LOS) in the United States have been becoming 49 shorter and shorter, with the average inpatient LOS now approximately five to six days 50 (Glick et al., 2011; Sturm and Bao, 2000). Although some studies indicate that shorter 51 LOS is as effective as longer LOS (e.g., 30 days) in reducing symptoms of dementia 52 (Kunik et al., 2001), major depression (Pettit et al., 2005), and schizophrenia (Johnstone 53 and Zolese, 1999), others have indicated that psychiatric morbidity is greater at 54 discharge among psychiatric patients with a shorter LOS than those with a longer LOS 55 (Glick et al., 2011; Ruaño et al., 2013). Meta-analytic results confirm that patients with a 56 history of psychiatric disorders are at an increased risk of suicide (Harris and 57 Barraclough, 1997) and the suicide risk is particularly high when the patients were 58 discharged from psychiatric hospitals where the average LOS is between 5 to 7 days 59 (Appleby et al., 1999; Goldacre et al., 1993). This is congruent with the literature 60 indicating that a LOS of less than 14 days in inpatient psychiatry was significantly 61 associated with a higher suicide risk (Desai et al., 2005). In addition to increased suicide 62 risk with reduced LOS, other studies demonstrated that shorter LOS predicts greater 63 risk of readmission (Lin et al., 2006; Ruaño et al., 2013). Despite the relationship 64 between poorer outcome and reduced LOS, economic pressures to limit LOS for 65 inpatient hospitalization have tended to tip the scale in favor of less time being needed 66 for inpatient treatment (Compton et al., 2006; Hirsch et al., 1979).

In addition to economic pressure, studies have identified other clinical factors which
appear to influence LOS for inpatient psychiatry. These include: prior hospitalization
(Stevens et al., 2001), level of psychopathology (Cohen and Casimir, 1989), medical

70 comorbidity (Lyketsos et al., 2002), suicide risk (Cohen and Casimir, 1989), substance 71 (Warnke and Rossler, 2008), and psychiatric diagnoses (Blader, 2011). use 72 Hospitalization history and diagnoses of schizophrenia, major depression, and other 73 psychotic disorders were correlated with extended LOS, whereas substance use was 74 associated with reduced LOS (Blader, 2011; Clapp et al., 2013; Cohen and Casimir, 75 1989; Hallak et al., 2003; Stevens et al., 2001; Warnke and Rossler, 2008). 76 Associations between LOS and demographic information (e.g., sex, marital status, 77 homelessness) were also noted in several studies but findings are mixed (Averill et al., 78 2001; Cohen and Casimir, 1989; Warnke and Rossler, 2008). A limitation of the 79 correlational approach taken to identifying predictors of LOS is that it assumes an 80 underlying homogeneity and explores primarily linear predictors of LOS. In principle it is 81 perfectly possible that in some instances, when it retards recovery severity predicts 82 increased LOS while in others, where high severity scores expedite rapid improvement, the association is reversed. Notably, there have been few studies that examined the 83 84 trajectories of recovery over the course of psychiatric hospitalization and how these 85 patterns can be predicted from patient characteristics, and how they relate to treatment 86 and follow-up outcomes. Being able to identify groups of patients based on information 87 available on admission (e.g. demographic, self-reported symptom data), allows 88 clinicians to anticipate likely treatment response and support effective decision making 89 by stratifying care (Saunders et al., 2019).

90 Growth mixture modeling (GMM) is a statistical method designed to identify classes of 91 individuals who are homogeneous in terms of their longitudinal change in a set of 92 variables of interest (Muthén, 2006; Ram and Grimm, 2009). GMM has been useful for

93 recognizing classes of patients who share similar longitudinal trajectories of depressive 94 symptoms (Bombardier et al., 2016; Gomez et al., 2017; Saunders et al., 2019; 95 Saunders et al., 2020). However, these were either studies of outpatient treatments or 96 did not consider other psychiatric disorders or their comorbidity. Another study, although 97 the finding could be limited to older adults, has also utilized GMM to examine the factors 98 related to longitudinal trajectories of PTSD symptoms after a hurricane disaster 99 (Pietrzak et al., 2013).

100 The present study applied GMM to longitudinal clinical data obtained from a large 101 sample of adults admitted to an inpatient psychiatric hospital. We considered six key 102 psychiatric indicators to agnostically determine trajectories of change in mental illness 103 outcomes during inpatient psychiatric treatment: psychological flexibility (Kashdan and 104 Rottenberg, 2010), emotion regulation problems (Gross and Muñoz, 1995), anxiety 105 (Ormel et al., 1994), depression (Ormel et al., 1994), suicide ideation severity (Kessler 106 et al., 1999), and disability (Ormel et al., 1994). The aims of this study were to (1) 107 establish the patterns of change in mental health, and (2) identify the variables 108 associated with the trajectories. It was hypothesized that different mental illness 109 trajectories would emerge (e.g., those that showed improvement, those that showed 110 treatment resistance), yet the specific number of trajectories was not assumed as this 111 was considered an exploratory approach. We also examined a number of demographic 112 and clinical variables (see Table 1 and 2) that are relevant to mental illness outcomes 113 and hospitalization to identify the demographic and clinical variables associated with the 114 identified patterns of change in mental illness. Lastly, we hypothesized that improved 115 mental illness outcomes would be associated with longer LOS, rather than shorter LOS.

116

117 Materials and methods

118 Participants

119 Study participants (N = 3,406) were adults admitted to a private inpatient psychiatric 120 hospital in Houston, Texas between 2012 and 2017. The patients were equally split by 121 sex (female: n = 1,637, 48.1%). On average, the patients were 35.24 years old (SD = 122 15.04; range = 17-89); and the majority of patients identified as White (n = 3,029,123 88.9%) followed by multiracial or "other" (n = 176, 5.2%) and Asian (n = 71, 2.1%). LOS 124 was defined as the number of days between admission date and discharge date for 125 each participant. The average LOS of the participants was 42.16 days (SD = 20.9; 126 range = 0-238 days). About half of the patients (n = 1,653, 48.5%) stayed at least six 127 weeks in the hospital.

128

129 Data procedures and measures

Data were collected as part of the clinic's outcomes study aimed to monitor longitudinal treatment response during inpatient psychiatric treatment (Allen et al., 2009). We administrated clinical measures at admission, bi-weekly, and discharge. The study protocols were approved by the Institutional Review Board at Baylor College of Medicine.

The severity of depressive symptoms, anxiety symptoms, psychological flexibility, emotion regulation and dysregulation, and disability were measured by using the Patient Health Questionnaire (PHQ-9) (Kroenke et al., 2001), General Anxiety Disorder Scale (GAD-7) (Spitzer et al., 2006), Acceptance and Action Questionnaire II (AAQ-II) (Bond

et al., 2011; Hayes et al., 2004), Difficulties in Emotion Regulation Scale (DERS) (Gratz
and Roemer, 2004), and WHO Disability Assessment Schedule 2.0 (WHODAS 2.0)
(Üstün et al., 2010), respectively. The Columbia-Suicide Severity Rating Scale (CSSRS) was also used to examine suicidal ideation and behavior in the past 30 days and
at present (Posner et al., 2011).

144 The Structural Clinical Interview for DSM-5 Disorders (SCID-5, SCID-5-PD) is the most 145 widely used semi-structured clinical interview administered by a trained interviewer 146 (First et al., 2015; First, 1997). The SCID-5-PD is used to assess for the presence of 147 DSM-5 personality disorders. The SCID-5 is a diagnostic measure assessing mood 148 disorders, psychotic disorders, substance use disorders, anxiety disorders, obsessive-149 compulsive disorders, eating disorders, somatic symptom disorders, insomnia, and 150 trauma-related disorders. The SCID-5 and SCID-5-PD were administered at admission 151 by trained clinical interviewers under supervision of a licensed psychologist.

152 As we have done previously (Hartwig et al., 2019; Oh et al., 2020), medication and 153 psychotherapy information were extracted from patient medical records and treatment 154 notes, respectively. Medication information (e.g., medication type, medication taken 155 during the treatment) was measured one week prior to patient discharge date, as 156 patients' medication regimen had been stabilized by this time point. Medications were 157 classified into the following categories: tricyclic antidepressants (TCA), selective 158 serotonin reuptake inhibitors (SSRI), antidepressant other (e.g., Wellbutrin, trazodone), 159 serotonin and norepinephrine reuptake inhibitors (SNRI), benzodiazepine, dopaminergic 160 stimulants agents, first- and second-generation antipsychotics, hypnotics, non-opioid 161 analgesics, and miscellaneous analgesics (e.g., gabapentin). Psychotherapy

information (e.g., type of psychotherapy and frequency) was recorded in the patient's
treatment notes after receiving psychotherapies. Psychotherapies were classified into
the following categories: psychodynamic, cognitive-behavioral therapy (CBT), and
family therapy.

166 Data analysis

167 GMM was used to classify patients into distinct classes, each manifesting a unique 168 pattern of change in mental illness measured by six indicators: AAQ, DERS, GAD-7, 169 PHQ-9, C-SSRS, and WHO-DAS. GMM postulates that similarities and differences in 170 longitudinal observations may be explained by the existence of a categorical latent 171 variable that represents a few mutually exclusive classes within the population (Nylund 172 et al., 2007). In this study, a series of third-order GMM models were fitted to the patents' 173 scores on the six mental illness indicators observed at four different time points 174 (admission, 2-week, 4-week, and 6-week; Fig. 1). Those six mental illness indicators 175 were loaded on a first-order latent variable, named Mental Illness, at each time point; 176 and the loadings of each indicator were constrained to be equal across time to achieve 177 temporal measurement invariance. Cross-time residual covariances were also specified 178 for each indicator in the models (they are omitted in Fig. 1 for the sake of simplicity). 179 The Mental Illness latent variables were loaded on two or three second-order growth 180 factors: Intercept and Linear Slope (2 latent variables); or Intercept, Linear Slope and 181 Quadratic Slope (3 latent variables). Those growth factors were then loaded on a third-182 order class factor (1 categorical latent variable). A total of eight models were fitted -1 to 183 4 unique patterns of linear growth (4 models) and another 1 to 4 unique patterns of 184 quadratic growth (4 models) (Table 3). Model parameters, including factor loadings,

185 means and variances/covariances of the growth factors, and class probabilities, were 186 estimated by using robust maximum likelihood (MLR) via accelerated expectation-187 maximization algorithm, which often yields estimates and sandwich standard errors that 188 are robust to non-normality and non-independence of data.

189 Next, the shape of growth (linear vs. guadratic) and the optimal number of growth 190 patterns (classes) were determined by comparing the models in terms of (a) entropy 191 that quantifies the amount of classification error (Celeux and Soromenho, 1996; 192 Ramaswamy et al., 1993) – a larger value and close to 1 indicates less classification 193 error made by the model; (b) average classification posterior probabilities (ACPP) 194 (Nagin, 1999) – high values at the diagonal of a classification table and low values at 195 the off-diagonal of the table indicate good classification guality; and (c) Bayesian 196 Information Criterion (BIC) (Sclove, 1987) – lower values indicate a better fitting model. 197 Note that Parametric bootstrap likelihood-ratio test (BLRT) (McLachlan et al., 2000) was 198 also performed, but the test results were discarded because they had local maxima 199 issues in the bootstrapping process. Entropy and ACPP could not be computed for the 200 case of single class and thus comparisons were not made against the 1-class models.

201 Once the GMM analysis successfully identified distinct classes of patients who share 202 similar longitudinal trajectories of mental illness, a post-hoc analysis was conducted to 203 understand the characteristics of these classes – or equivalently, to find potential 204 antecedents and consequences of differential treatment efficacy. In this second stage, 205 the identified classes were compared in terms of theoretically relevant and meaningful 206 variables such as age, sex, ethnicity, marital status, educational level, occupation, 207 previous experiences of psychotherapy, hospital care, medication use history, and

208 current length of stay, etc. Chi-square tests of independence were used for the 209 comparisons of categorical variables; independent-samples t-tests or analyses of 210 variance (ANOVA), depending on the number of identified classes, were performed for 211 the comparisons of continuous variables. Statistical significance was determined at a 212 0.05 alpha level, or at a level reduced to control for Type I error in multiple pairwise 213 comparisons (i.e., Bonferroni adjustment). All analyses were conducted using Mplus 8.0 214 (Muthén and Muthén, 2018) and SAS 9.4 (SAS Institute, 2013).

215

216 Results

217 Find the distinct patterns of recovery in mental illness

218 The patients were grouped into three mutually exclusive classes as optimally 219 representing three different patterns of (quadratic) change in mental illness over the 220 inpatient treatment period. Specifically, the entropy values of greater than .80 showed a 221 clear delineation of classes in the 'quadratic' GMM models with two or three classes 222 (Celeux and Soromenho, 1996). Further, the ACPP values from the 'quadratic' model 223 with three classes (.84-.92 at the diagonal of the classification table and 0-.08 at the off-224 diagonal) denoted the best classification quality among the models being tested. The 225 BIC values decreased as the number of classes increased in the case of either linear or 226 quadratic patterns of growth. However, caution should be exercised when interpreting 227 this result because BIC tends to overestimate the number of classes (Enders and 228 Tofighi, 2008; Henson et al., 2007; Nylund et al., 2007). Thus, considering the results of 229 entropy and ACPP, we chose the 'quadratic' GMM model with three classes as the final 230 solution.

231 The final model solution was substantively interpretable – the identified classes of 232 patients indeed represented three distinctive patterns of change in mental illness (see 233 Fig. 2). The first class, the largest class including 55.4% of the sample, showed more 234 severe mental illness at the beginning of the treatment (i.e., high latent scores) but their 235 mental illness improved substantially across inpatient psychiatric treatment, particularly 236 in the first two weeks of treatment. Thus, this class was referred to as High-Risk, Rapid 237 Improvement (HR-RI). The second largest class (28%) included patients with less 238 severe mental illness at the start of the treatment (i.e., low latent scores) who made 239 gradual improvement during the treatment. The extent of recovery was lessened at the 240 later stage of the treatment (i.e., the slope of the line is closer to 0). This second class 241 was referred to as Low-Risk, Gradual Improvement (LR-GI). The last class of patients 242 (16.7%) was characterized by more severe mental illness at the beginning of the 243 treatment, similar to the HR-RI class, but these patients made only gradual 244 improvements over the treatment period – High-Risk, Gradual Improvement (HR-GI). 245 The means of the individual indicators of mental illness and their changes at four 246 different time points are shown in Fig. 3. Additionally, follow-up assessments of anxiety 247 and depressive symptoms (at 2 weeks, 3 months, 6 months, and 1-year post-discharge) 248 are displayed in Fig. 3A. As expected, all observed scores were well aligned with the 249 'model-implied' longitudinal trajectories (i.e., estimated latent scores) of mental illness 250 during inpatient psychiatric treatment.

251

252 Identify patient characteristics associated with differential treatment efficacy

253 The post-hoc analyses revealed similarities and differences among the three identified 254 classes. Table 1 and 2 provide the descriptive statistics, the results of bivariate tests, 255 ANOVA, and follow-up pairwise comparisons. On average, the HR-GI class (33.24 ± 256 13.71) was significantly younger than both the HR-RI (35.26 \pm 14.99; adjusted p < 0.05) 257 and the LR-GI (36.38 \pm 15.79; adjusted p < 0.001) classes, while the age of the latter 258 two classes was not significantly different (adjusted p = 0.18). The LR-GI class had 259 more male patients (63.4%) than females (36.6%), while the HR-GI class had more 260 female patients (59.8%) than males (40.2%) (p < 0.001); The HR-RI class had an 261 equivalent number of males (49.6%) and females (50.4%). The three classes 262 demonstrated similar ethnic breakdowns (p = 0.33) – the majority of the class members 263 were white (87.9-90.1%) followed by multiracial or "other" category (5-5.2%) and Asian 264 (1.5-2.3%), as observed in the overall sample. The patients' marital status did not differ 265 among the three classes (p = 0.71). Also, the patients' education level (p = 0.23) and 266 occupation (p = 0.97; see Supplementary) were not significantly related to their patterns 267 of recovery.

268 Regarding clinical history and LOS in treatment, the patients in three different classes 269 reported similar previous experiences of psychotherapy, hospital care, and stopping 270 medication (all p > 0.05; see Table 2). However, significantly fewer patients in the HR-271 RI class took hypnotic medications (19.79 ± 19.72%) compared to the LR-GI class 272 $(27.97 \pm 27.29\%)$ (adjusted p < .01). Also, significantly fewer patients in the HR-RI class 273 took antidepressant other (e.g., Wellbutrin, trazodone) (37.84 ± 33.61%) and SSRI 274 $(34.82 \pm 24.28\%)$ medications compared to the HR-GI class $(44.68 \pm 39.03\%)$ and 41.33275 $\pm 28.59\%$, respectively) (both adjusted p < .01). The HR-RI class (41.49 \pm 20.25) stayed

276 in their current hospitalization for significantly shorter periods of time than the HR-GI 277 class (44.40 \pm 21.29; adjusted p < 0.05), with the HR-RI group staying approximately 3 278 days shorter than the HR-GI class. The average LOS for the LR-GI class (42.27 ± 279 21.90) was in between the average LOS for the two High-Risk classes and did not 280 significantly deviate (adjusted p = 1.00 and 0.17). SCID-5 and SCID-5-PD DSM-5 281 clinical diagnosis differences were also examined within each class (see 282 Supplementary). The most common diagnoses in both the HR-RI and LR-GI classes 283 were (1) major depressive disorder, recurrent (HR-RI = 42%; LR-GI = 39%), followed by (2) anxiety not otherwise specified (HR-RI = 27%; LR-GI = 31%), (3) substance 284 285 dependence disorder (HR-RI = 27%; LR-GI = 29%), (4) generalized anxiety disorder 286 (HR-RI = 19%; LR-GI = 20%), and (5) alcohol abuse disorder (HR-RI = 17%; LR-GI = 287 16%). The HR-GI class also had similar common diagnoses to the HR-RI and LR-GI 288 classes, but with substance dependence disorder more prevalent than anxiety not 289 otherwise specified: (1) major depressive disorder, recurrent (HR-GI = 45%), followed 290 by (2) substance dependence disorder (HR-GI = 28%), (3) anxiety not otherwise 291 specified (HR-GI = 26%), (4) generalized anxiety disorder (HR-GI = 21%), and (5) 292 alcohol abuse disorder (HR-GI = 16%). Finally, the type and frequency of 293 psychotherapy sessions received during inpatient psychiatric treatment did not differ 294 between the three classes (p = .37; see Table 2).

295

296 Discussion

Aims of the current study were to (a) find the distinct classes of patients who demonstrate similar trajectories of mental illness outcomes during inpatient psychiatric

299 treatment, and (b) identify patient characteristics associated with the identified patterns 300 of recovery over time. Through the use of growth mixture modeling, we were able to 301 describe three distinct trajectories (HR-RI, LR-GI, and HR-GI) of mental illness 302 improvement across inpatient psychiatric hospitalization as measured by six indicators: 303 psychological flexibility, emotion regulation problems, anxiety, depression, suicide 304 ideation severity, and disability. Patients' age and sex were significantly related to 305 treatment response, with significantly younger females in the HR-GI group and more 306 males in the LR-GI group.

307 The findings of the present study support our hypothesis that different mental illness 308 trajectories would emerge by identifying three classes of patients that are distinct in 309 terms of mental illness at the beginning of treatment and change over their 310 hospitalization. These findings suggest that the patient classes could be differentiated at 311 the beginning of the treatment and by the improvement of six indicators during inpatient 312 psychiatric treatment. The most common trajectory class was characterized by more 313 severe mental illness at the beginning of the treatment and rapid treatment 314 improvement during inpatient psychiatric treatment, referred as HR-RI (55.4%). The 315 second class, LR-GI (28%), was characterized by less severe mental illness than the 316 HR-RI group at the start of the treatment and gradual improvement during the inpatient 317 treatment. The last class of patients, HR-GI (16.6%), demonstrated the highest severity 318 of mental illness at admission and gradual improvement over the treatment period. Thus, 319 the majority of patients could be categorized as following a path to recovery from a fairly 320 severe experience of mental health and only a smallish group (16%) of severely 321 impacted patients could be categorized as poor responders.

322 Having said that, all trajectory classes demonstrated improvement (significant negative 323 slope of trajectories) in mental illness across treatment and their improvement was most 324 rapid after psychiatric hospitalization, consistent with earlier studies that reported 325 depressive symptoms and suicide ideation reduction following the first week of 326 admission or even brief intervention (Clapp et al., 2013; Czyz and King, 2015; Hirsch et 327 al., 1979; Hopko et al., 2001; Lieberman et al., 1998; Pettit et al., 2005; Prinstein et al., 328 2008; Rocca et al., 2010). Factors contributing to rapid improvement may include 329 medication effects (e.g., antidepressant, antipsychotic agents) (Agid et al., 2003; 330 Posternak and Zimmerman, 2005), removal of alcohol and substances, and 331 psychotherapy gains (Stiles et al., 2003). All classes sustained their improvement 332 related to anxiety and depression post-discharge from the hospital (i.e., anxiety and 333 depression scores post-discharge were in the normal range (Kroenke and Spitzer, 334 2002; Spitzer et al., 2006)), although generalization of these findings is limited as there 335 was participation attrition at follow-up.

Our results are consistent with other studies demonstrating the follow-up assessments of depressive symptoms after hospitalization for traumatic brain injury (Bombardier et al., 2016) and maternal depressive symptoms from pregnancy through 2 years postpartum (Mora et al., 2008). However, these prior studies also reported a trajectory showing gradual reemergence of depressive symptoms which may associate with a history of alcohol dependence and other mental illness disorders (Bombardier et al., 2016).

In the current study, we found that patients' age and sex best predict the particular patterns of mental illness recovery trajectories. The HR-GI class (33.24 ± 13.71) had patients who were significantly younger relative to the HR-RI (35.26 ± 14.99) and LR-GI

345 (36.38 ± 15.79) classes. The LR-GI class had fewer female patients than males, 346 whereas HR-GI class had more female patients than males. These findings are 347 consistent with previous studies that females have shown higher rates of depressive 348 symptoms (Kandel and Davies, 1982), generalized anxiety disorder (Alternus et al., 349 2014; Kessler et al., 1994), and suicidal ideation (Borges et al., 2006; Crosby et al., 350 1999). Additionally, previous studies have consistently demonstrated higher rates of 351 depressive symptoms (Kessler et al., 2010) and suicidal ideation (Crosby et al., 1999) 352 among young adults. For the LR-GI class that had more men than women, as they were 353 admitted to an inpatient psychiatric hospital, the likelihood that they did not present with 354 mental illness symptoms is unlikely. Instead, as men have a higher rate of minimizing 355 self-reported symptoms due to societal ideals of masculinity (Martin et al., 2013), it is 356 possible that symptoms are minimized by this patient group.

357 We examined the LOS in each trajectory class and found that the HR-RI class stayed in 358 the hospital less time (by a few days) than the HR-GI class (adjusted p < 0.05). Note 359 that the average LOS in this study was approximately 42.16 days which is a contrast to 360 the previous studies (five to six days being the average LOS) (Glick et al., 2011; Sturm 361 and Bao, 2000) that indicate negative treatment outcomes, including higher rates of 362 death by suicide post-discharge (Appleby et al., 1999; Goldacre et al., 1993). Therefore, 363 our results highlight not only better understanding of the distinct classes but emphasize 364 the importance of a longer LOS in generating positive treatment outcomes for inpatient 365 psychiatry. Previous studies have shown that patients with cognitive impairment had a 366 significantly increased LOS, but such differences in LOS were not found among patients 367 with depression and anxiety (Furlanetto and da Silva, 2003). However, other studies

368 have reported that depression, anxiety, schizophrenia, mood disorders, and alcohol and 369 drug related disorders are important predictors of LOS (Borchardt and Garfinkel, 1991; 370 Draper and Luscombe, 1998; Huntley et al., 1998; Jiménez et al., 2004; Sloan et al., 371 1999) and that psychiatric comorbidity could have contributed to prolonged LOS (Bressi 372 et al., 2006; Saravay, 1994). In addition to clinical diagnoses, a previous study reported 373 that longer LOS was associated with sex (i.e., females) (Averill et al., 2001). It should be 374 noted that the most common diagnoses in all three classes were recurrent major 375 depressive disorder, anxiety not otherwise specified, and substance dependence 376 disorder. The HR-GI class, which showed longer LOS than the HR-RI class, had more 377 female patients than males, as compared to other two classes. Clearly, the hypothesis 378 (i.e., improved mental illness outcomes would be associated with longer LOS) was not 379 supported by the data, but this group of patients deserves more careful scrutiny and 380 their treatment path (more SSRI and more antidepressant other) indicates that during 381 their slightly longer treatment a wider range of therapies were tried. Therefore, it is likely 382 that significant factors related to LOS for inpatient psychiatry may relate to demographic 383 factors, particularly sex. Perhaps, a bi-factor analytic approach would show them to 384 have higher p Factor scores than the other two groups indicating a higher level of 385 general psychopathology (Caspi and Moffitt, 2018; Smith et al., 2020).

A number of limitations of the present study should be addressed. First, the study sample was from an inpatient psychiatric hospital where most patients were white and had a higher level of education than the general population. Therefore, a future study needs to be replicated with a more diverse sample in order to ensure our findings are not limited to the present sample. Although follow-up assessments of depression and

391 anxiety showed that patients in this study maintained their improved status throughout 392 their first year after discharge, the follow-up results should be interpreted with caution 393 because there is potential for bias in the missing assessments (e.g., individuals who 394 were in other treatment facilities post-discharge) - missing data after discharge: 82.8% 395 at 2 weeks, 85.2% at 3 months, 88.3% at 6 months, and 91.1% at 1 year. While 396 obtaining a more comprehensive follow up sample will always be challenging in the 397 context of routine outcomes monitoring, data obtained from such a small proportion of 398 the sample is very likely to be irredeemably contaminated by non-random forces 399 associated with loss to follow-up (such as an understandable wish to obliterate an 400 experience that may have felt stigmatizing). Data with fewer missing and frequent 401 follow-up assessments of the indicators used in this study may help to improve our 402 understanding of distinct trajectory classes after discharge and how clinical outcomes 403 are/are not maintained following post-discharge.

404 In this study, we identified three distinct mental illness trajectory classes using a large 405 sample of 3,406 patients admitted to a private inpatient psychiatric hospital. The 406 majority of the patients with high mental illness at the beginning experienced substantial 407 improvement within 6 weeks of inpatient psychiatric treatment and maintained their 408 reduction in depression and anxiety throughout their first year after discharge. This 409 study highlights the importance of understanding the relevance of the distinction of 410 classes with meaningful patterns of mental illness treatment change over time. The 411 knowledge of differences between the classes may provide valuable information for the 412 clinicians as well as future researchers making predictions regarding the course of 413 mental illness improvement during inpatient psychiatric treatment and after

414 hospitalization, and ultimately to ensure that patients receive the adequate length of
415 stay for inpatient psychiatric treatment in order to maximize their mental illness
416 outcomes.

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423

424 Contributors

- 425 All authors have approved the final article.
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- 427 Data analysis: HO, JL, SK, KR, and MP.
- 428 Interpretation: HO, JL, KR, PF, JO, and MP.
- 429 Drafting the manuscript: HO, JL, KR, and MP.

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431 Conflict of interest

432 The author reports no conflicts of interest in this study.

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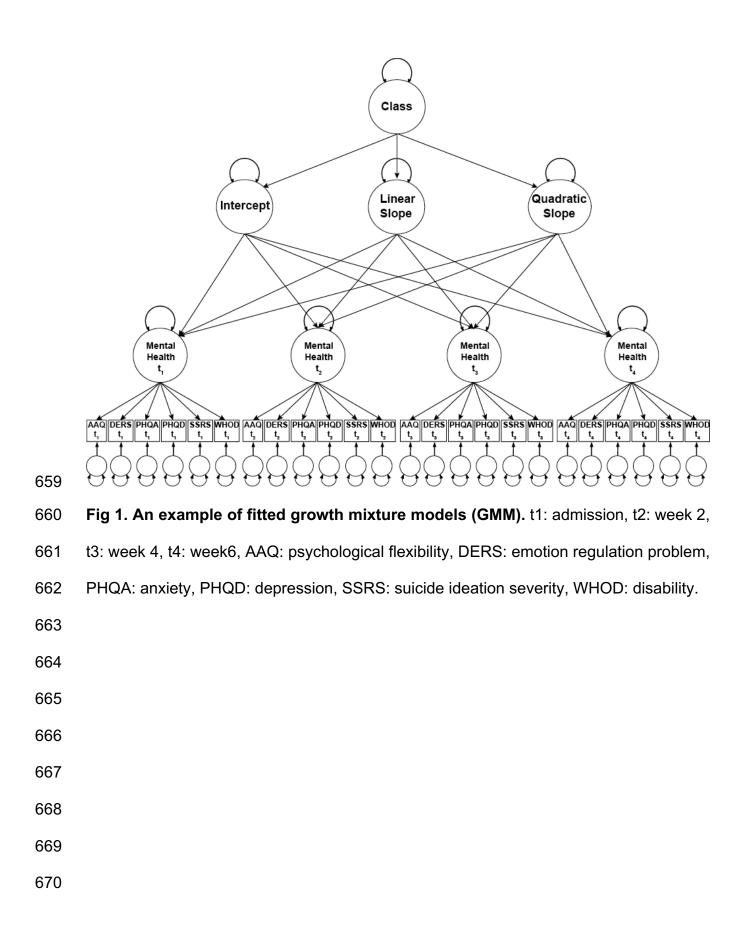
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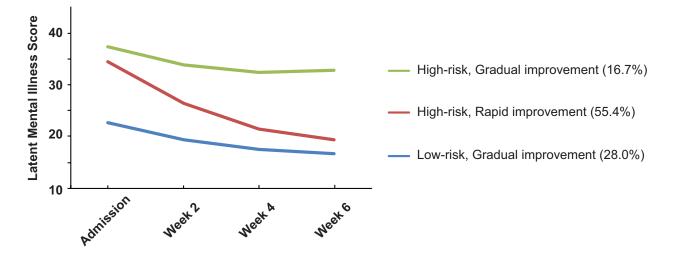
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672 Fig 2. Latent mental illness score in identified classes. Higher latent mental illness

673 scores indicate more disabled mental illness.

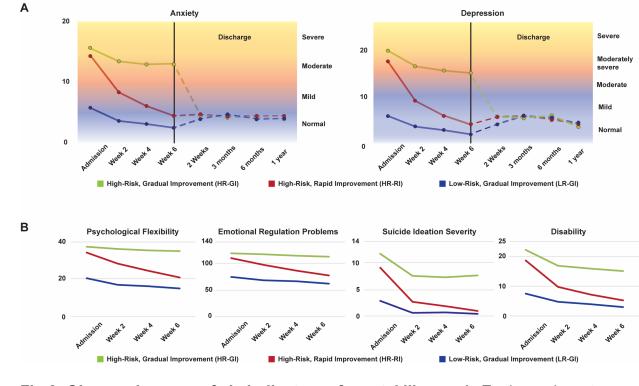


Fig 3. Observed scores of six indicators of mental illness. A. Each number at
Anxiety and Depression indicates the mean of PHQANXIETY and PHQDEPRESSION,
respectively. Follow-up assessments after discharge were characterized in the normal
range. B. Each number at Psychological Flexibility, Emotional Regulation Problems,
Suicide Ideation Severity and Disability indicates the mean of AAQ, DERS, SSRS and
WHOD scores, respectively.

	HR- (<i>n</i> = 1878		LR (<i>n</i> = 948		HR (<i>n</i> = 565		Ove	rall differe	nce	Pairwise difference (p)			
	(11 - 1070	, 55.4%)	(11 – 940	, 20.0%)	(11 – 505)	, 10.7%)					HR-RI		
Variable	M/N	SD / %	M/N	SD / %	M/N	SD / %	F/χ²	df	a	HR-RI vs LR-GI	vs HR- GI	LR-GI v HR-G	
Age	35.26	14.99	36.38	15.79	33.24	13.71	7.76	2, 3388	< .001	.182	.015	< .00	
Sex							85.18	2	< .001				
Female	946	50.4%	347	36.6%	338	59.8%							
Male	932	49.6%	601	63.4%	227	40.2%							
Ethnicity							11.37	10	.330				
American Indian or Alaskan	3	0.2%	3	0.3%	0	0.0%							
native													
Asian	44	2.3%	14	1.5%	13	2.3%							
Black or African-American	22	1.2%	22	2.3%	8	1.4%							
White/Caucasian	1683	89.6%	833	87.9%	509	90.1%							
Native Hawaiian or Pacific	9	0.5%	2	0.2%	2	0.4%							
Islander	9	0.5%	2	0.2%	Z	0.4 %							
Multiracial or other	98	5.2%	49	5.2%	28	5.0%							
Missing	19	1.0%	25	2.6%	5	0.9%							
Highest level of education							22.08	18	.228				
Some elementary school	1	0.1%	0	0.0%	1	0.2%							
Some middle school (junior high)	170	9.2%	87	9.4%	46	8.3%							
Some high school	0	0.0%	3	0.3%	0	0.0%							
High school diploma or	41	2.2%	22	2.5%	10	1.8%							
equivalent	41	Z.Z70	23	2.5%	10	1.0%							
Some college	161	8.7%	78	8.4%	50	9.0%							
Technical or associates degree	631	34.1%	309	33.3%	219	39.6%							
Bachelor's degree	81	4.4%	29	3.1%	22	4.0%							
Master's degree	552	29.8%	281	30.3%	143	25.9%							
Doctoral degree	190	10.3%	105	11.3%	57	10.3%							
Professional degree (JD, MD)	25	1.3%	12	1.3%	5	0.9%							
Marital status							7.15	10	0.711				
Married	475	25.8%	256	27.7%	130	23.8%							
Separated	90	4.9%	45	4.9%	27	4.9%							
Divorced	170	9.2%	85	9.2%	46	8.4%							
Widowed	26	1.4%	16	1.7%	4	0.7%							

	Never married1054Living with someone as married23	57.3% 510 1.3% 11	55.3% 1.2%	330 9	60.4% 1.6%
694	Table 1. Comparisons of demograph	ic variables be	etween the	e class	ses. HR-RI: High-risk, Rapid improvement, LR-GI:
695	Low-risk, Gradual improvement, HR-GI:	High-risk, Grad	lual improv	vement,	, M: mean, SD: standard deviation.
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	HR- (<i>n</i> = 1878)		LR- (<i>n</i> = 948		HR- (<i>n</i> = 565,		Ove	rall differen	се	Pairwi	se differen	ce (<i>p</i>)
Variable	 	SD / %	 	SD / %		SD / %	F/χ²	df	p	HR-RI vs LR-GI	HR-RI vs HR- GI	LR-GI vs HR-GI
Over the course of your life, how much psychotherapy (talk therapy) with a mental health professional have you had (in month)?	70.55	85.07	66.11	82.77	70.55	85.97	0.92	2, 3388	.398	.577	1.000	.987
How many different therapists have you seen?	4.27	4.33	3.96	2.97	4.15	3.67	1.91	2, 3388	.148	.152	1.000	1.000
How many different psychiatrists or health care providers have you seen (for medication) for a mental disorder?	3.22	3.64	2.97	2.32	3.10	2.56	2.04	2, 3388	.130	.137	1.000	1.000
How many times have you been admitted for acute/crisis psychiatric hospital care (i.e., 1-5 days)?	1.58	4.98	1.39	2.96	1.40	2.48	0.86	2, 3388	.425	.732	1.000	1.000
How many times have you been admitted for extended psychiatric hospital treatment (i.e., more than 5 days)?	1.19	4.02	1.11	2.10	1.16	2.11	0.18	2, 3388	.839	1.000	1.000	1.000
Have you stopped medication against or without your doctor's advice? Yes No	824 1028	44.5% 55.5%	395 532	42.6% 57.4%	252 301	45.6% 54.4%	1.43	2	.489			
Have you stopped therapy against or without your therapist's advice? Yes	635	34.3%	310	33.4%	188	34.0%	0.20	2	.906			

No	1217	65.7%	617	66.6%	365	66.0%						
Length of stay (in day)	41.49	20.25	42.27	21.90	44.40	21.29	4.20	2, 3388	.015	1.000	.011	0.170
SSRI Scheduled Taken	35.37 34.82	24.67 24.28	34.57 34.01	26.47 25.84	41.69 41.33	29.13 28.59	5.91 6.46	2 2	.0028 .0016	1.000 1.000	.0051 .0030	.0051 .0032
Antidepressant other Scheduled Taken	38.62 37.84	34.28 33.61	38.59 37.66	30.51 29.98	45.51 44.68	39.96 39.03	4.74 4.9	2 2	.0089 .0076	1.000 1.000	.0099 .0091	.0242 .019
Hypnotic Scheduled Taken	20.23 19.79	20.27 19.72	28.77 27.97	28.05 27.29	27.55 26.87	22 21.46	6.12 5.93	2 2	.0024 .0029	.0043 .0052	.0659 .0688	1.000 1.000
Psychotherapy sessions Psychodynamic Cognitive-Behavior Therapy Family Therapy	8.35 1.22 0.6	7.02 4.02 1.89	8.74 1.27 0.69	7.47 3.72 2.06	8.71 1.35 0.62	7.31 3.82 1.97	.99 .21 .51	2 2 2	.3731 .8114 .6012	.6292 1.000 .9431	1.000 1.000 1.000	1.000 1.000 1.000

711 Table 2. Comparisons of treatment variables and length of stay between the classes. HR-RI: High-risk, Rapid

712 improvement, LR-GI: Low-risk, Gradual improvement, HR-GI: High-risk, Gradual improvement, M: mean, SD: standard

713 deviation, SSRI: Selective Serotonin Reuptake Inhibitors.

Model		LL	# parameters	BIC	Entropy
Linear growth	1-class	-199336	78	399306	-
-	2-class	-195033	81	390724	1.000
	3-class	-194920	84	390522	0.781
	4-class	-194873	87	389921	0.720
Quadratic growth	1-class	-199130	82	398927	-
	2-class	-194674	86	390047	0.999
	3-class	-194509	90	389750	0.796
	4-class	-194412	94	389588	0.685

Table 3. Optimal number of growth patterns (classes). LL: log-likelihood, BIC: Bayesian information criterion.