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1 **Affective instability in those with and without mental disorders: a case control study**

2

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## 29 **1. Introduction**

30 Affective Instability (AI) is a transdiagnostic symptom (Broome et al., 2015b; Henry et al.,  
31 2001b). It has been defined as rapid oscillations of intense affect, with difficulty regulating  
32 these or their behavioural consequences (Marwaha, 2013). Multiple strands of evidence have  
33 associated AI with suicidal thinking (Palmier-Claus et al., 2012; Yen et al., 2004), health  
34 service use (Marwaha et al., 2013c), new onset of depression (Marwaha et al., 2015),  
35 psychotic symptoms (Marwaha et al., 2013a), onset of bipolar disorder and increasing time to  
36 recovery (Howes et al., 2011; Stange et al., 2016). It is also independently linked to greater  
37 medication use and detention under mental health legislation (Patel et al., 2015). It is  
38 associated with childhood trauma experiences (including abuse) and it is suggested that it  
39 may partly explain the connection between these and psychiatric disorders (Aas et al., 2016;  
40 Marwaha et al., 2016; Moffa et al., 2017). The estimated prevalence of AI in the general  
41 population is 14%, with levels being higher in younger people and women (Marwaha et al.,  
42 2013c; Patel et al., 2015). We (Broome et al., 2015a), and others (Harrison et al., 2017) have  
43 suggested that trans-diagnostic investigation of AI is compatible with the NIMH Research  
44 Domain Criteria project (Insel, 2014), a framework for understanding mental disorders by  
45 study of dysfunction in individual psychological and biological systems.

46

47 Through systematically reviewing the literature we have previously identified AI as  
48 having three core affect components: intensity, lability, and ability to control the oscillations  
49 or their behavioural consequences (Marwaha et al., 2014). The review also identified that  
50 comprehensive measurement of all three components is rarely undertaken. The current  
51 literature is limited in part by theoretical and methodological heterogeneity in how AI is  
52 understood and assessed. This means that studies of AI in different diagnostic groups cannot

53 be compared, and hence understanding whether AI is similar in different disorders and how it  
54 contributes to outcomes such as functioning are hard to ascertain. As such, there is a  
55 significant gap in understanding this clinical phenomenon. A second major shortcoming of  
56 the current literature is that nearly all studies to date, apart from a few notable exceptions  
57 (Ben-Zeev and Young, 2010; Ben-Zeev et al., 2009), have lacked comparisons with  
58 individuals without mental disorder. This means it is unclear how far AI represents  
59 psychopathology needing intervention or indeed whether it is a core aspect of abnormal  
60 mental states, or is a feature of normal mental life.

61

62 To our knowledge, only three studies to date have compared AI in different diagnostic  
63 groups using the same assessment procedures but limiting assessment of AI to two if its  
64 domains. Henry et al. (2001a) examined AI using the Affect Lability Scale (ALS) and Affect  
65 Intensity Measure (AIM) in out-patients with Borderline Personality Disorder (BPD) (N=29),  
66 bipolar disorder: type II without BPD (N=14), BPD and bipolar disorder: type II (N=12), and  
67 no BPD or bipolar disorder but other personality disorders (N=93). Lability scores were  
68 significantly ( $p < 0.05$ ) higher in BPD, whilst bipolar patients tended ( $p = 0.06$ ) to have higher  
69 lability scores than other personality disorders. No differences in affect intensity were  
70 observed.

71

72 In a similar study those with bipolar disorder had significantly higher scores on the  
73 euthymia-elation subscale of the ALS as well as significantly higher scores on affect  
74 intensity, whereas those with BPD experienced more shifts between anxiety-depression,  
75 euthymia-anger and significantly fewer shifts between euthymia-elation and depression-  
76 elation (Reich et al., 2012). Most recently Richard-Lepouriel et al (Richard-Lepouriel et al.,  
77 2016) compared ALS and AIM scores in people with bipolar disorder, attention deficit

78 hyperactivity disorder (ADHD) and controls (dentistry students and doctors). Those with  
79 ADHD and bipolar disorder scored higher on ALS than controls, with AIM scores being  
80 highest for ADHD. Whilst affective lability appears to be higher in BPD, and people with  
81 bipolar disorder, results for affective intensity are discrepant between studies with some  
82 evidence that affective intensity may be higher in people with mental disorders than in  
83 controls.

84

85         Given the paucity of previous research, we aimed to expand the diagnostic groups in  
86 which AI is examined (given the suggestion that AI is transdiagnostic), compare these  
87 “cases” with psychologically “healthy controls”, assess AI more comprehensively, and test  
88 whether AI is independently linked to functioning within a clinical population.

89

## 90 **2. Aims**

91

92 We aimed to answer the following research questions:

93

- 94         1. Does affective instability differ between clinical cases and controls and between  
95             diagnostic groups?
- 96         2. To what extent are measures of affective lability, intensity and ability to control affect  
97             correlated in a trans-diagnostic clinical sample?
- 98         3. Is affective instability associated with functioning in a clinical population independent  
99             of diagnosis?

## 100 **3. Methods**

101 We undertook a case-control study among users of secondary care mental health services  
102 (cases) and primary care attenders without evidence of current mental disorder (controls).

103 Ethical approval was obtained from the Coventry and Warwickshire Ethics Committee, UK.

104 Participant consent and data collection was completed by an experienced researcher with a  
105 psychology background.

106

### 107 *3.1 Participants*

108

109 Individuals with a range of diagnoses were recruited from secondary care mental  
110 health services within Coventry and Warwickshire, UK through convenience sampling. The  
111 aim was to include individuals who were representative of the ‘typical’ case mix of these  
112 services, so participants were recruited in out-patient departments, day hospitals, community  
113 mental health teams and a specialist personality disorder service. Inclusion criteria were: a]  
114 aged 18-65 years; b] capacity to give informed consent; c] the primary reason for attending  
115 the mental health service was for management of a clinical diagnosis of BPD, bipolar  
116 affective disorder, major depressive episode (moderate or severe depressive episode) or non-  
117 affective psychosis as reported by a Consultant Psychiatrist. The researcher confirmed the  
118 diagnosis with the Psychiatrist using ICD-10 criteria.

119

120 Exclusion criteria were: a] an acute illness episode (sufficient to require urgent or  
121 inpatient care) according to the patient’s Consultant Psychiatrist; b] unable or unwilling to  
122 complete the assessments (e.g. individuals with a clinically assessed learning disability, with  
123 insufficient command of the English language to understand and complete questionnaires); or  
124 c] individuals with a primary ICD-10 diagnosis of dependency to drugs or alcohol (to avoid  
125 confounding by drug or alcohol misuse).

126

127 Control participants were recruited from primary care (general practitioner surgeries).  
128 Physicians asked patients if they were interested, a researcher in the waiting room then  
129 consented the patients and completed the battery of questionnaires. Exclusion criteria for the  
130 control group were: a] presence of a current mental disorder (including common mental

131 disorders such as depression or anxiety disorders); b] dependency on substances or alcohol;  
132 c] previous diagnosis of BPD, bipolar disorder, or non-affective psychosis, according to their  
133 primary care records.

134

### 135 *3.2 Materials*

136

137 Details were collected on participants' diagnosis (for cases) and confirmed by their  
138 Consultant Psychiatrist. Details on duration of illness (cases only) and current medications  
139 were identified by a researcher, from clinical records. Medications were grouped as anti-  
140 psychotic, anti-depressant, anti-anxiety, mood stabiliser, anti-depressant/mood-stabiliser or  
141 'other' (medication not directly related to the patient's psychiatric diagnosis).

142 Sociodemographic information was collected on age (years), gender, marital status  
143 (married/cohabiting, single, separated/widowed), employment (employed, unemployed,  
144 other), ethnicity (White British, other), and highest education level (None/GCSE, A Level,  
145 Degree/higher degree) (see Table 1).

146

### 147 *3.3 Assessment scales*

148

149 Participants were asked to complete four questionnaires relating to affective / mood  
150 instability.

- 151 1. The Affective Lability Scale - short form (ALS-18) (Harvey et al., 1989), is a  
152 reliable and valid measure comprising 18 items coded 0-3. Overall score is  
153 obtained by taking the mean of the scores for each item as indicated by the scale  
154 developers. Three subscales can be derived; 'anxiety-depression', 'depression-  
155 elation', and 'anger'.
- 156 2. The Affective Intensity Measure-20 (AIM) (Larsen et al., 1986a) was used to  
157 measure affect intensity. The AIM contains 20 items coded 1 to 6. A person's

158 overall score is obtained by taking the mean of the scores for each item. The AIM  
159 has good internal consistency, test-retest reliability and construct validity (Larsen  
160 et al., 1986b).

- 161 3. The Affective Control Scale (ACS) (Williams et al., 1997) comprises 42 items,  
162 coded 1 to 7 (with some items requiring reverse scoring); it has good  
163 psychometric properties including construct validity. A higher ACS score  
164 indicates reduced ability to control affect. Four subscales can be derived; ‘anger’,  
165 ‘positive affect’, ‘depressed mood’, and ‘anxiety’.
- 166 4. Mood fluctuation rate: Because of the lack of a previously well validated scale for  
167 fluctuation rate (Marwaha, 2013) we used a new bespoke schedule for this study.  
168 Mood fluctuation rate was assessed using a question from the Structured Clinical  
169 Interview for DSM Disorders (SCID). It asks the respondent to state how often  
170 they experienced a sudden marked shift in mood. Study participants rated the  
171 number of significant mood fluctuations they experienced over a week.  
172 Respondents were asked to consider this for each one of the weeks in the last  
173 month prior to assessment, and possible responses were 0, 1-3, 4-7, or >7 mood  
174 changes over each week.

175

176 Functioning was measured using the WHO Disability Assessment Schedule 2.0 – 12  
177 item version (WHODAS; Üstün, 2010). This contains 12 items each coded 0 to 4. To obtain  
178 a person’s final score, the simple version entails summing the scores from each of the 12  
179 items, scores range from 0-48. For consistency in comparing with the other scores above, the  
180 mean rather than the sum was used in the current study.



181

182 *3.4 Data analysis*

183

184 Descriptive statistics including means/medians with standard deviations/interquartile ranges,

185 or frequencies with percentages where relevant, were used to investigate participant

186 demographics and characteristics of AI in the different diagnostic groups and controls. There

187 are no clear rules about the acceptable fraction of missing data to justify imputation. As such,

188 we decided on 10%, as a level that would allow imputation, thus enabling us to use as much

189 of the data as possible, whilst also retaining reliability and accuracy (Steyerberg, 2008). As

190 such scores were imputed if the patient had less than 10% missing items. This translates as:

191 AIM: Up to 2 missing values, ALS-18: Up to 2 missing values, ACS: Up to 4 missing values,

192 WHODAS: 1 missing value.

193

194 Two sample t-tests were used to compare means between the cases and controls after

195 verifying that relevant assumptions were valid. Proportions were compared using chi-squared

196 tests. General linear models (GLMs) were used to compare the mean lability (ALS), intensity

197 (AIM), and subjective ability to control affect (ACS) outputs across cases (different

198 diagnostic groups) and the control group. Adjustment was made for age, sex and educational

199 level if necessary. Model assumptions were checked and, in the case of an overall significant

200 difference in mean score across the diagnosis groups, pairwise post-hoc comparisons of

201 adjusted mean scores were performed with a Bonferroni correction. To investigate how far

202 the different aspects of affective instability correlate with each other, the linear association

203 between each pair of measurement scales for the full sample and for the cases only was

204 assessed using Pearson's product moment correlation. Association between each

205 measurement scale and the mood fluctuation rate was assessed using Spearman's rank

206 correlation.

207  
208 Multiple regression was used to examine the association between affective instability  
209 and general assessment of functioning as measured by the WHODAS (Üstün, 2010) in  
210 clinical cases, adjusting for diagnosis and other patient characteristics. A purposeful selection  
211 approach was used to fit the model. Manual backward elimination was first used to remove  
212 variables based on Wald statistics using  $p = 0.05$  as the cut-point for removal. Removed  
213 variables were then re-entered into the model one-by-one to check their significance.  
214 Variables initially considered in the model included: (a) socio-demographics: age, sex,  
215 ethnicity, education level, marital status, (employment status was not considered in the model  
216 since the WHODAS incorporates this parameter in ratings), (b) illness characteristics:  
217 diagnosis, duration of illness, total number of medications, (c) AI measures: mood fluctuation  
218 rate and the mean scores for the ACS, ALS-18 and AIM. All analyses were conducted in  
219 IBM SPSS Statistics 24.

220

## 221 **4. Results**

222

### 223 4.1 Participant characteristics

224

225 The initial dataset comprised 101 participants, but 9 individuals were excluded due to  
226 missing data (3 bipolar, 2 major depression, and 2 controls). Hence the final sample (N=94)  
227 comprised of 69 cases and 25 controls.

228

229 Table 1 describes the socio-demographic characteristics of included participants by  
230 group (case versus control), and diagnostic subgroups (bipolar disorder (n=11), BPD (n=12),  
231 psychosis (n=21), and major depression (n=25)). There was a significant difference between  
232 cases and controls in mean age ( $p=0.001$ ), employment status ( $p=0.001$ ) and marital status  
233 ( $p<0.001$ ). Age was controlled for during regression analysis as AI is influenced by this

234 (Marwaha et al., 2013b). Duration of illness was recorded in the dataset for 67 out of the 69  
235 cases and was positively skewed with the sample having been ill for a median duration of 36  
236 months (interquartile range (IQR) 15-156 months). Across the diagnostic groups, participants  
237 with depression reported the longest duration of illness (median 120 months, IQR 12-258),  
238 followed by participants with BPD (median 36 months, IQR 24-120), psychosis (median 27  
239 months, IQR 20.5-111), and bipolar disorder (median 24 months, IQR 9-36), respectively.

240

241 *Table 1 about here*

242

243 The commonest class of medication prescribed was anti-depressants, and most patients  
244 reported being prescribed one (25%) or two (25%) medications. Seventeen percent of cases  
245 (N=12) were not taking any medication. We did not explore, type of medications and their  
246 impact on our results because of the lack of a robust typology of the effectiveness of  
247 medications indicated for affective instability (Lieb et al., 2010).

248

249 4.2 Comparison of questionnaire scores between cases and controls

250 Differences in the unadjusted mean scores between cases and the controls for all  
251 measures are presented in Table 2. Age and sex were found not to be significant across the  
252 sample in the general linear model (GLM) for the ACS, AIM, and WHODAS scores; whilst  
253 there was trend towards significance for the effect of age on ALS score ( $P = 0.068$ ).

254

255 *Table 2 about here*

256

257 Affect lability (ALS): When adjusted for age, a statistically overall significant  
258 difference was observed in mean ALS-18 scores between cases and controls ( $F(4,88) =$

259 7.195,  $p < 0.001$ ). Post-hoc pairwise comparisons of mean scores revealed significantly lower  
260 mean ALS-18 scores for the control group compared to each diagnosis group but no  
261 significant differences between diagnoses.

262

263 Affect intensity (AIM): There was little difference in the mean AIM scores between  
264 groups, with slightly higher mean scores found for controls compared to cases. These  
265 differences were not statistically significant ( $p = 0.867$ ).

266

267 Ability to control affect (ACS): An overall significant difference was found between  
268 mean ACS scores across the different diagnostic groups, including controls ( $F(4,89) =$   
269  $14.520$ ,  $p < 0.001$ ). Post-hoc pairwise comparisons of the mean scores revealed significantly  
270 higher mean ACS scores (meaning lower control) for each diagnostic group compared to  
271 controls ( $p < 0.05$ ). A significant difference was also found between the mean scores in  
272 borderline personality disorder patients and patients with non-affective psychosis ( $p = 0.010$ ).

273

274 Mood fluctuations in the last week: Table 2 shows the number of participants (i.e.  
275 frequency with percentage) who reported each number of mood fluctuations over the past  
276 week prior to assessment. This revealed that cases tended to have more changes in their mood  
277 state than controls, although no overall differences were found in rate of mood fluctuation  
278 between groups ( $p=0.310$ ). Those with major depression reported the greatest number of  
279 mood fluctuations in the last week, followed by non-affective psychosis, borderline  
280 personality disorder and then bipolar disorder.

281

282 *4.3 Correlations between different components of AI, mood fluctuation rate and functioning*

283 Correlations are shown in table 3. Strong positive correlations were found between  
284 the ALS and the ACS in the full and cases only analysis. Weak to moderate correlations were  
285 found between the AIM and the ALS. When assessing the association between each  
286 measurement scale and mood fluctuation rate ‘last week’, moderate to strong positive  
287 correlations were found between mood fluctuation and ALS and ACS. There was a weak  
288 correlation between AIM and mood fluctuation rate. All correlations were weaker when  
289 focusing on the cases only.

290

291 *Table 3 about here*

292

#### 293 *4.4 AI and functioning*

294

295 In the clinical sample, an overall significant difference was observed between mean  
296 WHODAS scores across the different diagnosis groups,  $F(4,89) = 11.454$ ,  $p < 0.001$  ( $p <$   
297  $0.05$  for bipolar disorder). Post-hoc pairwise comparisons revealed significantly lower mean  
298 WHODAS scores for the control group compared to each diagnosis group, as might be  
299 expected, but differences between diagnostic groups were not significant.

300

301 A multiple regression model investigating factors associated with the WHODAS  
302 score, demonstrated that both ALS-18 and ACS scores were significantly associated with  
303 current level of functioning. After correcting for multicollinearity, ALS-18 score was retained  
304 in the final model ( $\beta=0.845$ ,  $p<0.001$ ), along with the total number of medications  
305 ( $\beta=0.107$ ,  $p<0.046$ ). All other variables considered, including diagnosis, were not  
306 significantly associated with WHODAS score in the final model.

307

## 308 **5. Discussion**

309

### 310 *5.1 Main findings*

311  
312           This is the first study, to our knowledge, that has comprehensively assessed the core  
313 components of affective instability in a trans-diagnostic clinical population and compared  
314 clinical cases with a control group without mental disorder. We found only affective lability  
315 and affective control is significantly different in people with a range of mental disorders in  
316 comparison to those without. No differences were observed between people with and without  
317 mental disorder in the intensity of affect experienced or the rate of mood fluctuation in the  
318 last week. Two of the three components of affective instability (lability and intensity) did not  
319 differ significantly between individuals with different psychiatric diagnoses, although ability  
320 to control affect was significantly different in individuals with BPD in comparison to non-  
321 affective psychosis. Whilst the small numbers within each diagnostic group mean that  
322 interpretation can only be exploratory, contrary to expectation, we found that the greatest  
323 number of mood changes in a week was experienced by people with major depression,  
324 followed by non-affective psychosis, BPD and then bipolar disorder.

325  
326           In terms of the affective instability construct, the strongest inter-correlation was found  
327 between lability and ability to control affect, with much weaker (modest) correlations  
328 between affective intensity and ability to control affect (or lability and control). Finally, only  
329 affective lability, but not affective intensity, ability to control affect or mood fluctuation rate  
330 was associated within functioning independent of diagnosis and other important confounders.

331

### 332 *5.2 Limitations*

333  
334           Our sample size was relatively modest (just under 100). This limited the statistical  
335 power of our analyses and increased the risk that our results might be due to type II error.  
336 This means that comparisons of affective instability between diagnostic groups in particular,

337 should be considered entirely exploratory, and other interpretations tentative. Another caveat  
338 to comparisons between diagnoses is that we did not complete inter-rater reliability  
339 assessments. However, this is the largest study to date exploring our questions.

340

341         Our observations related to affective instability are limited to the four mental  
342 disorders that we sampled. We cannot therefore generalize our findings to other disorders,  
343 where affective instability is known to be important such as OCD (Bowen et al., 2015) or  
344 ADHD (Asherson et al., 2007). Furthermore, we could not take into account the contribution  
345 made by mental or physical comorbidities in our sample. However, given our sample of cases  
346 were those in contact with secondary mental health services there are likely to be high levels  
347 of comorbidity. Therefore, it is possible that high levels of affective lability and problems  
348 with affective control are linked to comorbidity and this should be the focus of future studies.  
349 In our regression modelling we were not able to control for some factors known to impact  
350 functioning such as cognition, illness severity, premorbid functioning and depressive  
351 symptoms.

352

353         The cases sampled were not in an acute illness episode and it is conceivable that this  
354 biased estimate of group difference towards the null, that is, there is no difference between  
355 the cases and controls on affective instability measures. Affect intensity (and possibly  
356 instability) might vary with illness acuity, which might explain why differences between  
357 cases and controls in the present study were smaller than those reported in an in-patient  
358 sample (Henry et al., 2008; Reich et al., 2012). Whilst we did not assess illness severity, we  
359 adjusted for illness duration and number of medications, both of which might be expected to  
360 be associated with illness severity. More specifically, we also did not assess current mood  
361 state using standardised measures and therefore do not know how far the severity of current

362 mood (e.g depth of depression) could have impacted on our results. There is little current  
363 evidence on how far AI changes, as mood becomes lower or more elated to guide how this  
364 could have influenced our main findings. Indeed, in bipolar disorder, AI is found in both  
365 euthymic and periods of acute illness (Harvey, 2008). We explored whether AI is different  
366 between cases and controls. Future studies should also aim to explain the differences between  
367 affective instability in people with mental disorders and without.

368

369         We used assessment measures which require recall of affective experiences. These  
370 may be prone to bias, particularly when compared to ecological momentary assessments  
371 (EMA) (Broome et al., 2015b). How accurately people with mental disorders recall their  
372 affective experiences might differ depending on diagnosis. The ratings themselves at an  
373 individual level may also be dependent on an initial calibration to understand what is meant  
374 by a “marked” shift in mood (Holmes et al., 2016). Therefore, paradoxically individuals with  
375 fewer mood fluctuations may better report retrospective fluctuations as they would have  
376 stood out in their experience, whilst those with more frequent fluctuations may only report  
377 “marked” ones, as small fluctuations were perhaps normalised by their experience. This is  
378 one potential explanation of why depressed patients reported more fluctuations than other  
379 groups, though this was not statistically significant. The question used to assess mood  
380 fluctuation didn’t specify type of affect and therefore could have excluded swings in anger  
381 and irritability, which have been shown to differentiate between diagnosis (Tsanas et al.,  
382 2016). We also recognise that current mood state may have impacted on assessment.

383

384         Whilst momentary assessment of psychopathology appears feasible using  
385 smartphones (Tsanas et al., 2016), it is as yet unclear whether retrospective affective  
386 assessments and EMA relate to the same underlying psychological or biological processes,



387 especially as the former will be subject to important cognitive processes (e.g contextual  
388 processing), which control how mood is experienced (Dubad et al., 2018). There is also the  
389 issue of how far individuals recognize and name affective states in the same way.

390  
391 *5.3 Theoretical and clinical implications*  
392  
393

394 Our findings only partly validated our original definition of affective instability as a  
395 trans-diagnostic parameter incorporating affect lability, ability to control and intensity  
396 (Marwaha et al., 2014). Affect lability and the ability to control these were indeed found to  
397 occur at higher levels than in controls and at similar levels across the different diagnostic  
398 groups. Scores on both measures were also relatively strongly correlated with each other re-  
399 enforcing the notion that they are facets of the same or similar underlying latent construct.  
400 Affective intensity was only relatively weakly associated with other affective instability  
401 measures. Replication in a much larger sample is required to understand how far this pattern  
402 holds true. In the current study affective intensity was no different between cases and controls  
403 or between the cases themselves consistent with previous literature (Henry et al., 2001b).  
404 Whilst caution is necessary in interpretation, this does suggest that intensity of affect may not  
405 be a feature that may help delineate the boundaries of “normal” or “abnormal” affective  
406 experience, or at least in the way that it was measured here. Again, a study with a larger  
407 sample size is required.

408  
409 Mood fluctuation rate (as measured by our bespoke instrument) showed some  
410 concurrent validity with two measures of affective instability, and surprisingly, fluctuation  
411 rate was no different between cases and controls. This may be a function of our sample size,  
412 but this finding should prompt larger studies, with more comprehensive fluctuation change

413 assessments to investigate this area. Crucially, these studies need to include people without  
414 mental disorders as controls.

415

416         We used a comprehensive way to measure affective instability in people with  
417 different diagnoses and the current results as well as previous research provides some  
418 counterbalance to the notion that affective instability is specific to or more severe in people  
419 with bipolar disorder or borderline personality disorder. The challenge now is to understand  
420 whether more subtle differences exist that may be clinically useful, such as whether a  
421 particular valence change is more or less common in different disorders (Reich et al., 2012)  
422 or whether richer, digitally captured mood data is helpful in differentiating disorders. Current  
423 evidence indicates clinicians do not use diagnostic criteria effectively to distinguish disorders  
424 such as BPD and bipolar disorder in which affective instability symptoms are seen to overlap  
425 (Saunders et al., 2015). Further research into common and uncommon valence changes in the  
426 disorders, perhaps incorporating digital mood monitoring, may help to resolve this clinical  
427 difficulty.

428

429         Finally, we demonstrate that affective instability independently adversely impacts  
430 functioning in people with mental disorders, and this is independent of diagnosis. The  
431 measure of functioning that we used suggests the impact could be on multiple domains  
432 including learning new tasks, joining in community activities, day to day work and  
433 maintaining friendships. We have previously found that interpersonal conflict is part of the  
434 pathway from affective instability and incident depression (Marwaha et al., 2015) and the  
435 current study is also consistent with other work highlighting the impact of affective instability  
436 on functioning in bipolar and transdiagnostically (Patel et al., 2015; Strejilevich et al., 2013).

437 We extend these previous findings by identifying that affective lability, as opposed to other  
438 aspects of AI such as ability to control affect or intensity, has the greatest impact.

439

440 As such affective lability has the potential for being a therapeutic target that could  
441 improve functional outcomes in mental disorders. Pharmacological interventions that are  
442 widely used (e.g mood stabilising antipsychotics) and emotional regulation training (Berking  
443 et al., 2008) need more robust trial evidence, but could have a significant impact on distress  
444 and outcomes.

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**Table 1: Descriptive statistics by group; case versus control and diagnosis subgroups (n = 94)**

Characteristic	Control (n = 25)	Case (n = 69)	Cases by diagnosis				Total (n = 94)	Case versus control	
			Bipolar (n = 11)	Borderline (n = 12)	Psychosis (n = 21)	Depression (n = 25)		P value	Test
Age (years); mean (SD)	48.5 (10.8)	38.2 (12.8)	35.6 (13.3)	33.9 (11.0)	34.9 (9.8)	44.2 (14.1)	41.0 (13.1)	0.001	t test
Male; n (%)	9 (36.0%)	36 (52.2%)	6 (54.5%)	2 (16.7%)	12 (57.1%)	16 (64.0%)	45 (47.9%)	0.165	Chi square test
Employment; n (%):									
– Employed	19 (76.0%)	26 (37.7%)	5 (45.5%)	4 (33.3%)	6 (28.6%)	11 (44.0%)	45 (47.9%)	0.001	Chi Square test (employed vs unemployed, n = 90)
– Unemployed	5 (20.0%)	40 (58.0%)	6 (54.5%)	8 (66.7%)	14 (66.7%)	12 (48.0%)	45 (47.9%)		
– Other	1 (4.0%)	3 (4.3%)	0 (0.0%)	0 (0.0%)	1 (4.8%)	2 (8.0%)	4 (4.3%)		
Ethnicity; n (%)									Chi square test
– White British	18 (72.0%)	57 (82.6%)	9 (81.8%)	10 (83.3%)	14 (66.7%)	24 (96.0%)	75 (79.8%)	0.259	
– Other	7 (28.0%)	12 (17.4%)	2 (18.2%)	2 (16.7%)	7 (33.3%)	1 (4.0%)	19 (20.2%)		
Education; n (%)									Chi square test (n = 89)
– None/GCSE	7 (28.0%)	35 (50.7%)	5 (45.5%)	5 (41.7%)	12 (57.1%)	13 (52.0%)	42 (44.7%)	0.061	
– A level	3 (12.0%)	16 (23.2%)	4(36.4%)	2 (16.7%)	5 (23.8%)	5 (20.0%)	19 (20.2%)		
– Degree/higher degree	11 (44.0%)	17 (24.6%)	2 (18.2%)	5 (41.7%)	3 (14.3%)	7 (28.0%)	28 (29.8%)		
– [Missing]	4 (16.0%)	1 (1.4%)	-	-	1 (4.8%)	-	5 (5.3%)		



Marital status; n (%)									Chi square test
– Married/cohabiting	20 (80.0%)	22 (31.9%)	1 (9.1%)	6 (50.0%)	5 (23.8%)	10 (40.0%)	42 (44.7%)	< 0.001	
– Single/separated/ widowed	5 (20.0%)	47 (68.1%)	10 (90.9%)	6 (50.0%)	16 (76.2%)	15 (60.0%)	52 (55.3%)		

**Table 2: Unadjusted mean scores (with standard deviation) and count (%) for each measurement scale and subscales**

Measurement scale	Controls (n = 25)	Cases (n = 69)				All (n = 94)	Case versus control (General linear models, F test)
		Bipolar (n = 11)	Borderline (n = 12)	Psychosis (n = 21)	Depression (n = 25)		
ACS (scale 1-7)	3.36 (0.56)	4.39 (0.66)	5.14 (0.58)	4.23 (0.86)	4.52 (0.86)	4.21 (0.92)	p<0.001
ACS: Anger	3.06 (0.60)	4.31 (0.83)	5.21 (0.96)	4.07 (0.89)	4.08 (1.15)	3.97 (1.11)	
ACS: Positive-affect	3.48 (0.63)	4.17 (0.73)	4.12 (1.02)	4.05 (0.89)	3.87 (1.03)	3.88 (0.89)	
ACS: Depressed	3.36 (0.75)	4.96 (0.88)	5.92 (0.72)	4.40 (1.01)	5.38 (1.04)	4.64 (1.27)	
ACS: Anxiety	3.42 (0.57)	4.31 (0.84)	5.63 (0.60)	4.42 (1.23)	4.93 (0.95)	4.43 (1.13)	
ALS-18 (scale 0-3)	0.64 (0.58)	1.47 (0.62)	1.66 (0.49)	1.53 (0.67)	1.50 (0.62)	1.29 (0.71)	p<0.001 <sup>1</sup>
ALS-18: Anxiety/Depression	0.55 (0.65)	1.49 (0.69)	2.23 (0.79)	1.60 (1.01)	1.88 (0.87)	1.46 (1.00)	
ALS-18: Depression/Elation	0.86 (0.70)	1.66 (0.57)	1.45 (0.58)	1.65 (0.73)	1.52 (0.53)	1.38 (0.70)	
ALS-18: Anger	0.38 (0.48)	1.16 (0.95)	1.42 (0.96)	1.27 (0.88)	1.08 (1.03)	0.99 (0.92)	
AIM (scale 1-6)	3.50 (0.48)	3.45 (0.37)	3.37 (0.44)	3.42 (0.56)	3.37 (0.39)	3.42 (0.45)	p=0.867
Number of mood fluctuations reported in the last week							P=0.310
0	13 (52%)	2 (18.2%)	0 (0%)	5(23.8%)	2 (8%)	22 (23.4%)	
1-3	10 (40%)	5 (45.5%)	3 (25%)	4 (19%)	8 (32%)	30 (31.9%)	
4-7	2 (8%)	3 (27.3%)	7 (58.3%)	7 (33.3%)	7 (28%)	26 (27.7%)	
>7	0 (0%)	1 (9.1%)	2 (16.7%)	5 (23.8%)	8 (32%)	16 (17.0%)	
WHODAS <sup>2</sup> (scale 0-4)	0.54 (0.11)	1.43 (0.61)	1.83 (0.69)	1.75 (1.00)	1.89 (0.91)	1.44 (0.96)	p<0.001

<sup>1</sup> Adjusted for age.

**Table 3: Correlation coefficients between each pair of measurement scales**

<b>Full -sample (N=94)</b>					
		<b>AIM</b>	<b>ALS-18</b>	<b>ACS</b>	<b>Mood fluctuation (last week)</b>
<b>AIM</b>		1	0.210	0.188	0.12
<b>ALS-18</b>			1	0.776	0.61
<b>ACS</b>				1	0.53
<b>Mood fluctuation (last week)</b>					1
<b>Cases only (N=69)</b>					
		<b>AIM</b>	<b>ALS-18</b>	<b>ACS</b>	<b>Mood fluctuation (last week)</b>
<b>AIM</b>		1	0.322	0.265	0.157
<b>ALS-18</b>			1	0.666	0.45
<b>ACS</b>				1	0.29
<b>Mood fluctuation (last week)</b>					1