INTRODUCTION

Bipolar spectrum disorders are characterized by extreme emotional highs and lows while unipolar depressive disorders share only the lows. Although many studies have reported transdiagnostic deficits across both disorders in domains like threat-related processing, executive control, and working memory (see Nusslock & Alloy, 2017, for review), identifying markers of differential risk that can separate the two can improve early detection and facilitate early intervention. For example, bipolar patients typically present to clinicians when they are depressed (Hirschfeld, Cass, Holt, & Carlson, 2005), leading to a common misdiagnosis of major depressive disorder up to 66% of the time (Hirschfeld et al., 2003). Cardinal

Hypomania and depression associated with distinct neural activity for immediate and future rewards

James E. Glazer1 | Nicholas J. Kelley1 | Narun Pornpattananangkul2 | Robin Nusslock1

1Department of Psychology, Northwestern University, Evanston, Illinois
2Emotion and Development Branch, U.S. National Institute of Mental Health, Bethesda, Maryland

Correspondence
James E. Glazer, Department of Psychology, Northwestern University, 2029 Sheridan Roan, Evanston, IL 60208.
Email: glazerja@u.northwestern.edu

Funding information
National Science Foundation, Grant/Award Number: DGE-2016231981 to J.E.G

Abstract
Bipolar spectrum and unipolar depressive disorders have been associated with distinct and opposite profiles of reward-related neural activity. These opposite profiles may reflect a differential preexisting vulnerability for both types of disorders. In support, recent ERP studies find that, following reward feedback, a larger reward positivity (RewP) is associated with greater vulnerability for bipolar spectrum disorders, whereas a smaller RewP is associated with greater vulnerability for depression. However, prior studies have investigated only immediate rewards and have not examined dimensions of both bipolar disorder and unipolar depression within the same sample. The present study is the first to investigate feedback-related ERP correlates of proneness to hypomania and unipolar depressive tendencies within the same sample and to expand our scope to include future rewards. Participants completed a modified time estimation task where the same monetary reward was available immediately or at one of five different future dates. Results revealed proneness to hypomania and unipolar depressive tendencies were related to an elevated and blunted RewP, respectively, but only following immediate rewards (i.e., today). Following rewards in the distant future (e.g., 8 months), proneness to hypomania and depressive tendencies were associated with elevated and blunted amplitudes for the P3, respectively, a subsequent ERP component reflecting motivational salience during extended feedback processing. Furthermore, these opposing profiles were independent of, and significantly different from, one another. These results suggest that feedback-related ERPs following immediate and future rewards are candidate biomarkers that can physiologically separate vulnerability for bipolar spectrum from unipolar depressive disorders.

KEYWORDS
bipolar disorder, depression, ERPs, individual differences, motivation
features of bipolar spectrum disorders absent in unipolar depression are manic and hypomanic symptoms (see Urošević, Abramson, Harmon-Jones, & Alloy, 2008, for review) that manifest as increased approach motivation, goal pursuit, and elevated mood (American Psychiatric Association, 2013). These symptoms fall on a continuum of severity that extends to the general population (Angst et al., 2003) where individuals with subthreshold symptoms may express increased vulnerability for developing bipolar or unipolar depressive disorders (Cuijpers & Smit, 2004; Kwapil et al., 2000).

1.1 | Distinct profiles of reward sensitivity

Prevailing theories suggest that distinct and opposite profiles of reward sensitivity constitute a preexisting risk factor for, rather than a consequence of, bipolar spectrum and unipolar depressive disorders (see Alloy, Olino, Freed, & Nusslock, 2016, for review). Reward sensitivity relates to the value an individual places on rewards, the perceived expectation of reward receipt, and the mechanisms by which rewards are processed. Numerous neuroimaging studies have linked vulnerability for bipolar spectrum and unipolar depressive disorders to respective elevated and blunted profiles of reward-related neural activity (see Nusslock & Alloy, 2017). These differential profiles are driven in part by abnormal phasic dopamine signaling (see Whitton, Treadway, & Pizzagalli, 2015, for review) among frontostriatal neural pathways implicated in reward processing (Berridge, Robinson, & Aldridge, 2009; Haber & Knutson, 2010). EEG studies utilizing ERPs have linked individual differences in reward sensitivity to modulation in an ERP known as the reward positivity (RewP: Bress & Hajcak, 2013; Holroyd, Pakzad-Vaesi, & Krigolson, 2008; Lange, Leue, & Beauducel, 2012; Threadgill & Gable, 2016). The RewP is a positive ERP deflection elicited approximately 300 ms following positive (vs. negative) reward feedback and has been linked to covariation within the frontostriatal circuit, including the ventral striatum (Becker, Nitsch, Miltner, & Straube, 2014; Carlson, Foti, Mujica-Parodi, Harmon-Jones, & Hajcak, 2011; Foti, Weinberg, Dien, & Hajcak, 2011). Reinforcement learning theories argue that the RewP may be associated with a phasic increase in mesencephalic dopamine signaling within the frontostriatal pathway, called a reward prediction error, that tracks violations in reward expectation between actual and predicted outcomes (Holroyd & Coles, 2002; Montague, Hyman, & Cohen, 2004; Schultz, 2002).

Recent research has found a blunted RewP amplitude in individuals with a family history of unipolar depression (Kujawa, Proudfit, & Klein, 2014), which may prospectively predict risk for depressive disorders (Bress, Foti, Kotov, Klein, & Hajcak, 2013; Bress, Meyer, & Proudfit, 2015). Although less research has investigated the relationship between the RewP and bipolar spectrum symptoms, two recent studies suggest proneness to hypomania may be related to an increased RewP positivity (Mason, O’Sullivan, Bentall, & El-Deredy, 2012; Mason, O’Sullivan, Blackburn, Bentall, & El-Deredy, 2012). However, no prior study has investigated ERP correlates of hypomanic and depressive tendencies within the same sample. While each disorder displays opposite profiles of reward-related neural activity, hypomanic and unipolar depressive symptom profiles likely represent separable, rather than opposite, psychometric dimensions (Johnson et al., 2011; Solomon et al., 2003; Youngstrom, Murray, Johnson, & Findling, 2013). In this way, the same individual can score high or low on one or both dimensions. One of the primary goals of the present study was to, within the same sample, test the hypothesis that tendencies toward unipolar depression versus hypomania are characterized by a distinct and opposite RewP amplitude.

1.2 | Future rewards

However, most prior neuroscientific research investigating reward processing has focused on immediate rewards. Real-world rewards are often delayed over long periods of time, and it remains unknown whether neural activity elicited by future rewards may be related to hypomanic or depressive symptoms. Affective forecasting research has shown that, as rewards approach the future, emotional reactions become less tangible and more difficult to predict (Rick & Loewenstein, 2008; Van Boven, White, & Huber, 2009). As a result, experiences following rewards in the present are used as an affective reference point to signal the value of future rewards (Gilbert & Wilson, 2007; Rick & Loewenstein, 2008). This line of research suggests that opposite profiles of reward sensitivity in the present among individuals prone to either hypomania versus unipolar depression may facilitate opposite profiles of neural activation for future rewards as well.

Prior studies have found that future rewards recruit distinct neural regions relative to their immediate counterparts (Ballard & Knutson, 2009; McClure, Laibson, Loewenstein, & Cohen, 2004). For example, immediate rewards have been associated with the midbrain dopamine system, whereas future rewards have been associated with frontoparietal regions relevant for abstract mental representations and extended cognitive processing in McClure et al. (2004). Therefore, for more distant future rewards, we predicted that hypomanic and depressive symptoms would be associated with respective increases and decreases in two later ERP components linked to extended feedback-related processing: the P3 and late positive potential (LPP). First, the P3 is a positive ERP component reflecting enhanced incentive salience of the feedback stimulus (Yeung & Sanfey, 2004). Prior work has linked the P3 to motivational salience and individual differences in reward sensitivity (Van den Berg, Franken, & Muris, 2011). Second, directly following
the P3, the LPP reflects extended cognitive and attentional processing (Althaus et al., 2010; Groen, Tucha, Wijers, & Althaus, 2013; Pornpattananangkul & Nusslock, 2015; Van Meel, Heslenfeld, oosterlaan, Luman, & Sergeant, 2011) and has been referred to as the “emotional counterpart” of the P3 (Groen et al., 2008).

The present study is the first to investigate feedback-related ERP correlates of proneness to hypomania and unipolar depressive tendencies within the same sample and to expand our scope to include future rewards. We recruited participants from the undergraduate Northwestern community who completed a time estimation task where the same reward (e.g., $10) was available today or at one of five future delays (see Figure 1). First, following immediate outcomes, we predicted that proneness to hypomania and depressive tendencies would be related to an elevated and blunted RewP, respectively, consistent with prior studies (Mason, O’Sullivan, Bentall, et al., 2012; Proudfit, 2015). Second, a recent study reported no relationship between the P3 and vulnerability for hypomania for future rewards up to 1 month in the future (Mason, O’Sullivan, Blackburn et al., 2012). Prior research indicates that rewards in the near future may not be distant enough to require extended feedback processing (Freitas, Salovey, & Liberman, 2001; Gilbert & Wilson, 2007; McClure et al., 2004; Rick & Loewenstein, 2008; Trope & Liberman, 2003). Therefore, only following feedback in the 8-month condition did we predict that proneness to hypomania would be associated with increases in the P3 and LPP, while depressive tendencies would be related to reductions in both components. Finally, we did not predict that either symptom dimension would be related to ERPs in any of the intermediate delay conditions.

2 | METHOD

2.1 | Participants

Fifty-eight right-handed, medication-free, native English speakers (28 female) reported individually to a study described as an investigation of how people process words, sentences, and pictures. All participants received credit toward a course requirement for their participation but were also told that they could win money based on their performance. Participants’ ages ranged from 18 to 22 ($M = 18.68$, $SD = 0.91$). Participants were excluded for excessively noisy EEG data (quantified using visual inspection, $N = 5$), excessive EEG artifact rejection (greater than 50% for any feedback condition, $N = 3$), and not looking at the screen during the time estimation task ($N = 2$). One additional participant was excluded because they did not believe the experiment would result in real money. After these exclusions, 47 participants (25 female) remained for analysis. This study was approved by the Northwestern University Internal Review Board.

2.2 | Procedure

Following informed consent, participants completed self-report questionnaires assessing proneness to hypomania and unipolar depression. Next, participants completed a short behavioral task as part of a larger study whose results were not investigated in the current study. After questionnaires, participants were set up with EEG and completed a modified time estimation task where, on each trial, they could win the same amount of money (e.g., $10) either today or at one of five future delays (Miltner, Braun, & Coles, 1997). Following this

![Figure 1](image-url)  
**Figure 1** Trial structure for the modified time estimation task. Before each trial, a fixation cross was displayed during a jittered time window. On each trial, one of six cues was presented indicating the time of reward receipt (e.g., today, 2 days, 2 weeks, 1 month, 3 months, or 8 months). Cues were displayed until a response was received and then became bold and italicized. After 2,000 ms, reward feedback indicated a gain (Win) or a no-gain (Fast or Slow).
task, participants were unhooked from EEG, compensated, and debriefed.

2.3 | Self-report measures

Individual differences in proneness to hypomania were measured using the Hypomanic Personality Scale (HPS: Eckblad & Chapman, 1986). The HPS was developed to identify individuals at risk for bipolar disorder, and increased scores indicate a greater presence of hypomaniac personality traits, such as elevated mood and racing thoughts. Increased hypomaniac personality scores have been shown to prospectively predict bipolar disorder onset over a 10-year follow-up period (Kwapil et al., 2000) and has been linked to elevated reward-related activation in electrophysiological and neuroimaging studies (Damme, Young, & Nusslock, 2017; Peterson, Gable, & Harmon-Jones, 2008; Pornpattananangkul & Nusslock, 2015). This questionnaire includes 48 true or false questions that measure hypomaniac personality traits (e.g., “I often feel excited and happy for no apparent reason” and “I frequently find that my thoughts are racing”) and is scored by adding up the number of true statements. To assess proneness to unipolar depressive symptoms, we utilized the 7 Down questionnaire from the 7 Up 7 Down Inventory (Youngstrom et al., 2013), a subscale used to measure traitlike depressive tendencies. Higher scores on the 7 Down indicate a greater presence of depressive symptoms, such as worthlessness and hopelessness. This seven-item questionnaire was developed from the Generalized Behavioral Inventory to psychometrically separate unipolar depressive from hypomaniac symptoms (Depue et al., 1981; Youngstrom et al., 2013). Participants are instructed to answer how often they experience a list of depression symptoms on a scale from 0 (never, or hardly ever) to 3 (very often or almost constantly), for example, “Have you had periods when it seemed that the future was hopeless and things could not improve?” Importantly, both HPS and 7 Down are trait measures of propensity to hypomaniac symptoms and depressive tendencies, respectively, and represent separable psychometric dimensions (Johnson et al., 2011; Solomon et al., 2003; Youngstrom et al., 2013). Mean scores were used to quantify the totals for both questionnaires. Specifically, HPS was scored as a value from 0 to 1 where larger values indicate higher proneness to hypomania ($M = 0.47, SE = 0.14, \alpha = 0.76$), and 7 Down was scored as a value from 0 to 3 where larger values indicate higher presence of depressive tendencies ($M = 0.52, SE = 0.08, \alpha = 0.84$).

2.4 | Time estimation task

To assess individual differences in reward-related EEG activity, participants completed a modified time estimation task (Damen & Brunia, 1987; Kotani et al., 2003; Miltner et al., 1997; see Figure 1) programmed using E-Prime while EEG was recorded. Specifically, participants were instructed to press a button exactly 3 s following the presentation of a cue and received feedback after every trial. Each trial began with a fixation cross presented for a jittered length of time randomly chosen between 750 and 1,250 ms, followed by a text cue indicating when participants could win money (today, 2 days, 2 weeks, 1 month, 3 months, and 8 months). Following cue presentation, participants estimated 3 s as closely as possible. Once the participant estimated that 3 s had elapsed, they were instructed to press a button with their right index finger on a response box. The cue remained on the screen unchanged for the duration of the time estimation window. Once the button was pressed, the cue text turned bold-italicized and remained on the screen for 2,000 ms to ensure participants remained aware of the condition of that particular trial prior to feedback presentation (e.g., the date at which they can potentially win money). Finally, positive (Win) and negative (Fast or Slow) feedback was presented for a total of 1,000 ms.

Following prior studies (Kotani et al., 2003; Ohgami et al., 2006), an adaptive algorithm was used to keep gains and no-gains constant at approximately 50%. This method was chosen to limit variation in feedback probability between gains and no-gains that has been found to modulate feedback-related ERPs (Holroyd & Coles, 2002). Specifically, accurate responses were defined as button presses within a sliding time window that adaptively shortened or lengthened by 10 ms following correct or incorrect responses, respectively. The time window began at ±200 ms centered around 3 s (e.g., 2,800–3,200 ms). Participants first completed a practice set of 30 trials to familiarize themselves with the task. The practice block contained five trials of each cue presented in a random order. In addition, response times from the practice block were used to tailor the appropriate time window for accurate responses to be used at the start of the first block. Before and after practicing, participants were permitted to press a button and hear a beep sound exactly 3 s later as many times as they desired to familiarize themselves with the task. Instructions indicated that gains resulted in $10 while fast or slow feedback carried no monetary gain or loss. A constant magnitude of $10 per trial was chosen to keep participants motivated on every trial. Participants were told that, following the experiment, four trials would be randomly chosen to determine their total earnings. If a gain trial was chosen, the participant was instructed they would receive $10 for that trial at the delay given on that trial, while if a no-gain trial was chosen, the participant would not gain or lose any money. However, in the interest of fairness, all participants received exactly $20 in cash following the experiment and were thoroughly debriefed.
2.5 | Electrophysiological recording

Continuous EEG data were recorded during the time estimation task using NeuroScan amplifiers (DC to 100 Hz online, NeuroScan Inc.) within an electromagnetic-shielded booth. Fifty-eight passive Ag/AgCl scalp electrodes were used (following the International 10–20 standard; Jasper, 1958) with four additional placed above and below the left eye, and beside both eyes. A nylon cap with conductive gel was used. EEG data were digitized at a 500 Hz sampling rate, online referenced to the left mastoid, and re-referenced offline to average of both mastoids. Impedance was kept below 5 and 10 kΩ for the scalp and eye electrodes, respectively. Offline, all EEG processing was done using EEGLAB (Delorme & Makeig, 2004) and ERPLAB (Lopez-Calderon & Luck, 2014) in MATLAB (Mathworks, 2017b). Data were resampled at 250 Hz and high-pass filtered at 0.01 Hz. Large scalp artifacts were rejected using continuous automated artifact detection, and noisy channels were identified by excessive kurtosis and interpolated. Next, independent components analysis was carried out to correct for ocular/muscular artifactual components. Once artifactual independent components were removed, data were low-pass filtered at 30 Hz, and 1,100 ms epochs were extracted for feedback-locked activity. These epochs were then de-trended, baseline-corrected at 100 ms prior to feedback, and rejected if containing artifacts exceeding ±75 μV. In addition, epochs associated with response times greater than 5,000 ms or less than 1,000 ms were removed to control for large outlier responses with approximately 1% (~2.5 trials) of the epochs removed per participant. After artifact rejection, the average number of trials for gains and no-gains in each delay condition was approximately equal (gains: M = 24.8, SE = 0.21; no-gains: M = 24.27, SE = 0.18).

Finally, single-trial EEG epochs were averaged separately for all 12 feedback conditions to extract time-locked ERPs. To isolate reward-related neural activity specific to gaining rewards (vs. no-gain outcomes), we utilized a difference wave approach by subtracting gains from no-gains. Difference waves are useful because they avoid ongoing neural activity that does not differentiate between gains and no-gains and may not be specific to reward-related modulation (Luck, 2014). Following prior studies, the RewP was measured as the mean activity from 250 to 350 ms following feedback onset at electrode site Cz, consistent with visual inspection of the grand average difference waveform (Foti et al., 2011; Threadgill & Gable, 2016). The P3 was measured as the mean activity between 400 and 600 ms after feedback onset at site Pz where positivity was maximal. Finally, from visual inspection, the LPP was quantified at electrode site Pz as the mean activity from 600 to 800 ms time-locked to feedback onset. This time window was chosen for the LPP as the end of the P3 component (i.e., 600 ms postfeedback) to the point where the difference between gains and no-gains approached baseline (i.e., 800 ms postfeedback), consistent with prior studies (Althaus et al., 2010; Groen et al., 2013). RewP difference waves were calculated by subtracting no-gains from gains while P3 and LPP difference waves were calculated by subtracting gains from no-gains.

3 | RESULTS

3.1 | Behavioral variables

Accuracy was quantified as the total number of gains divided by the total number of trials and confirmed that our adaptive algorithm kept gains and no-gains at approximately 50% (M = 50.1%, SE = 0.002). Response times were calculated separately for each delay condition. First, trials with response times greater than 5,000 ms or less than 1,000 ms were removed to control for large outlier responses, with an average of less than 1% of all trials (~2.5 trials) removed per participant. Second, all response times were calculated as the absolute value of the difference between the total response time and 3,000 ms to capture how close each response was to the 3,000-ms target response time. Response times for gains (M = 99.33, SE = 4.49) and no-gains (M = 383.19, SE = 17.62) were entered into a 2 (Outcome) × 6 (Delay) repeated measures analysis of variance (ANOVA). Results revealed a significant main effect of outcome, F(1, 46) = 456.78, p < 0.001, η² = 0.91, and an Outcome ×Delay interaction, F(5, 230) = 2.86, p = 0.020, η² = 0.06, but no main effect of delay (p > 0.200). Follow-up t tests revealed that response times for gains were significantly closer to the target of 3,000 ms than response times for no-gains, ts(46) > 18.03, ps < 0.001. To unpack the interaction, response times for gains were subtracted from no-gains at each delay to attain a difference score reflecting the relative improvement in response time (i.e., response times closer to 3,000 ms) on gain trials over no-gain trials. Follow-up t tests on response time difference scores revealed that response times for immediate gains (vs. no-gains) were significantly more accurate than every delayed reward, ts(46) < −2.12, ps < 0.034, except nonsignificantly different from 2 days (p > 0.430).

3.2 | ERPs

We conducted a series of 2 (Outcome: gain vs. no-gain) × 6 (Delay: today, 2 days, 2 weeks, 1 month, 3 months, 8 months) repeated measures ANOVAs on the RewP, P3, and LPP (see Figures 2 and 3, and online supporting information Table S1 and Figure S1). For the RewP, results revealed a significant effect of outcome, F(1, 46) = 53.88, p < 0.001, η² = 0.54, with t tests indicating that gains were significantly more positive than no-gains for all delays, ts(46) > 3.81, ps < 0.001. A main effect of delay was also significant,
F(5, 230) = 19.41, p < 0.001, $\eta^2_p = 0.30$, with $t$ tests showing that the average RewP voltage on immediate outcomes was significantly more positive than all future outcomes, $t(46) > 5.69, ps < 0.001$. In addition, paired $t$ tests between future outcomes revealed several significant effects: First, 2 days was significantly more positive than 2 weeks, $t(46) = 2.02, p = 0.049$; 1 month, $t(46) = 2.31, p = 0.025$; and 3 months, $t(46) = 4.11, p < 0.001$. Second, 2 weeks was significantly more positive than 3 months, $t(46) = 2.10, p = 0.041$; and third, 1 month was significantly more positive than 3 months, $t(46) = 2.09, p = 0.042$. No other significant effects emerged for the RewP ($ps > 0.060$).

For the P3, a significant effect of outcome emerged, $F(1, 46) = 33.08, p < 0.001, \eta^2_p = 0.42$, with $t$ tests revealing that gains were significantly less positive than no-gains for all outcomes, $t(46) < -2.14 = 9, ps < 0.034$. A significant effect of delay also emerged, $F(5, 230) = 9.31, p < 0.001, \eta^2_p = 0.17$, with $t$ tests showing that immediate outcomes were significantly more positive than all future outcomes, $t(46) > 3.71, ps < 0.001$. Finally, the Outcome × Delay interaction was also significant, $F(5, 230) = 3.10, p = 0.013, \eta^2_p = 0.06$. Follow-up $t$ tests performed on difference waves at each paired delay revealed that immediate outcomes were significantly larger than all future outcomes, $t(46) > 2.14, ps < 0.038$, although this effect only approached significance at 2 days ($p = 0.066$). No other significant effects emerged for the P3 ($ps > 0.060$). For the LPP, a significant effect of outcome was found, $F(1, 46) = 34.42, p < 0.001, \eta^2_p = 0.43$, with $t$ tests showing that gains were significantly less positive than no-gains, $t(46) < -2.12, ps < 0.040$. In addition, a main effect of reward delay was also significant, $F(5, 230) = 6.78, p < 0.001, \eta^2_p = 0.13$, with $t$ tests showing
that immediate outcomes were significantly more positive than all future outcomes, \( t_s(46) > 3.22, ps < 0.010 \). No further significant associations emerged for the LPP \( (ps > 0.100) \).

### 3.3 Associations between ERPs, self-report, and behavior

First, and in line with expectation, we confirmed that self-report scores for HPS and 7 Down were not significantly related to one another \( (r = 0.13, p = 0.37) \). Next, we examined the Pearson correlations between self-report measures (i.e., HPS and 7 Down) and each ERP difference wave (i.e., the RewP, P3, and LPP) at each delay (see Figure 4 and Table S2). The RewP following immediate outcomes showed a significant correlation with HPS \( (r = 0.30, p = 0.043) \) and a significant but opposite correlation with 7 Down \( (r = -0.313, p = 0.032) \). Furthermore, the difference between both these opposite correlations was significant \( (z = 2.91, p < 0.010) \). The P3 also significantly correlated with HPS \( (r = 0.31, p = 0.037) \) and, in the opposite direction, 7 Down \( (r = -0.32, p = 0.031) \), but only for the 8-month delay. Again, the difference in correlation at 8 months between HPS and 7 Down for the P3 difference wave was significant \( (z = 3.01, p < 0.010) \). However, the LPP following 8 months was only significantly correlated with HPS \( (r = 0.35, p = 0.017) \), but not with 7 Down \( (r = -0.24, p = 0.111) \), although the correlational difference between HPS and 7 Down was again significant \( (z = 2.82, p < 0.010) \). There were no other significant correlations between ERPs and either HPS or 7 Down \( (ps > 0.050) \), although the association between 7 Down and the RewP at the 2-week delay approached significance \( (r = 0.28, p = 0.053; \text{see Table S2}) \).

Finally, we investigated the relationships between HPS and 7 Down with behavioral variables. Neither self-report...
scale was related to accuracy \((p_s > 0.300)\); however, 7 Down was significantly associated with worse response times for no-gains, relative to gains, following 8-month cues \((r = 0.35, p = 0.017)\). No other significant associations emerged \((p_s > 0.060)\).

### 3.4 Regression analyses

To test the independence of correlations between ERP difference waves and questionnaire measures, linear regressions were performed with the inclusion of both HPS and 7 Down within the same model. In each regression, mean-centered scores for HPS and 7 Down were entered as predictors of our specific hypotheses: (a) the RewP following immediate outcomes, and (b) the P3 and LPP following 8-month outcomes. Results revealed significant independent effects for both HPS and 7 Down in predicting the RewP \((HPS: \beta = 0.35, p = 0.014; 7 \text{ Down: } \beta = -0.36, p = 0.011)\), P3 \((HPS: \beta = 0.35, p = 0.012; 7 \text{ Down: } \beta = -0.36, p = 0.010)\), and LPP \((HPS: \beta = 0.39, p < 0.010; 7 \text{ Down: } \beta = -0.29, p = 0.040)\). These results indicate that, for each ERP component, the unique associations between proneness to hypomania and depressive tendencies were independent of one another. In addition, we performed identical regression analyses for each ERP difference wave separately at every other delay. Of interest, 7 Down significantly predicted the RewP following 2-week outcomes \((\beta = 0.311, p = 0.034)\), but HPS did not \((\beta = -0.20, p = 0.170)\). No other significant effects emerged \((ps > 0.100, \text{ see Table S2})\).

## 4 DISCUSSION

The present study is the first to investigate multiple feedback-related ERP correlates of hypomanic and depressive symptoms within the same sample and to expand our scope to include future rewards. First, our results offer novel insights into the electrophysiological correlates of reward processing and how these are modulated by future rewards. The RewP, P3, and LPP were all modulated by reward delay and displayed increased positivity for immediate over future rewards. However, only the P3 showed a significant Outcome ×Delay interaction such that immediate rewards generated larger P3 difference waves than future rewards. Second, we found that proneness to hypomania and depressive tendencies were associated with opposite profiles of feedback-related ERPs for immediate and distant future rewards. Proneness to hypomania and depressive tendencies were related to an increased and decreased RewP difference wave, respectively, but only following immediate rewards (delivered today). In contrast, proneness to hypomania and depressive tendencies were related to an elevated and reduced P3 difference wave, respectively, but only following distant future rewards (delivered in 8 months). Importantly, proneness to hypomania and depressive tendencies were unrelated to each other, and their associations with ERP components remained significant while controlling for the opposing symptom dimension, suggesting that these relationships are independent of one another.
4.1 ERP correlates of immediate and future rewards

Our results shed new light on the electrophysiological correlates of reward processing following feedback for immediate and future rewards. Consistent with prior studies, the RewP and P3 displayed greater average positivity for immediate over future outcomes (Blackburn et al., 2012; Cherniak et al., 2013; however, see Qu, Huang, Wang, & Huang, 2013), a pattern that we show for the first time also extended to the LPP (see Figure 2). These results suggest that immediate reward feedback was evaluated as “better than expected,” elicited enhanced motivational salience, and displayed extended attentional and affective processing compared to their future counterparts. Interestingly, the RewP tended to display greater positivity for sooner (vs. later) rewards, while the P3 and LPP did not show any significant differences between future outcomes, suggesting that positivity in the RewP time window may in part track reward delay.

In addition, all three components were modulated by feedback valence following immediate and future rewards (see Figure 2); whereas the RewP was greater for gains, the P3 and LPP were greater for no-gains. These results support prior studies suggesting that the LPP following reward feedback may reflect a “negativity bias” where negative feedback elicits greater extended affective and attentional processing than positive feedback (Smith, Cacioppo, Larsen, & Chartrand, 2003). We show for the first time that this negativity bias extended to future rewards. In addition, although many studies have not found that the P3 is sensitive to valence (Yeung & Sanfey, 2004), and some found that positive feedback was more positive than negative feedback (Bellebaum & Daum, 2008; Hajcak, Moser, Holroyd, & Simons, 2007; Zhou, Yu, & Zhou, 2010), other studies reported that negative (vs. positive) feedback was more positive when tasks implement feedback that contains performance information (Chase, Swainson, Durham, Benham, & Cools, 2011; Cohen & Ranganath, 2007; Frank, Worch, & Curran, 2005). In the present study, no-gains may have generated greater P3 amplitudes than gains to motivate more accurate responding on subsequent trials and maximize upcoming rewards.

For ERP difference waves, we report for the first time that the P3 difference wave was significantly modulated by reward delay, whereas the RewP and LPP difference waves were not (see Figure 2). Specifically, the P3 difference wave following immediate rewards was significantly greater than delayed outcomes later than 2 days in the future. An enhanced P3 for no-gains over gains likely reflects an increased motivational signal to facilitate subsequent behavioral adjustments following no-gains to maximize upcoming rewards (Chase et al., 2011; Cohen & Ranganath, 2007; Frank et al., 2005). This finding indicates that the P3 may track differences in reward delay such that the motivational salience of no-gains (vs. gains) is reduced for future rewards in comparison to their immediate counterparts. These results are consistent with a recent electrophysiological study that suggested it is the motivational salience of immediate rewards, rather than their hedonic impact, that biases the motivational value of future rewards during delay discounting (Blackburn et al., 2012).

4.2 Distinct profiles of RewP amplitude

Our results also support models of reward sensitivity that posit that differential profiles of reward-related neural activity may predict clinical risk for bipolar spectrum versus unipolar depressive disorders (see Nusslock & Alloy, 2017, for review). Despite numerous studies that separately investigated relationships between neural activity and vulnerability for hypomania and depression (see Alloy et al., 2016, for review), the present results are the first to report that hypomanic and depressive symptoms are associated with differential profiles of feedback-related ERPs within the same sample (see Figures 3, 4) and to show that these associations are independent of, and significantly different from, one another (see Figure 5). Although these measures are associated with opposite profiles of reward-related neural activity, previous research indicates that hypomanic and depressive symptoms are independent constructs and represent separable, rather than opposing, psychometric dimensions (Johnson et al., 2011; Solomon et al., 2003; Youngstrom et al., 2013). Our results suggest that feedback-related ERPs may independently identify risk for bipolar spectrum and unipolar depressive disorders within the same individuals. These findings not only have implications for understanding the differential pathophysiology underlying bipolar spectrum and unipolar depressive disorders, but also may help identify electrocortical biomarkers that could facilitate more accurate and timely diagnoses.

As expected, depressive tendencies were related to a blunted RewP difference wave following immediate outcomes (see Figures 3, 4). This finding supports emerging research suggesting that reduced RewP amplitude may constitute a potential biomarker of risk for depressive disorders (Bress et al., 2013; Bress et al., 2015; see Proudfit, 2015, for review). Surprisingly, regression analyses showed that depressive tendencies were also significantly related to a blunted RewP difference wave following 2-week outcomes (although correlational results only approached significance), suggesting rewards available in the near future may also elicit blunted neural responses to feedback. In contrast, proneness to hypomania was associated with an elevated RewP difference wave following immediate outcomes, converging with a prior study that found more positive RewP amplitudes following gains and losses in hypomania-prone individuals during a reinforcement learning task (Mason, O’Sullivan, Bentall, et al., 2012). These results are consistent with recent neuroimaging
Evidence that vulnerability for hypomania and unipolar depressive disorder are related to opposite profiles of reward prediction error (Greenberg et al., 2015; Kumar et al., 2008; O’Sullivan, Szczepanowski, El-Deredy, Mason, & Bentall, 2011). Consistently elevated or blunted reward prediction errors in the present may convey a teaching signal that biases the predicted value of upcoming rewards and alters their motivational salience. For example, elevated reward prediction errors among those prone to hypomania have been linked to an expectancy bias toward positive outcomes that fails to update future expectations accordingly (Eisner, Johnson, & Carver, 2008; Johnson, Ruggero, & Carver, 2005; Mason, O’Sullivan, Bentall, et al., 2012; O’Sullivan et al., 2011) and has prospectively predicted increases in hypomanic symptoms (Stange et al., 2012). In contrast, blunted reward predictions errors among individuals with higher levels of...
unipolar depressive symptoms have been linked to an inability to effectively integrate reward feedback and alter behavior to maximize future rewards (Chase et al., 2010; Greenberg et al., 2015; Huys, Vogelstein, & Dayan, 2009). Our results suggest that this blunted reward prediction error may extend to rewards available in the near future as well. Together, these results suggest that abnormal reward prediction errors in the present may index a potential biomarker of differential risk for bipolar spectrum and depressive disorders.

### 4.3 Distinct profiles of P3 and LPP amplitude

Proneness to hypomania and depressive tendencies were also associated with an elevated and blunted P3 difference wave (see Figures 3, 4), respectively, but only following the most distant possible future reward (e.g., 8 months). Importantly, these associations were independent of each other and significantly different from one another (see Figure 5). These results suggest that opposite profiles of reward-related neural activity associated with proneness to hypomania and depressive tendencies emerge for future rewards, but disrupt extended feedback evaluation, indexed by the P3, rather than reward prediction error, indexed by RewP. Whereas proneness to hypomania was associated with increased motivational salience for no-gains (vs. gains) following 8-month outcomes, depressive tendencies were associated with the opposite pattern. Increased motivated salience following no-gains (vs. gains) among those with elevated hypomanic symptoms may facilitate enhanced goal pursuit for especially distant future rewards. On the other hand, depressive symptoms were associated with reduced salience of rewards 8 months in the future and may decrease motivation to pursue especially distant future rewards. In support of this perspective, increased depressive tendencies were related to less accurate response times for no-gains (relative to gains) in the present study, but only following 8-month cues. This result suggests that depressive tendencies are related to deficits in time estimation performance when rewards are available in the distant future, which may result from a decreased motivation to pursue them. Finally, Pearson correlations revealed that only proneness to hypomania was associated with an elevated LPP following 8-month outcomes (see Table S2), although regression analyses showed a significant but opposite correlation with depressive tendencies (see Figure 5). This result indicates that proneness to hypomania was associated with elevated affective and attentional processing for distant-future rewards, while depressive tendencies were associated only with a reduction in motivational salience.

While no prior studies have investigated depressive symptoms and feedback-related ERPs following rewards in the future, one study investigated vulnerability for hypomania and the P3 following future rewards, but found no such relationship (Mason, O’Sullivan, Blackburn et al., 2012). However, the most distant future reward in that study was 1 month in the future while we found effects only for 8 months. Prior studies have found that future rewards recruit discrete neural systems that involve abstract mental representations and extended feedback processing (Ballard & Knutson, 2009; McClure et al., 2004). Importantly, these distinct neural systems may only be recruited following especially distant future rewards that are sufficiently intangible and abstract, whereas near-future rewards may not rely on extended feedback processing to properly evaluate outcomes (Freitas et al., 2001; Gilbert & Wilson, 2007; Rick & Loewenstein, 2008; Trope & Liberman, 2003). When rewards become distant enough in the future, emotional reactions become increasingly difficult to predict and, as a result, the present is used as a reference point to evaluate their significance (Gilbert & Wilson, 2007; Rick & Loewenstein, 2008). Therefore, opposing profiles of P3 amplitude between hypomanic and depressive symptoms may emerge only following feedback for rewards in the distant future while rewards in the near-future remain unaffected.

### 4.4 Limitations and future directions

The present study has some important limitations. First, our results revealed significant associations only between ERPs and symptom dimensions among immediate and 8-month outcomes with no significant effects for the four future rewards in between. It is possible our sample size may be underpowered to detect significant effects for near-future rewards for the RewP and distant rewards closer to 8 months for the P3. Future studies should continue to probe the relationships between symptom dimensions and neural responses to immediate and varying delayed reward (including larger delays than 8 months) in larger samples. Second, depressive tendencies predicted blunted RewPs following 2-week outcomes, but not 2-day. This may be due to our sample population, which consisted of undergraduates who were not at the extremes for hypomanic personality traits or unipolar depressive tendencies. Future studies should investigate whether individuals with clinical levels of hypomania or depression may display more robust relationships with feedback-related ERPs following immediate and future rewards. Third, due to experiment-length considerations to prevent participant fatigue, we did not include a loss or neutral condition (e.g., feedback that is not linked to monetary gains or losses), so the present results are limited to gain-specific modulation of ERPs. Future studies should investigate how future rewards affect neutral- and loss-related ERPs during feedback processing and whether these may be related to dimensions of hypomanic and depressive symptoms.
4.5 Conclusion

Together, our results provide novel insights into the neural mechanisms underlying reward dysfunction across hypomanic and depressive symptom clusters for both immediate and future rewards. Proneness to hypomania and depressive tendencies were related to elevated and blunted RewP difference waves, but only following immediate rewards. This result suggests that the RewP following immediate rewards may be a good candidate biomarker of differential risk that separates both disorders. In addition, both measures displayed similar relationships with the P3 difference wave, but only following distant rewards 8 months in the future, indicating disruptions of extended feedback processing following distant-future rewards. Together, our findings reveal new avenues to investigate differential electrophysiological correlates of risk that may help physiologically bifurcate the fine line between bipolar spectrum and depressive disorders.

ACKNOWLEDGMENTS

This material is based upon work supported by the National Science Foundation Graduate Research Fellowship (DGE-2016231981 to J.E.G.).

REFERENCES


Delorme, A., & Makeig, S. (2004). EEGLAB: An open source toolbox for analysis of single-trial EEG dynamics including independent...


**SUPPORTING INFORMATION**

Additional supporting information may be found online in the Supporting Information section at the end of the article.

**Figure S1:** A. ERP waveforms showing the RewP at Cz (left) and the P3 and LPP at Pz (right) for all six delays (listed on the far left). Gains (black) and no-gains (red) are plotted separately as well as their difference wave (blue). Note that all difference waves are plotted here as no-gain – gain for consistency. B. Displays average scalp topographies for the RewP (250-350 ms), P3 (400-600 ms), and LPP (600-800 ms) at each of the six time delays labeled on the far left of the figure.
Table S1: Displays means and standard errors for each of the ERPs (RewP, P3, and LPP) for all 12 of the 2 (outcome) by 6 (delay) ANOVA (top) and marginal values for delay (middle) and outcome (bottom) separately.

Table S2: Displays results for each regression analysis performed separately on each ERP difference wave at each of the six delays (beta-values, t-values, p-values) for proneness to hypomania (i.e. the hypomanic personality scale, left) and depressive tendencies (i.e. 7-down, right) as simultaneous predictors within each model. The final column of both questionnaires contains r-values for pelson correlations between each ERP difference wave and either proneness to hypomania or depressive tendencies. For both the regression (“p” column) and correlation (“r” column) analyses * denotes significance below .05, + denotes significance below .06.

How to cite this article: Glazer JE, Kelley NJ, Pornpattananangkul N, Nusslock R. Hypomania and depression associated with distinct neural activity for immediate and future rewards. Psychophysiology. 2018;e13301. https://doi.org/10.1111/psyp.13301