Individual Differences in Typical Reappraisal Use Predict Amygdala and Prefrontal Responses

Emily M. Drabant, Kateri McRae, Stephen B. Manuck, Ahmad R. Hariri, and James J. Gross

**Background:** Participants who are instructed to use reappraisal to downregulate negative emotion show decreased amygdala responses and increased prefrontal responses. However, it is not known whether individual differences in the tendency to use reappraisal manifests in similar neural responses when individuals are spontaneously confronted with negative situations. Such spontaneous emotion regulation might play an important role in normal and pathological responses to the emotional challenges of everyday life.

**Methods:** Fifty-six healthy women completed a blood oxygenation-level dependent functional magnetic resonance imaging challenge paradigm involving the perceptual processing of emotionally negative facial expressions. Participants also completed measures of typical emotion regulation use, trait anxiety, and neuroticism.

**Results:** Greater use of reappraisal in everyday life was related to decreased amygdala activity and increased prefrontal and parietal activity during the processing of negative emotional facial expressions. These associations were not attributable to variation in trait anxiety, neuroticism, or the use of another common form of emotion regulation, namely suppression.

**Conclusions:** These findings suggest that, like instructed reappraisal, individual differences in reappraisal use are associated with decreased activation in ventral emotion generative regions and increased activation in prefrontal control regions in response to negative stimuli. Such individual differences in emotion regulation might predict successful coping with emotional challenges as well as the onset of affective disorders.

**Key Words:** Amygdala, cognitive control, emotion, fMRI, regulation, reappraisal

Emotions play a crucial role in adaptation (1), shaping a wide range of cognitive (2) and behavioral (3) responses. However, even as emotions influence how we respond to adaptive challenges, they are themselves subject to regulation. The past decade has seen a dramatic increase in research on emotion regulation (4), and it is now clear that emotion regulation influences emotional experience, peripheral physiology, and neural dynamics (5–7) and impacts a wide range of physical (8) and mental health (9) outcomes. One type of emotion regulation that has been a particular research focus is “reappraisal,” which involves altering the meaning of a potentially emotion-eliciting situation in order to change its emotional impact (6).

**Instructed Reappraisal**

To examine the effects of reappraisal, one common approach has been to instruct participants to use reappraisal to downregulate responses to emotion-eliciting stimuli such as films (10) or slides (11–15). Behaviorally, instructed reappraisal decreases emotion experience (12,13). Neurometrically, instructed reappraisal decreases activation in emotion-generative brain regions such as the amygdala (11–15) and increases activation in a distributed network of brain regions associated with verbal processing, attention, and cognitive control such as the anterior cingulate cortex (ACC), ventral lateral prefrontal cortex (vPFC), dorsal lateral prefrontal cortex (dlPFC), medial prefrontal cortex (mPFC), and orbitofrontal cortex (OFC) (see [7] for review). Functional connectivity analyses suggest that activity in prefrontal regions is inversely related to amygdala activation (10,16).

**Spontaneous Reappraisal**

Although reappraisal is occasionally triggered by explicit instructions in everyday life (such as when a parent coaches a child about how to think about an upcoming exam), it more commonly arises spontaneously, without explicit instructions from another person. Studies of spontaneous (automatic) emotion regulation have typically adopted an individual-difference strategy (17,18), considering individual differences in the use of regulation strategies such as reappraisal, as indexed by the Emotion Regulation Questionnaire (ERQ) (18). This measure has been widely used across a large number of cultures to examine emotion regulation in everyday life (19). Such studies have found that individuals who use reappraisal more frequently have lower levels of negative affect, greater interpersonal functioning, and greater psychological and physical well-being (18). These studies suggest but do not directly demonstrate that spontaneous use of reappraisal engages the same neural systems revealed by studies of instructed regulation.

Three studies have provided initial evidence that spontaneous emotion regulation engages the same neural systems as instructed regulation. Jackson et al. (20) found that individual differences in left-sided frontal electroencephalogram activation negatively correlated with startle magnitude, suggesting that individuals automatically engaged left-sided PFC regions to decrease emotional reactivity. Haas et al. (21) found that individual differences in agreeableness predicted activation in lateral PFC in response to threatening faces, suggesting that agreeable individuals might be more likely than less agreeable individuals to engage in self-regulation. Aber et al. (22) found that depressed individuals who frequently use reappraisal in everyday emotion regulation have decreased amygdala activity and increased prefrontal activity during the processing of negative emotional facial expressions.
life showed decreased amygdala responses during the anticipation of emotion.

These findings are suggestive, but these studies have key limitations. First, these studies did not consider individual differences in emotional reactivity. This is an important limitation, because differences in reactivity (as indexed by neuroticism and trait anxiety) have been shown to predict greater amygdala reactivity (23–26). Second, these studies did not examine both emotion-generative and cognitive-control brain regions. Third, the first two studies did not assess individual differences in the tendency to use different regulation strategies and instead indirectly inferred regulation from other types of individual differences. Only the third study (22) directly assessed individual differences in reappraisal. However, the sample was small (n = 12), and the role of cortical control regions was not reported. Furthermore, the sample contained only depressed individuals, who might not exhibit typical neural correlates of reappraisal (27), and examined anticipation rather than emotion processing per se.

The Present Study

The goal of the present functional magnetic resonance imaging (fMRI) study was to examine the relationship between individual differences in self-reported use of reappraisal and the spontaneous response of the amygdala and control-related PFC regions during the perceptual processing of negative emotional facial expressions. We employed a large healthy sample to avoid confounds with mental illness. We limited analyses to female participants to maximize homogeneity of emotional responses (28,29). We used a well-characterized perceptual face processing task (30,31) to ensure robust activation of the amygdala and interconnected PFC regions. We assessed typical reappraisal use by administering the ERQ, which assesses individual differences in everyday use of reappraisal. We controlled for individual differences in emotion reactivity by assessing neuroticism and trait anxiety and controlled for the use of another common strategy to downregulate emotion by assessing suppression. We hypothesized that participants with higher reported use of reappraisal would be more likely to engage in spontaneous reappraisal and would thus show: 1) decreased amygdala activity, and 2) increased activity in control-related PFC regions. We also hypothesized that these effects would not be due to individual differences in reactivity or the use of suppression.

Methods and Materials

Participants

Fifty-six right-handed female volunteers (mean age 44.0 ± 6.7 years) participated after providing informed consent according to the guidelines of the University of Pittsburgh Institutional Review Board. All participants were recruited from a larger parent study, the AHAB (Adult Health and Behavior) project, which assesses behavioral and biological traits among a community sample of nonpatient, middle-age volunteers. Participants were in good general health and did not exhibit conditions affecting cerebral blood flow and metabolism or current Axis I disorders as diagnosed by the Structured Clinical Interview for DSM-IV.

Experimental Task

The experimental fMRI paradigm (30,31) consisted of four blocks of a perceptual face processing task interleaved with five blocks of a sensorimotor control task. During the face processing blocks, subjects viewed a trio of faces of the same gender, all of which expressed either anger or fear. Subjects selected one of two different faces (presented on the bottom half of the screen) that matched an identical target face (presented on the top half of the screen). Each face processing block consisted of six face trios balanced for gender and emotion (angry or fearful), all derived from a standard set of pictures of facial affect (32). During the sensorimotor control blocks, subjects viewed a trio of simple geometric shapes (circles, vertical, and horizontal ellipses) and selected one of two shapes (bottom) as matching an identical target shape (top). Each image (faces or shapes) was presented for 4 sec with a variable inter-stimulus interval (2–6 sec, mean 4 sec), resulting in a total of 9 blocks lasting 48 sec each. Because we were interested in eliciting a maximal amygdala response across all subjects that we could then interrogate for relationships with reappraisal, we chose to not use neutral faces as control stimuli, because neutral faces can be subjectively experienced as affectively laden or ambiguous and thus engage the amygdala (33,34).

Individual Difference Measures

Individual difference measures were administered before fMRI scanning. The primary measure of interest was the reappraisal scale of the ERQ (18). This scale consists of six items designed to assess individual differences in reappraisal use (e.g., “I control my emotions by changing the way I think about the situation I’m in”). This scale previously has been shown to have good internal consistency and test-retest reliability and to be independent from intelligence (18). To ensure independence of ERQ reappraisal from both intelligence and socioeconomic status in our sample, we obtained two markers of intelligence (highest level of education and IQ score from the Wechsler Adult Intelligence Scale) as well as two markers of socioeconomic status (median household income from the 1999 census and current self-reported family income). Consistent with prior reports, ERQ reappraisal was not correlated with any of these indices (all p values > .38). In addition, we administered control measures, including: 1) the neuroticism scale (N) of the Neuroticism Extroversion Openness Personality Inventory—Revised (NEO-PIR) (35), which assesses an individual’s tendency to experience psychological distress; 2) the trait version of the State Trait Anxiety Inventory (STAI trait version) (36), which assesses relatively stable individual differences in anxiety proneness; and 3) the suppression scale of the ERQ, which assesses use of expressive suppression in everyday life.

Image Acquisition

Each subject was scanned with a Siemens 3T Allegra scanner. Blood oxygenation-level dependent (BOLD) functional images were acquired with a gradient echo planar imaging sequence and covered 34 axial slices (3-mm thick) beginning at the cerebral vertex and encompassing the entire cerebrum and the majority of the cerebellum (repetition time/echo time = 2000/25 msec, FOV = 20 cm, matrix = 64 × 64). Before the collection of fMRI data for each subject a reference echo planar imaging scan was acquired and visually inspected for artifacts (e.g., ghosting) as well as good signal across the entire volume of acquisition, including the medial temporal lobes and ventromedial PFC. Data from all 56 participants were free from artifacts and exhibited good signal in our regions of interest.

Functional MRI Statistical Analysis

Whole brain image analysis was completed with SPM2 (http://www.fil.ion.ucl.ac.uk/spm). Images for each subject were re-
aligned to the first volume in the time series to correct for head motion, spatially normalized into a standard stereotactic space (Montreal Neurological Institute template) with a 12 parameter affine model, and smoothed with a Gaussian filter, set at 6 mm full-width at half-maximum. Voxel-wise signal intensities were ratio normalized to the whole-brain global mean. Condition effects at each voxel were calculated with a t statistic, and regionally specific effects were compared with linear contrasts.

We conducted group level, random-effects analyses to investigate the relationship between individual differences in reappraisal and brain activation during face processing. The ERQ reappraisal scores were entered into a correlation analysis (simple regression) examining the contrast between face processing and sensorimotor control blocks. Unless indicated otherwise, whole brain regression analyses employed a significance threshold of \( p < .001 \) (uncorrected) with a five voxel extent threshold \( (\text{Montreal Neurological Institute template}) \) with a 12 parameter motion, spatially normalized into a standard stereotactic space.

Given our a priori hypotheses regarding the relationship between reappraisal scores and amygdala activation, an anatomical mask of the amygdala (dilated by a factor of 1) \( (38) \) was applied bilaterally. A significance threshold of \( p < .01 \) (uncorrected) with a five voxel spatial extent threshold was applied in this a priori region of interest. We then defined a functional region of interest (ROI) within this anatomical ROI, which encompassed the amygdala voxels above our significance threshold at the group level.

The average contrast values (faces > shapes) for the amygdala voxels in the functional ROI were extracted for each individual with MarsBar \( (39) \). Average values were also extracted from functional clusters in prefrontal and parietal regions shown to be positively correlated with reappraisal. Within each region, contrast values \( > \pm 2.5 \text{ SDs} \) from the mean were excluded from subsequent analyses. To determine the correct voxel-level threshold and cluster extent for amygdala activation, AlphaSim, a Monte Carlo simulation bootstrapping program in the Analysis of Functional NeuroImage (AFN) library (Medical College of Wisconsin), was employed to identify a joint probability consisting of a voxel-wise threshold and a minimum cluster volume threshold to establish a cluster-wise \( p \) value that protects against false-positives. At a voxel-wise threshold of \( p < .01 \), AlphaSim determined that a cluster size of 12 voxels is required to produce an overall \( p < .05 \).

Selective between-subjects functional connectivity analyses were conducted with a simple regression model in SPSS. Average contrast values were extracted from the amygdala functional ROI in the procedure described in the preceding text. These values were correlated with average extracted values from the prefrontal and parietal regions identified in the group level whole-brain analyses as significantly positively correlated with reappraisal.

A final set of regression analyses were conducted to assess whether self-reported reappraisal use predicted brain activation over and above emotional reactivity (neuroticism and anxiety) and another form of regulation (suppression).

**Results**

We hypothesized that self-reported reappraisal use would be negatively correlated with activity in the amygdala and positively correlated with activity in control-related prefrontal regions. We did not expect that these associations would be attributable to individual differences in emotional reactivity or in the use of another common regulation strategy, namely suppression.

**Main Effect of Task**

As expected, we replicated previous findings, showing that this task robustly activates the amygdala. For the contrast faces > shapes, bilateral amygdala activation can be seen in Figure 1.

**BOLD Responses in Emotion-Generative Regions**

As hypothesized, we observed a negative correlation between self-reported reappraisal use and BOLD responses within the right \( (T = 2.40, p < .01, \text{uncorrected}) \) and left \( (T = 2.40, p < .01, \text{uncorrected}) \) anatomical amygdalae. Follow-up analyses confirmed that individuals who reported using reappraisal more frequently in everyday life showed lesser activation in both the right \( (r = −.45, p < .001) \) and left \( (r = −.37, p < .006) \) amygdala during face processing. Activation extent and scatterplot for this negative correlation is reported in Figure 2.

**BOLD Responses in Control-Related Prefrontal Regions**

As hypothesized, we observed a positive correlation between self-reported reappraisal use and BOLD responses in a subset of regions previously identified in studies of instructed reappraisal, including dorsal medial prefrontal cortex (dmPFC) \( (37) \) Brodmann area [BA] 8; \( r = .33, p = .016 \), lateral OFC (BA 10; \( r = .42, p = .002 \)), and dlPFC (BA 9; \( r = .39, p = .004 \)) with a whole-brain approach. Selected activation extent and scatterplot for dmPFC is reported in Figure 3, and all activations are reported in Table 1. In addition to these control-related PFC effects, whole-brain analyses also revealed a positive correlation between reappraisal use and BOLD responses in both inferior \( (BA 40; r = .34, p < .012) \) and superior parietal lobule \( (BA 7; r = .31, p = .025) \).

**Between-Subjects Functional Connectivity Between Amygdala and Prefrontal Regions**

Correlational functional connectivity analyses revealed that activation in the right amygdala was negatively correlated with dmPFC \( (BA 8; r = −.28, p = .041) \) and lateral OFC \( (BA 10; r = −.27, p = .045) \). Activation in the left amygdala was negatively correlated with dlPFC \( (BA 9; r = −.25, p = .049) \).
A contrast value that of reappraisal, and we found that the relationship between it bears noting that this relationship is in the opposite direction to ERQ reappraisal scores did change the model (STAI did not change the model (predicted neural response (BA 8, BA 9, BA 10, BA 40). With one exception, no significant relationships emerged (all $p < .05$). These findings indicate that our primary results were not due to individual differences in emotional reactivity.

Are Findings Specific to Reappraisal?

Address this question, we examined whether the use of suppression was associated with BOLD responses in brain regions associated with reappraisal (left amygdala, right amygdala, BA 7, BA 8, BA 9, BA 10, BA 40), and whether the relationship between reappraisal and these regions was significant for suppression use ($r$ change = .256, $p < .001$).

Discussion

A growing literature demonstrates that instructed reappraisal is associated with decreased activity in the amygdala and increased activity in a prefrontal cognitive control network. In the current study, we examined whether individual differences in everyday reappraisal use would predict activity in the same neural systems identified in studies of instructed regulation. Our findings revealed that self-reported reappraisal use predicted lesser amygdala activation and greater activity in prefrontal and parietal regions, and activity in these regions was inversely correlated across subjects. Importantly, these effects were significant and dependent on individual differences in emotional reactivity as assessed by neuroticism and trait anxiety and for another common form of emotion regulation (suppression).

Amygdala Activation and Individual Differences in Reappraisal Use

Studies of instructed reappraisal have shown reliable down-regulation of amygdala activation in response to negative stimuli (11–15). In the current study, we found that greater everyday reappraisal use also predicted decreased amygdala activation. Given that amygdala responses are one important central index of emotional responding, as demonstrated by the amygdala’s role in animal and human fear conditioning and coordinating downstream fear responses (see [40] for review), these results suggest an important role for individual differences in reappraisal in underlying emotional behavior.

Control-Related Activations in Instructed Versus Spontaneous Emotion Regulation

We found that, consistent with studies of instructed reappraisal, greater everyday reappraisal use predicted increased activation in regions associated with deploying a cognitive strategy (dLPFC) (41), reappraising emotional states of self and others (dmPFC) (42), evaluating emotional value on the basis of context (lateral OFC) (43), and directing spatial attention (parietal cortex) (44).
Given that the regions of neural activation observed here closely resemble those found in instructed reappraisal (see [7] for a review) and that these activations are specifically correlated with self-reported reappraisal use, one interpretation is that these activations result from spontaneous use of reappraisal during the experimental task. It bears noting, however, that whereas the current study reveals activation in many of the same regions previously associated with instructed reappraisal, the spatial extents of these activations are dramatically smaller than the typical activations seen in studies of instructed reappraisal, particularly in dlPFC. Why might this be?

One possibility is that, although initial use of reappraisal requires some effort, with frequent use it might become less effortful and thus require fewer cortical resources to achieve the same reduction in amygdala activation. This idea is consistent with research showing that frequent use of a working memory strategy results in decreased activation in dlPFC during subsequent memory trials. This decrease in neural activation in dlPFC is accompanied by an increase in working memory performance (45), suggesting that frequent use of the strategy results in greater neural efficiency. In the current study, it is possible that the reduced activation seen in the cognitive control network indicates that spontaneous reappraisal is more automatic and neurally efficient.

**Methodological Implications**

The current findings have important implications for the interpretation of results from simple tasks thought to index “pure” emotional reactivity. Our task robustly engages the amygdala and is therefore thought to be an excellent probe of the neural basis of emotional reactivity. Indeed, the task employed here has become a staple in affective neuroimaging in recent years as an assay of emotional reactivity in a wide array of populations. However, the current findings suggest that even “pure” reactivity tasks such as the one used here might actually be intertwined with emotion regulation, which is dissociable from emotional reactivity to the stimuli. This should be taken into account when interpreting findings from these types of tasks.

For example, several researchers have used this task to examine the differences in a putative emotion regulation circuit between amygdala and PFC on the basis of genetic background (e.g., [46]), personality (47), and clinical diagnosis (48). Findings from these studies have been interpreted as between-group differences in emotion regulation ability, given the association between amygdala and PFC regions in studies of instructed regulation. The present results buttress such accounts and provide the first empirical evidence for the claim that individual differences in emotion regulation might mediate the decreases in amygdala activity and increases in PFC activity observed during this task.

**Clinical Implications**

There is an increasing appreciation that emotion dysregulation underlies a variety of psychiatric illnesses, including mood and anxiety disorders (49). Empirical studies that aim to characterize mood and anxiety disorders have begun to focus on how individual differences function as risk factors for such disorders. Decreased use of reappraisal in particular, as indexed by the ERQ, is associated with increased depressive symptoms (18). In addition, neuroimaging research has shown that increased amygdala activation in response to negative stimuli is associated with depression (50). Our results represent a crucial link between

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Activations for the contrast faces > shapes for whole brain analysis. The display threshold was \( p < .001 \) with an extent threshold of five voxels. Coordinates are listed in Talairach space. The \( r \) and \( p \) values represent correlation between extracted values and reappraisal. BA, Brodmann area; PFC, prefrontal cortex; OFC, orbitofrontal cortex.

Figure 3. Positive correlation between reappraisal and prefrontal activations. (A) From the simple regression (correlation) between Emotion Regulation Questionnaire (ERQ) reappraisal score and the contrast faces > shapes. The display threshold was \( p < .001 \) with an extent of five voxels. (A) Activation in the medial frontal gyrus (Brodmann area [BA] 8; \( x = −2 \), \( y = −22 \), \( z = −47 \); 20 voxels) identified within the whole brain analysis (other activations reported in Table 1). (B) Scatter plot of ERQ reappraisal score and average contrast values extracted from voxels identified in panel A. BOLD, blood oxygenation-level dependent.
increased amygdala activity and the decreased use of reappraisal during basic emotion processing. The present study, therefore, offers a potential neural mechanism by which individual differences in emotion regulation result in greater neural responses to emotional stimuli, which might characterize both risk for (30) and pathophysiology of (51) depression and related affective disorders.

One of the most widely used treatment interventions for mood and anxiety disorders is cognitive behavioral therapy (CBT) (52,53). One feature of CBT is that relatively brief interventions (e.g., 10 weeks) can have long-lasting effects (54). A key element to understanding the effects of CBT involves characterizing the transition from therapist-guided reinterpretations of negative cognitions to automatic reinterpretations of negative cognitions. Given the relationship between instructed reappraisal and the therapist-guided reinterpretations that are part of CBT, the present finding highlights a potential mechanism by which the reframing taught in CBT becomes more self-generated and automatic. Future work should track the neural changes associated with reappraisal use in relation to progress made over the course of CBT.

Limitations and Future Directions

The present study has several important limitations. First, only healthy female participants were studied, and it is therefore unknown whether our results are generalizable to clinical samples or to men. Because gender differences are often observed in studies of emotional reactivity (28,29) and sometimes observed in studies of emotion regulation (55), it will be important to examine the neural correlates of emotion regulation use in men as well as women. Given the relationship between emotion dysregulation and mood and anxiety disorders, studying emotion regulation in various clinical populations is necessary to improve our understanding of emotion regulation deficits and offer insight into treatment implications.

Second, we examined only one task context—namely, simple perceptual face processing. This task robustly activates corticolimbic circuitry, providing a strong target for downregulation without specific instructions to regulate, and thus was an ideal choice for this investigation of the relationship between everyday reappraisal use and spontaneous neural responses to negative emotions. However, it is important to go beyond basic processing to better characterize the interplay between emotion generative and emotion regulatory brain systems across different contexts. Subsequent studies could examine emotion regulation in relation to positive as well as negative faces, subliminal face presentation, or a direct comparison of instructed versus uninstructed contexts.

Third, reappraisal was assessed with one self-report measure—namely the ERQ. Previous research has shown that the ERQ predicts important real-world outcomes such as well-being and depressive symptomatology (18), and there is ample precedent for using single self-report measures to examine relations between individual differences in emotional reactivity and brain activity (23–26). In future work, however, it will be useful to also employ non-self-report measures of emotion regulation when examining brain responses to look for convergence across diverse measures.

Fourth, we did not directly assess which emotion regulation strategies participants were using, if any, during the task. We felt it was important to avoid calling attention to any potential regulation processes by having participants indicate which strategy they were using. However, future studies might ask individuals to retrospectively report what strategies (if any) they used to impact their emotional responses to these faces. This might further clarify the differences in task processing associated with self-reported reappraisal use. Other types of processing could result in similar activation patterns, such as distraction (56), decreasing attention to the affectively salient features of the stimuli (15), or simply labeling the affective expressions depicted (57–60). However, it is noteworthy that in the present study these effects were not explained by individual differences in suppression.

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