

Systems/Circuits

for choice comparison based on reward value [e.g., the frontoparietal network and the dorsal anterior cingulate cortex (dACC)–anterior insular cortex (AIC) network; Monterosso et al., 2007; Hoffman et al., 2008]. In addition, task-related activity in the frontoparietal network and the dACC–AIC network is engaged in deliberation and emotional processes, respectively, during decision-making (Sanfey et al., 2006), and the rsFC in the frontoparietal network and dACC–AIC network is correlated with executive task performance and anxiety, respectively (Seeley et al., 2007). The present study investigated whether the rsFC between brain regions recruited during the DDT is associated with preference for immediate reward. Moreover, given the different function roles of the DDT-related neural networks, we examined whether the rsFC within these neural networks is associated with the impulsivity of economic decision-making in a similar vein.

Materials and Methods

Experiment I

Participants

Twenty-three Chinese adults (16 males; mean \pm SD age, 22.8 \pm 1.6 years, ranging from 20 to 25 years; mean \pm SD years of education, 16.5 \pm 1.6, ranging from 13 to 19 years) participated in experiment I. All participants reported no history of neurological or psychiatric disorders. Written informed consent was obtained before the study. This study was approved by the Human Research Ethics Committee of the University of Science and Technology of China.

Experimental procedure

DDT

The DDT used a jittered single-trial design, similar to Ballard and Knutson (2009) (Fig. 1a). To ensure that each future reward option would be presented only once during the entire task and that each block would cover a similar size of a two-dimensional space for monetary magnitude and time delay of future reward options, future reward options were combined with two sets of monetary magnitudes (M1: 50, 55, 70, 90, 110, 140, 175 Chinese yuan; M2: 50, 60, 68, 75, 100, 125, 150 Chinese yuan) and two sets of time delays (T1: 0, 7, 30, 60, 90, 180 d; T2: 0, 10, 28, 50, 100, 160 d) as M1T1, M1T2, T1M2, and T2M2, yielding four

blocks with 42 trials each. The first and second combinations (“money-to-time” blocks) and the third and fourth combinations (“time-to-money” blocks) were counterbalanced in terms of the order in which future reward options were presented. The order of the four blocks was counterbalanced across participants.

Each participant performed a short version of the DDT to get familiar with the task (\sim 5 min and without any payment) before scanning. During scanning, to ensure incentive compatibility and to verify that discounting measures were reliable, participants were informed that they would receive actual payment (between 50 and 175 Chinese yuan) based on their choice on one randomly drawn trial of the task. Participants came to our laboratory and got the payment within 24 h after the scanning if the outcome of the selected trial was a monetary gain in 0 d or in the specified delay for the delayed reward outcome of the selected trial. Participants were paid in cash on the same day if the outcome of the selected trial was immediate monetary gain or after a specified period of delay if the selected trial was a delayed reward. Participants were paid with cash. There was no extra payment to the participants.

MRI data acquisition

Imaging data were obtained with a 3 T Siemens Magnetom Trio scanner in the Anhui Provincial Hospital (Hefei, Anhui, China). A circularly polarized head coil was used, with foam padding to restrict head motion. Functional images were acquired with a T2*-weighted echo-planar imaging sequence (echo time, 30 ms; repetition time, 2 s; field of view, 24 cm; matrix, 64 \times 64) with 33 axial slices (slice gap, 0 mm; one voxel: 3.75 \times 3.75 \times 3.7 mm), covering the whole brain. Before entering the MR scanner, all the participants were told to keep their heads steady during all scans. Resting-state fMRI data were first acquired with one functional 8 min scan (240 epochs) when participants were asked to keep their eyes closed. This was followed by four functional scans corresponding to the four blocks of the DDT. Each lasted for 8 min, 40 s, during which participants performed the DDT. Between every two scans, there was an interval of \sim 1 min. Corresponding high-resolution T1-weighted spin-echo (for anatomical overlay) images and three-dimensional gradient-echo (for stereotaxic transformation) images were also collected.

Association analyses

Behavioral data analyses

Each participant's discounting rate was assessed based on the choices made in the DDT (Ballard and Knutson, 2009). For each delay, the choices were defined as 0 for choosing the immediate option and 1 for choosing the future option and then were fit with a logistic function to

called discounting curve (Fig. 1*b*), in which D refers to the delay in days, and k is an individual's discounting rate, with larger k indicating higher impulsivity in decision-making. Trials with missed choices were excluded from this processing and from subsequent processing. Trials with response times (RTs) < 100 ms were also excluded, because these indicated responses before the choice screen being presented. The other trials were defined as hit trials and were included in the analyses.

Easy trials and hard trials were defined to parse the difficulty level of decision-making, similar to Hoffman et al. (2008). For each trial, distance to discounting curve is defined as $DV' - D\hat{V}$, where

$$DV' = \frac{\text{magnitude of immediate reward (50 yuan)}}{\text{magnitude of future reward}},$$

and $D\hat{V}$ is the value on an individual's discounting curve in the corresponding delay. RTs were fit against the distance to each individual's discounting curve (dist) by a normal distribution function:

$$RT = C + A \cdot \exp\left(-\frac{1}{2} \left(\frac{\text{dist}}{w}\right)^2\right).$$

Trials located within the area $\pm w$ were defined as hard trials, and other trials outside this area were defined as easy ones (Fig. 1*c*). For trials with future reward options involving either 50 yuan or 0 d, which are the same as the immediate reward option in either monetary magnitude or time delay, participants could make their choices by simply comparing the digits of the other information of the options (i.e., comparing time delay for the 50 yuan future reward options, and comparing monetary magnitude for the 0 d future reward options), without integrating the information of money magnitude and time delay that is considered as an essential process in the DDT (Green and Myerson, 2004). Thus, these trials, labeled as control trials, were considered as the baseline of the decision-making demand (Boettiger et al., 2007; Hoffman et al., 2008) and were excluded from hard and easy trials.

fMRI data analysis

Preprocessing. The imaging data were processed with AFNI (Analysis of Functional Neuroimages) (Cox, 1996) and MATLAB (version 7.6.0.324; MathWorks). Each participant's raw data were corrected for temporal shifts between slices, corrected for motion, spatially smoothed with a Gaussian kernel (full width at half maximum = 4 mm), and temporally normalized (for each voxel, the signal of each epoch was divided by the temporally averaged signal). We scanned a total of 27 participants in experiment 1, and four were excluded from imaging data analysis as a result of head motion being larger than 2.0 mm. To reduce the effect of head motion and obtain low-frequency fluctuation from resting-state fMRI data, we regressed the motion data out of the time series and then preformed bandpass temporal filtering (0.01–0.08 Hz) on the residual signals (Birn et al., 2006; Auer, 2008). Then, to further reduce nuisance signals, we regressed out the average signals in the white matter and the CSF (Ma et al., 2010, 2011). The mask of white matter for each participant was determined from the high-resolution structural image using FAST segmentation program of Functional MRI of the Brain software library (www.fmrib.ox.ac.uk). The resultant white matter segmentations were merged by 80% tissue type probability. The CSF mask for each participant was manually drawn according to the anatomical boundaries of the cortical structures of a standardized Talairach atlas brain (Talairach et al., 1992), transformed onto the image space of the individual, and then modified according to the cortical structures of the individual brain by referencing to the anatomical boundaries in the high-resolution three-dimensional structural image. These nuisance signals were used to account for fluctuations unlikely to be relevant to neuronal activity (Fox et al., 2005; Birn et al., 2006; Di Martino et al., 2008). The resultant resting-state fMRI data were then subjected to functional connectivity analysis.

Localizing regions of interest. We first defined the regions of interest (ROIs) using the DDT-related fMRI data for additional rsFC analysis. Because decision-making engages both valuation and choice processes (Kable and Glimcher, 2009; Peters and Büchel, 2011), we used a multiple regression analysis to localize the ROIs related to valuation and choice

processes in the DDT, respectively. This analysis included the following psychological regressors: a "money" regressor (defined as 1 to 7 according to the seven levels of monetary magnitudes of future rewards for each scan when the money information was presented and 0 for the other epochs), a "time" regressor (defined as 1 to 6 according to the six levels of time delays of future rewards for each scan when the time information was presented and 0 for the other epochs), two "choice" regressors for hard and easy trials, respectively (defined as 1 for hard/easy trials when participants were required to choose between the options and 0 for the other epochs), and "period" regressors (defined as 1 for period effects for the epochs when the immediate reward was presented or when the first information of future reward was presented or when choice was required, respectively, and 0 for the other epochs). These regressors were convolved by the gamma function to approximate the hemodynamic response of the brain to a stimulus. The regression analysis also included six regressors for head motion and two regressors for linear trends and constant for each scan.

In localization of the ROIs for valuating monetary magnitude in the DDT, each participant's preprocessed time series of the money-to-time scans were concatenated and the regressors were applied in the multiple regression analysis. The resultant β -value map of the money regressor for each participant was transformed to the Talairach space (resampled voxel size: $2 \times 2 \times 2$ mm) according to the spatial transformation between the anatomical data and the Talairach space and was then entered into group-level one-sample t test. The group-level t test map was masked by the gray matter of the brain (Ballard and Knutson, 2009) and then merged at the threshold of cluster size 20 voxels (160 mm^3) and uncorrected $p < 0.005$ for single voxels. Clusters in which the activation showed a positive correlation with monetary magnitude were defined as the money-related ROIs (Ballard and Knutson, 2009). The regression analysis showed activations in brain regions of the mesolimbic cortices [e.g., the striatum, the medial prefrontal cortex (mPFC), and PCC], which were found to be related to processing reward in previous studies (McClure et al., 2004; Kable and Glimcher, 2007; Ballard and Knutson, 2009; Pine et al., 2009), as well as one cluster in the occipital cortex that was, however, excluded in the analysis of resting-state data because the occipital activation might simply arise from differential visual processing of monetary digits. The procedure of localizing the ROIs for valuating time delay was similar to that for the money-related ROIs but used the imaging data in the time-to-money scans as the method implemented by Fitzgerald et al. (2010). Because shorter time delay represents placing a greater value on future reward (Ballard and Knutson, 2009), the time-related ROIs were defined as the clusters in which the activation for the time regressor showed negative correlation with time delay.

The procedure of localizing the ROIs for the choice process was similar to that for the money-related ROIs (both the money-to-time and time-to-money scans were used). In the group-level one-sample t test, clusters that showed a positive activation in the contrast between choice regressors of hard and easy trials were merged at the significance level of $p < 0.05$ [familywise error corrected, i.e., uncorrected $p < 0.005$ for single voxels and a minimum cluster of 53 voxels (424 mm^3)]. Most of the activated clusters were well separated from each other in the functional statistic map, and only a few activation areas in dorsal part of the PFC were further segmented into subregions because they may be involved in different functions (Aron et al., 2004; Goldstein et al., 2007; Heekeren et al., 2008; Rangel et al., 2008; Rowe et al., 2010) by referencing them to the anatomical boundaries in the Talairach atlas. The resultant ROIs were further separated into two subsets, the frontoparietal network and the dACC-AIC network, because brain regions in the two networks have different functions in decision-making (Sanfey et al., 2006) and are dissociated in their rsFC (Seeley et al., 2007).

In total, we localized the brain regions that were involved in valuating monetary magnitude and time delay, and we labeled them as "the money network" and "the time network," because according to our localization procedure, which was similar to that by Ballard and Knutson (2009), the two networks were dissociated in their functions during the DDT. In addition, the brain regions associated with the choice process were divided into two subsets mainly according to previous evidence of functional differences in decision-making (Sanfey et al., 2006). Thus, the two

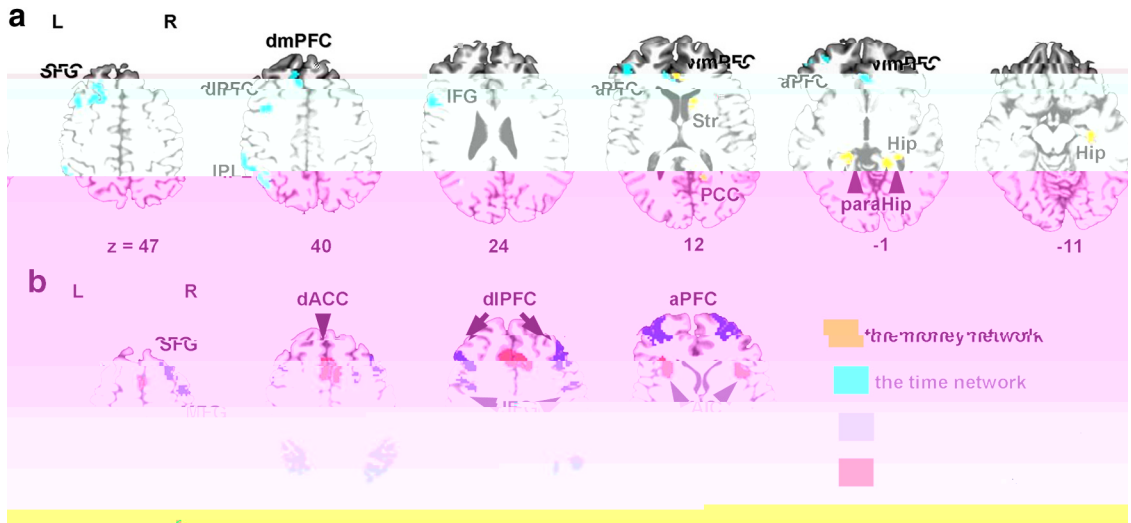


Figure 2. NNe3v@vus rz vxz uvwvus r•3v@vurtz rz 4a2L rz vxz v@vu yv r@r z tv 4eyv v v •t z vu w rz vxz z yzy yvrtzz r zzv@t v@vu zy vr rx z uv ww vvr uEyv z v v •t z vu w rz vxz z yzy yvrtzz r vxr zv@t v@vu zy z vuv@ ww vvr u4b2 L rz vxz v@vu zy yvtv zv tv 4eyv v vuvuvus t r zxyru v vr x@z yvNNe r u vvw yv uz avuz yvw r zr@v •r uyvuKMM KTM v • w yvz uz w v w tz r@ z vtz z 3 r•z x4d 2d x ESz 2yz tr ErrSz 2 rryz tr EX PR2 zuu@w r@ 4

subsets were labeled more conventionally, as “the frontoparietal network” and “the dACC–AIC network.” The four networks were well separated from each other in their containing brain regions (Fig. 2, Table 1). Although the brain regions in the time network were close to those in the frontoparietal network, the majority of the regions in the two networks were dissociated. The subsequent analysis was based on the rsFC within and between these four DDT-related networks.

Defining functional connectivity intensities for the ROIs. The rsFC of the ROIs was analyzed using the algorithm from graph theory (He et al., 2008; Liu et al., 2008; Bullmore and Sporns, 2009) that allows the definition and calculation of the properties of a network involving multiple brain regions along a series of predefined thresholds (Achard and Bullmore, 2007; Stam et al., 2007). This has the advantage of providing a global perspective on the functional organization of the brain and enables the discovery of sensitive thresholds for measuring brain networks. The present study used the parameter known as K_{cost} in graph theory as a measurement for the functional connectivity intensity (FCI) of the DDT-related brain regions. This parameter measures the sparseness of the connections in a network (Latora and Marchiori, 2003).

For each participant, the ROIs defined in Talairach space were transformed to one’s original image space and then modified according to the cortical structures of the individual brain by referencing to the anatomical boundaries in the high-resolution three-dimensional structural image. The preprocessed resting-state fMRI data were averaged within each ROI. Correlation coefficients (r values) were calculated between each pair of the averaged time series and then transformed to Fisher z values, yielding an $N \times N$ functional connectivity matrix for each participant. We merged each z value matrix with a T value as threshold and obtained an undirected binary graph G , consisting of nodes (ROIs) and edges/functional connections (functional connectivity between pairs of ROIs), with the edge between the i th and the j th node in G defined by the formula

$$e_{ij} = \begin{cases} 1 & \text{if } |z(i,j)| \geq T, i \neq j \\ 0 & \text{otherwise} \end{cases},$$

which means that, if the absolute z value of functional connectivity between the i th and the j th ROI is larger than T , the edge between the i th and the j th node in graph G exists and vice versa. Because both positive and negative connections contribute to the functional organization of a network (Fox et al., 2005), most previous studies defined the connections between two brain regions using the absolute value of the correlation coefficient. Thus, a larger number of connections surviving in a network could represent more information exchange within that brain network under a certain threshold (He et

Table 1. Regions of interest defined by the brain activations during the delay discounting task

cvxz	L	u	r	r	v	r	g	@	v.	9/	x ^d	y	z
gr@r z													
X v													
c aPM	8:						7B:					:4	984
c d x							:66					764	764
c aMM	89597						;6:					764	764
c Sz							8: B					884	884
							7B:					984	984
W r rSz	96596						;8B					7A4	7A4
c r rSz							8A8					7@4	7@4
ez v													
Wu aPM	B						B6B					9A4	9A4
W aPM	8:						8: B					9A4	9A4
c aPM	8:						8: 6					9A4	9A4
WIPR	@B						776:					864	864
Wu@PM	C5: @						B98					9: 4	9: 4
WIPR	::						7698					8A4	8A4
W aPM	76						7: A@					9: 4	9: 4
W aW	A59C						89B:					8: 4	8: 4
My zv tv b													
uKMM	8: 598						AB7@					74	74
WKTIM	79						87C8					964	964
c KTM	79						8@ B					994	994
c dPR	B						8666					884@	884@
c X PR ^c	B						786B					9@C	9@C
Wu@PM	C5: @						7B6B					8: 4@	8: 4@
c u@PM	C5: @						999@					8: 4@	8: 4@
WIPR ^c	@:						78: 6					8: 4	8: 4
c TPR	@:						C86					8: 4	8: 4
W aPM	76						98BB					984	984
c raPM	76						: A78					964	964
W aW	A59C						;6@					9: 4	9: 4
c TaW	A59C						ABC@					9B4	9B4

W2WwEc2 xy ESz 2yz tr ErrSz 2 rryz tr EX PR2 zuu@w r@ 4
M uz r v z er@rty rtv .x2@w xy Ey2 vz r v z Ez2 w z vz /4
b'cvxz wyv r@r z tv D z zvt v@z zy v rx z uE vxr zvt v@z zy z vuv@E
yvy @r t vt vup<646;2t@ v zv7@ 94cvxz wyvty zv tv D@xv rt z r z z yv u
x@yr z vr x@Eyvy @r t vt vup<646;2t@ v zv:8: 92t vt vup<64;4
d'vx v v vxz 4

al., 2008; Liu et al., 2008; Bullmore and Sporns, 2009). The present research also used the absolute value of correlation coefficients to define the global functional organization of a network.

For an individual's functional connectivity matrix, we varied the threshold value T from 0 to the maximum of the absolute z value with a step of 0.05, which produced a series of graphs from those with all possible edges between the nodes to those with no edges between any nodes. For a graph G under a certain threshold T , the FCI within the network was defined as the cost of the graph, K_{cost} , which is the proportion of the total number of edges in the maximum possible number of edges:

$$\text{FCI}_{\text{within}_G} \equiv K_{\text{cost}} = \frac{1}{N(N-1)} \sum_{i \in G, j \in G} e_{ij},$$

and could represent the sparseness of functional connections in a graph. Moreover, we further defined the FCI between two networks (e.g., G and H) as the proportion of survived edges:

$$\text{FCI}_{\text{between}_G \& H} \equiv K_{\text{cost_inter}}$$

$$= \frac{1}{N_G \times N_H} \sum_{i \in G, j \in H} e_{ij}.$$

Because we defined four brain networks related to the DDT, 10 FCIs were calculated, including the following: (1) three FCIs for networks of the valuation process: the FCI within the money network, the FCI within the time network, and the FCI between the money and time networks; (2) three FCIs for networks of the choice process: the FCI within the frontoparietal network, the FCI within the dACC–AIC network, and the FCI between the frontoparietal and dACC–AIC networks; and (3) four FCIs for networks of the interaction between valuation and choice processes: the FCI between the money and frontoparietal networks, the FCI between the money and dACC–AIC networks, the FCI between the time and frontoparietal networks, and the FCI between the time and dACC–AIC networks.

Correlation between FCIs and discounting rates. To investigate the association between the rsFC of the ROIs and individuals' discounting rates, multiple linear regression analyses were conducted along all the T steps with FCIs as independent variables (regressors) and discounting rates [log transformed (Hariri et al., 2006)] as dependent variable. (For some T steps, some FCIs might be zeros or ones across all the participants, so these FCIs did not contribute to the variation of discounting rates, which were excluded from the multiple linear regression model, leaving the other FCIs in the model.)

We determined a proper set of FCIs that could be used as regressors in the regression model in the following way. We subjected all possible combinations within the 10 FCIs (1023 combinations in total) to the multiple linear regression analysis across the thresholds. We used adjusted R^2 ($\text{adj-}R^2$) to measure the contribution of an FCI combination to the regression model. This is the ratio of the variance explained by the regressors to the total amount of variance in the dependent variable adjusted for the number of explanatory terms in the model, and this statistic has been shown to be an unbiased estimator of the contribution of a set of explanatory variables x to the explanation of y (Ohtani, 2000).

For each combination of FCIs, we calculated the median of the $\text{adj-}R^2$ values of the regression models across all the thresholds. If all FCIs reached zeros or ones at a specific threshold, the multiple linear regression model cannot be constructed and the $\text{adj-}R^2$ values would be zero. The $\text{adj-}R^2$ values on these thresholds were excluded from the median calculation. Moreover, for some thresholds, the $\text{adj-}R^2$ values might be negative. This suggests that the explanatory variables explain less variation than random normal variables would. In this case, the $\text{adj-}R^2$ values would be modified to zero (Legendre, 2008) and passed into the median calculation.

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The combination with the largest $\text{adj-}R^2$ median value consisted of five FCIs, including the FCI within the money network, the FCI between the money and time networks, the FCI within the frontoparietal network, the FCI within the dACC–AIC network, and the FCI between the money and dACC–AIC networks. Meanwhile, these FCIs were contained in bo262.(value)-5(the)-297

with discounting rates as the dependent variable, forming the correlation model as the norm model for prediction. Next, FCIs of the 23rd participant (the “to-be-predicted” participant) were calculated using the brain regions defined by data from the other 22 participants and then averaged across all the thresholds weighted by the same z values, which were defined by data from the other 22 participants. Finally, the weighted averaged FCIs of the 23rd participant were substituted into the correlation model to calculate the predicted discounting rate for the participant.

In prediction analysis, both the norm model and FCIs of the to-be-predicted participant were independent of the participant’s task-state data, making the predicted discounting rate independent of his/her task-state data. The predicted discounting rates for all 23 participants were entered into a correlation analysis with their actual discounting rates (both were log transformed).

Correlation between discounting rate and head motion during resting-state fMRI scan

Because head motion during scanning might be a trait and thus could be confounded with individual differences (Van Dijk et al., 2012), we conducted a correlation analysis of the participants’ head motion during the resting-state scanning and their discounting rates to examine whether head motion confounded the results of the association and prediction analyses. With the motion parameters generated in the motion correction phase of preprocessing, four separate metrics of head motion were calculated, including mean motion, maximum motion, number of movements, and rotation (Van Dijk et al., 2012). All four metrics were entered into correlation analysis with participants’ discounting rates (log transformed).

Experiment II

The prediction analysis in experiment I suggested that the FCI-discounting rate association model could be used to predict an individual’s discounting rate. However, there is evidence that the generalizability of an internally validated prediction may be poor for a new sample (Bleeker et al., 2003), and external validation with independent samples is considered to be more relevant than internal validation when the prediction model is applied in another practical setting (Steyerberg et al., 2001; Toll et al., 2008). Thus, we conducted a second experiment to provide external validation with an independent sample of rsFC as a predictor of discounting rate.

Participants

An independent group of 38 Chinese adults (32 males; mean \pm SD age, 23.5 ± 1.8 years, ranging from 20 to 28 years; mean \pm SD years of education, 16.7 ± 1.6 years, ranging from 13 to 19 years) participated in experiment II. All participants reported no history of neurological or psychiatric disorders. Written informed consent was obtained before the study. This study was approved by the Human Research Ethics Committee of the University of Science and Technology of China.

Experiment procedure

Resting-state fMRI data were from another study of our group and were obtained in the same manner as in experiment I. Corresponding high-resolution T1-weighted spin-echo (for anatomical overlay) images and three-dimensional gradient-echo (for stereotaxic transformation) images were also acquired. After fMRI data acquisition and outside the scanner, each participant completed the same version of the DDT used in experiment I. Some participants completed other cognitive tasks in the scanner (data will appear elsewhere).

Data analyses

The DDT-related brain regions and the norm model obtained in experiment I were used to define ROIs in experiment II. The five FCIs selected in association analysis in experiment I were used to predict discounting rates following the same procedure as in experiment I. These participants’ predicted discounting rates were entered into a correlation analysis with their actual discounting rates (both were log transformed). Correlations between head motion and participants’ discounting rates (log transformed) were also calculated.

Results

Experiment I

In experiment I, participants’ discounting rates in the DDT ranged from 0.00281 to 0.05529 (mean \pm SD, 0.01872 ± 0.01228), similar to previous results (Ballard and Knutson, 2009). The proportion of hit trials was high ($98.8 \pm 2.2\%$). Within the hit trials, except control trials (for details, see above, Behavioral data analyses), there were $65.7 \pm 4.6\%$ easy trials and $34.3 \pm 4.6\%$ hard trials. Participants’ choices for immediate and future reward did not significantly differ between easy and hard trials ($F_{(1,22)} < 0.001$, $p = 0.985$, ANOVA; Fig. 1c), indicating that the difficulty of the trials was orthogonal to choice of reward.

The analysis of the DDT-related fMRI data yielded the ROIs of four networks. The money network, in which activations were positively correlated with monetary magnitude and thus were engaged in the valuation process, consisted of the right ventral medial PFC (vmPFC), right striatum, right PCC, right hippocampus (two clusters), and bilateral parahippocampus. The time network, in which activations were negatively correlated with time delay, consisted of the left anterior PFC (aPFC), left superior frontal gyrus (SFG), left dorsolateral PFC (dlPFC), left inferior frontal gyrus (IFG), bilateral vmPFC, left dorsal mPFC (dmPFC), and the left inferior parietal lobe (IPL) (Fig. 2a, Table 1). These two networks are similar to those previously observed to be engaged in a similar paradigm (Ballard and Knutson, 2009).

Two networks related to the choice process were obtained in which activations were identified by the contrast of hard versus easy trials during the DDT. The frontoparietal network consisted of the bilateral aPFC, dlPFC, IFG, IPL, the right SFG, and the right middle frontal gyrus. The dACC–AIC network consisted of dACC and the bilateral AIC (Fig. 2b, Table 1). The activation of these two networks is consistent with previous observations using similar paradigms (Monterosso et al., 2007; Hoffman et al., 2008).

Multiple linear regression analysis using the five selected FCIs and individuals’ discounting rates in experiment I showed that individuals’ discounting rates were positively correlated with the FCI within the money network, the FCI within the dACC–AIC network, and the FCI between the money and dACC–AIC networks. However, discounting rates were negatively correlated with the FCI between the money and time networks and the FCI within the frontoparietal network (Fig. 4a). Multiple linear regression analysis using all the possible combinations of the 10 FCIs showed a pattern similar to that obtained from the above five FCIs (Fig. 4b), suggesting robust correlations between impulsivity and the five FCIs. These results revealed the association between intrinsic functional organization and individuals’ impulsivity in decision-making.

Prediction analysis using the leave-one-out procedure in experiment I revealed that the predicted discounting rates based on the five FCIs selected in association analysis were significantly correlated with actual discounting rates ($r = 0.426$, $n = 23$, $p = 0.043$; Fig. 5b), indicating that the norm model based on resting-state activity can predict the discounting rate for a given participant.

Experiment II

In experiment II, the delay discounting rates of the independent sample ranged from 0.00067 to 0.04302 (mean \pm SD, 0.01439 ± 0.01100), similar to that in experiment I. The proportion of hit trials was high ($99.8 \pm 0.3\%$). In external validation, the estimated multiple linear regression model used to predict discounting rates was

predicted discounting rate =

$$\begin{aligned}
& 2.67 \times \overline{\text{FCI}}(\text{within_the_money_network}) \\
& - 12.15 \times \overline{\text{FCI}}(\text{between_the_money \& time_networks}) \\
& - 2.05 \times \overline{\text{FCI}}(\text{within_the_frontoparietal_network}) \\
& + 1.64 \times \overline{\text{FCI}}(\text{within_the_dACC-AIC_network}) \\
& + 2.61 \times \overline{\text{FCI}}(\text{between_the_money \& dACC-AIC_networks}) - 3.21,
\end{aligned}$$

where FCI represented the weighted average for the corresponding FCI, and the β values in the model were consistent with the association results in experiment I. Based on this model, predicted discounting rates were derived for the participants. A correlation analysis showed that the predicted discounting rates were significantly correlated with participants' actual discounting rates ($r = 0.453$, $n = 38$, $p = 0.004$; Fig. 5c). These results constitute external validation and suggest that the norm model of the FCI-discounting rate association can predict the discounting rate for a given participant based on his resting-state fMRI data in an independent sample.

Correlation between head motion and discounting rate

The correlations between the four metrics of head motion and discounting rates were not significant in either experiment I (all p values > 0.270) or experiment II (all p values > 0.643), indicating that the relationship between resting-state brain function and impulsivity cannot be attributed to individuals' head motion during the resting state.

Discussion

The previous fMRI studies found associations between impulsive decision-making behavior and DDT-related brain activation (McClure et al., 2004; Boettiger et al., 2007; Kable and Glimcher, 2007) but leave open the question of whether resting-state activity can predict impulsive decision-making. The current work provides the first evidence linking individuals' impulsivity and rsFC in brain regions involved in the DDT. Moreover, we found distinct patterns of association between resting-state FCIs in different neural networks and individuals' impulsivity in decision-making.

The resting-state FCI within the money network was positively correlated with discounting rates. As noted previously, the money network consists of the vmPFC, striatum, PCC, hippocampus, and parahippocampus and is engaged in valuation of both immediate and future rewards (McClure et al., 2004).

parietal cortex, are consistently engaged in deliberation processes, such as problem solving, planning, and cognitive control (Davidson and Irwin, 1999; Miller and Cohen, 2001; Kim and Lee, 2011). Furthermore, temporarily disrupting the function of lateral PFC increases choice of immediate rewards over larger delayed rewards but does not impact valuation of these rewards, which indicates that impaired self-control could increase impulsivity in the DDT (Figner et al., 2010). Therefore, the negative correlation between the FCI within the frontoparietal network and discounting rates suggests that weakened intrinsic functional organization in the frontoparietal regions may induce insufficient cognitive control during deliberation (a tendency linked to impulsive choices) and thus result in greater preference for immediate reward (a sign of high impulsivity; Patton et al., 1995; Bechara, 2005).

Conversely, the dACC and AIC are the key nodes of the emotional salience network (Seeley et al., 2007; Menon and Uddin, 2010). The insula is a critical neural substrate for conscious urges that reflect an emotional state elicited by external value and corresponding bodily responses, and the insula activation can intensify one's impulsivity (Naqvi and Bechara, 2010). The dACC and AIC are coactivated during emotion-inducing tasks (Sanfey et al., 2003; Bartels and Zeki, 2004; Johnstone et al., 2006), and it has been suggested that the ACC and insular cortex engender the feeling and motivation that constitute emotion (Craig, 2002, 2009). During decision-making, the ACC and insula may be engaged during emotional processes (Sanfey et al., 2006), which are rapid and highly automatic and bias the decision-making behavior toward cognitive less demanding strategies (Botvinick, 2007; Kuo et al., 2009; Rushworth et al., 2011). Thus, the positive correlation between the FCI within the dACC–AIC network and discounting rates observed in our work suggests that enhanced intrinsic functional organization in the dACC and AIC may lead to conscious urges, bias decisions toward less cognitive demand for future planning, and thus lead to

viduals with stronger FCI between the money and time networks during the resting state may use more frequent information exchange between the two networks during the DDT, and this may result in less impulsive behavior during decision-making.

In addition to the valuation process, the choice process (e.g., self-control) also plays a pivotal role in the DDT (Luo et al., 2009; Figner et al., 2010). Our fMRI results showed that the FCIs within the frontoparietal network and within the dACC–AIC network were correlated with discounting rates in opposite directions. The anterior and dorsolateral regions of PFC, as well as posterior

and the FCI within the dACC–AIC network, suggests that the dACC–AIC network may serve as a hub linking valuation and choice processes and thus may play an essential role in generating impulsivity. The results from neural networks involved in both valuation and choice processes provide a comprehensive picture of the association between resting-state FCI of DDT-related networks and impulsivity.

The observed association between rsFC and individuals' discounting rates raised the question of whether the resting-state activity can predict an individual's impulsivity in the DDT. This was first verified in experiment I using the leave-one-out procedure. Moreover, experiment II used external validation in an independent sample and showed evidence for a significant correlation between predicted and actual discounting rates. These results indicate that the FCI-discounting rate association model was able to predict an individual's discounting rate. People's impulsivity is routinely evaluated with behavioral assessments (Kirby and Finch, 2010). The results from the two samples in our present study demonstrate that an individual's discounting rate in the DDT can be predicted by his resting-state brain activity, which provides a biomarker and a task-free assessment for impulsivity in economic decision-making.

The current work has several limitations. First, the current work used only the monetary DDT task. It is unknown whether resting-state activity can predict impulsivity in the DDT using other rewards, such as food (McClure et al., 2007), nor is it clear whether these findings generalize to impulsivity related to environmental outcomes and health (Hardisty and Weber, 2009) and impulsivity in other decision-making tasks, such as risk discounting (Christopoulos et al., 2009). Second, like most previous fMRI studies that have used graph theory, the present study only analyzed the absolute value of the functional connections (Achard and Bullmore, 2007; Liu et al., 2008; Bullmore and Sporns, 2009). It is unclear how to relate the sign of connections in graph theory (positive/negative; Fig. 6) to individual difference of impulsivity. Third, because some of the identified neural networks play additional functional roles (e.g., general cognitive abilities for the frontoparietal network; Shmash and Gray, 2008), the functional significance of these networks in the DDT is unclear. Finally, although we found an association between rsFC and impulsivity in decision-making, the mechanisms by which these patterns of rsFC lead to task-related neural activation and impulsive behavior remain unknown. These questions should be addressed in future research.

In conclusion, we found evidence for the association between impulsivity in decision-making and the rsFC between DDT-related brain regions, which showed distinct patterns for different functional networks. Moreover, the rsFC between DDT-related brain regions was able to predict behavioral impulsivity during the DDT. Our results suggest that the intrinsic functional organization of the brain may underlie individual differences in impulsivity. These findings extend our perspective on the neural basis of impulsivity and provide evidence for a biomarker of impulsivity in economic decision-making.

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