## Summary

Training can lead to long-lasting improvement in our perceptual ability, which is referred to as perceptual learning. Unraveling its neural mechanisms has proved difficult. With functional and structural magnetic resonance imaging (MRI), we addressed this issue by searching for the neural correlates of perceptual learning of face views over a long time course. Human subjects were trained to perform a face view discrimination task. Their behavioral performance and MRI signals were measured before, immediately after, and 1 month after training. We found that, across individual subjects, their behavioral learning effects correlated with the stability improvementhref shalt and strucfusiform cortex immediatelyrattercanof pencephualtleatraing og face views (Figure 1A) We also found that the ethic knegston the defise us Werne sed rites paradigm not only before training could guradiset periodeguratisal behaving ad fleace in the second seco effects. These findings fourthanfirstaling anasteristissupplesterceptual learningthat, remarkably, the simplificated patternest stability edutibalizes because face view to the long-term meastrantisting is interoretatual of the long-term meastrantistic and social also provide strong and countier fing evidence for the pivotal role of the left fusiform cortex in adaptive face processing.

Psychophysical Results Results and Discussion

Perceptual learning has been studied extensively both because of its close links to cortical plasticity and because it reflects an inherent property of our perceptual systems and thus must be studied to understand perception [1]. Not surprisingly, visual object recognition and discrimination rely critically on learning [2]. Although behavioral characteristics of

<sup>9</sup>These authors contributed equally to this work. \*Correspondence: ffang@pku.edu.cn object perceptual learning have been well recognized [3, 4], its neural mechanisms remain elusive. A popular view is that training could induce strong and focal changes in the strength and/or the selectivity of neural responses to trained stimuli. However, evidence from both functional magnetic resonance imaging (fMRI) and neurophysiology is inconclusive, even contradictory [5–10]. An alternative view is that perceptual learning introduces moderate and distributed effects that modulate a preexisting, rich, and flexible set of neural object representations [11].

Notably, two critical issues in object perceptual learning (and other kinds of visual perceptual learning) are left unaddressed. First, we know almost nothing about the mechanisms mediating long-lasting learning effects. Previous studies usually focused on neural changes immediately after training, and attempts to reveal the mechanisms did not succeed [12, 13]. For example, the learning effect with a texture segmentation task could be retained for at least 2-3 years [12]. Yotsumoto and colleagues [13] measured the dynamics of subjects' behavioral performance with the task and their V1 activation over a long time course of perceptual learning. Within the first few weeks of training, V1 activation in a subregion corresponding to the trained visual field quadrant and task performance both increased. However, while the improved performance was maintained 2 weeks after training, the V1 activation decreased to the level observed before training. Thus, the long-term mechanisms of perceptual learning are still mysterious. Second, there is no existing knowledge of neuroanatomic correlates of visual perceptual learning, although considerable progress has been made with other types of learning (e.g., music learning, language learning) [14-17]. These two issues are not only theoretically interesting, as they inform us about brain organization and limits of plasticity, but they also have significant clinical implications as they can assist the development of optimal training/rehabilitation programs [18].

To address the progra thoutve-214(lewe227(adtook229(as)dvt)-e)-3423of l

1 consisted of four phases—pretraining test (Pre), face view discrimination training, posttraining test 1 (Post1), and posttraining test 2 (Post2). Pre, Post1, and Post2 took place on the days before, immediately after, and 1 month after training (Figure 1B). During the training phase, subjects underwent eight daily training sessions (1,000 trials per session) to discriminate face views around the in-depth face orientation of 30°. In a trial, two face views with slightly different orientations were presented sequentially. Subjects were asked to make a two-alternative forced-choice (2-AFC) judgment of the orientation of the second view relative to the first (left or





[22] was used to control the two views adaptively discrimination thresholds g course, their thresholds after day 6 (Figure 1D). hophysical and MRI tests face view discrimination ,  $30^{\circ}$ , and  $60^{\circ}$ , similar to the discrimination threshantly lower for the trained 79, p < 0.001), but showed (i.e., the  $0^{\circ}$  and  $60^{\circ}$  views; This result demonstrated ng effect, which was sped up to 1 month.

easured blood-oxygenaresponding to the three h consisted of three 12 s , interleaved with three lock contained five trials. ry similar to those in the he orientation difference discrimination threshold ysical test, which caused ss blocks and tests. as to investigate whether I change associated with ect. We focused the data egions of interest (ROIs), visual cortex (EVC, Experimental included

correlation was found (all abs(r) < 0.26, p > 0.05), suggesting that these areas might not play an important role in the face learning. These results do not support the view that focal changes in the strength of neural responses to trained stimuli are critical mechanisms for perceptual learning.

The multivariate pattern analysis was a standard correlation analysis of spatial activity pattern [24]. For each ROI, we computed the correlation coefficient between the spatial activity patterns evoked by the same face view in different runs (Figure S1B). We defined the LMI for pattern correlation as [Coef (trained view posttraining) – Coef (trained view pretraining)] – [Coef (untrained views posttraining) – Coef (untrained views pretraining)]. An index significantly above zero indicates that training improves the stability of the spatial activity pattern evoked by the trained view. We found that only the IFFA exhibited a significantly positive index at both Post1 and Post2 (both t(17) > 3.16, p < 0.05) (Figure 2B), demonstrating that the improved stability of the spatial activity pattern in the IFFA persisted over the long time course of perceptual learning.

The MVPA performed above used a single statistical threshold to define the ROIs, which usually had different numbers of voxels. It is likely that the correlation coefficient in an ROI was affected by its voxel number. To rule out this explanation, for each subject, we adjusted the thresholds for the rFFA, rSTS, ISTS, rOFA, IOFA, and EVC individually to ensure that they had the same voxel numbers as the IFFA (the mean voxel number in the IFFA across subjects was 73). With these redefined ROIs, we performed the same analysis. None of these areas exhibited a significantly positive LMI at Post1 and Post2 after ROI resizing (Figure S2A).

To further evaluate the role of the improved IFFA pattern sta-



(Figure S2B). The null result in EVC also excludes the possibility that the observed learning effect here could be due to some kind of feature learning, which is consistent with our previous psychophysical findings [19].

## **Cortical Thickness Results**

The second purpose of experiment 1 was to explore the structural correlates of the face view discrimination learning. Because of the close relationship between brain structure and brain function, it is natural to ask whether or not, after training, there was any structural change accompanying the functional change in the IFFA, as well as in the other ROIs. In each test phase, subjects' structural images were acquired before the fMRI runs. We measured the thickness of the cortex before and after training, as cortical thickness is a sensitive index of brain plasticity [31, 32]. To remove the fluctuation in the average thickness of the whole cortex among different scanning sessions, we subtracted the average thickness in each session for each subject from the original thickness of each vertex. All the subsequent analyses were based on the relative cortical thickness. We found that the average cortical thickness for each ROI did not change after training (all t(17) < 1.40, p > 0.05) (Figure 3A). We then searched the whole cortical surface and still found no reliable change in cortical thickness. This result was in sharp contrast with previous learning studies

Figure 3. Cortical Thickness Results of Experiment 1

(A) Cortical thickness of the ROIs at Pre, Post1, and Post2. Cortical thickness analyses were performed with the relative cortical thickness (i.e., the original thickness of each vertex minus the average thickness of the whole cortex). A positive or negative value of a ROI thickness means that the ROI is thicker or thinner than the average thickness of the whole cortex. Error bars indicate 1 SEM.

(B) Correlation between the cortical thickness of the IFFA and the average psychophysical learning index of Post1 and Post2 across subjects.

(C) Regions with a significant correlation between cortical thickness and the average psychophysical learning index across subjects. Positive correlations are shown in hot colors, and negative correlations in cool colors. Images were thresholded at p < 0.01 (uncorrected) and cluster size >100 vertices. See also Figure S3.

in other functional domains, in which regional increases in cortical thickness were detected after memory, music, language, or navigation training [14–17].

Although there was little cortical thickness change after training, a surprising finding in our analysis was that the cortical thickness of the IFFA before training (at Pre) was predictive of the behavioral learning effect across individual subjects. Figure 3B shows an inverse correlation between the cortical thickness of the IFFA and the average psychophysical learning index of Post1 and Post2 (r = -0.70, p < 0.01). In other words, the thinner the cortex of the

IFFA, the greater the behavioral learning effect. Other ROIs did not show a significant correlation (all abs(r) < 0.49, p > 0.05) (Figure S3B). It is important to note that, before training, the cortical thickness of the IFFA did not correlate with the discrimination threshold for the trained view (r = -0.15, p = 0.565). This observation ruled out a possible explanation that the thinner IFFA cortex caused a lower performance before training and therefore allowed more room for improvement. We further searched the entire cortical surface and computed correlations between the cortical thickness of each vertex and the average psychophysical learning index. Figure 3C shows vertex-based, correlational maps across the entire cortical surface depicting the topography of significant correlations at a statistical threshold of p < 0.01 (uncorrected). The area showing the strongest correlation was in the left fusiform cortex (peak vertex: r = -0.73, p < 0.001; Talairach coordinates: -50, -48, -17), which was strikingly overlapped with the IFFA defined by the functional localizer (Talairach coordinates: -41, -46, -14). Two additional regions were detected by this analysis, the right parahippocampal gyrus (peak vertex: r = 0.71, p < 0.001; Talairach coordinates:13, -49, 4) and the right anterior cingulate cortex (peak vertex: r = -0.70, p < 0.01; Talairach coordinates: 6, 31, -5). Thus, this unbiased, post hoc approach supported the specificity of our ROI findings to the IFFA.



Figure 4. Correlation between the Cortical Thickness and the Average LMI for Pattern Correlation of Post1 and Post2at the IFFA across Subjects See also Figure S4.

The inverse relationship between the cortical thickness of the IFFA and the behavioral learning effect is counterintuitive. It challenges a common assumption that a greater cortical thickness is associated with better processing efficacy of that region [18]. Shaw et al. [31] found a negative correlation between IQ and cortical thickness in early childhood. In agreement with our study, children who had thinner cortex in frontal regions gained more in an intelligence measure. Although the microstructure and cellular events contributing to cortical thickness are largely unknown, a compelling explanation of our finding is cortical pruning. Cortical pruning is a process of removing inefficient synapses and neurons, especially during adolescence [33]. It is conceivable that thinner cortex as a consequence of cortical pruning leads to more efficient processing, as well as strong learning ability. We further speculate that the thinner IFFA cortex, after training, could become more functionally stable because of fewer inefficient synapses and neurons, consequently leading to a greater behavioral learning effect. The significant correlation between the cortical thickness and the average LMI for pattern correlation of Post1 and Post2 at the IFFA (r = -0.49, p < 0.05) (Figure 4) provides tentative evidence for this idea and suggests a close correspondence between cortical thickness and increased brain function.

## **Control Experiments**

Experiments 2 and 3 were designed to investigate whether the neural changes found in experiment 1 depend on subjects' attention to the face stimuli and the face view discrimination task. These two experiments were identical to experiment 1 except that subjects performed a luminance discrimination task at fixation (instead of the face view discrimination task) in the fMRI tests (experiment 2) or during training (experiment 3). In an fMRI test trial of experiment 2, fixation point was presented at two different luminances during the presentation intervals of two face views. Subjects needed to judge which interval contained a brighter fixation point. In a training trial of experiment 3, subjects performed the same task.

In experiment 2, the face view discrimination training led to a similar behavioral learning effect as that in experiment 1. Relative to Pre, the discrimination thresholds at Post1 and Post2

were significantly lower for the  $30^{\circ}$  view (both t(14) > 7.43, p < 0.001), but showed little change for the  $0^{\circ}$  and  $60^{\circ}$  views (all t(14) < 3.06, p > 0.05) (Figure S2C). In experiment 3, because subjects were trained with the luminance discrimination task, after training, subjects' face view discrimination performance showed little improvement for all the three face views (all t(11) < 1.58, p > 0.05) (Figure S2D). In both experiments, LMIs for BOLD amplitude and pattern correlation were not significantly greater than zero in any ROI at Post1 and Post2 (experiment 2: all t(14) < 2.36, p > 0.05, Figure S2C). These results demonstrated that only exposure to the trained view (without performing the face view discrimination task) during training and test was not able to induce the neural changes observed in experiment 1.

We also performed the cortical thickness analysis with the anatomical data in experiments 2 and 3. In experiment 2, a significant correlation between the cortical thickness and the average psychophysical learning index of Post1 and Post2 was found only in the IFFA, but not other ROIs (Figure S4A). Before training, the cortical thickness of the IFFA did not correlate with the discrimination threshold for the trained view in both experiment 2 (r = -0.11, p = 0.685) and experiment 3 (r = -0.10, p = 0.753). The cortical thickness analyses above were performed with the relative cortical thickness data. It should be noted that these findings can be replicated with the raw cortical thickness data (Figures S4B and S4C). Taken together, these results strengthen the cortical thickness conclusion in experiment 1.

# The Role of the Left Fusiform Cortex in Face Processing

Our finding that both the function and structure of the left fusiform cortex are closely associated with the long-lasting effect of face view discrimination learning has important implications on face processing and its plasticity. Using fMRI adaptation and MVPA methods, previous studies [34, 35] have demonstrated that face views are represented in the FFA and STS. But it was unknown whether and how the neural representation of face views can be shaped by visual experience. The current study provides strong and converging evidence that the face view representation is plastic even in the adult brain and suggests that the IFFA plays a pivotal role in adaptive face view processing.

Our findings shed new light on the hemispheric asymmetry in face processing. Although the essential role of the rFFA in face recognition has been extensively documented in literature over the past few decades [36], we still know little about the exact function of the IFFA and the functional difference between these two areas. Recently, Meng and colleagues [37] proposed that the IFFA performs the graded analyses of faces, while the rFFA performs the categorical analyses. They also found that the IFFA is more susceptible to contextual information than the rFFA. Based on the findings of the current study, we argue that the IFFA is more susceptible to perceptual learning and is more plastic than the rFFA. Our view is in line with two recent event-related potential studies. First, Rossion et al. [38] found that training with novel objects (i.e., Greebles) led to a left-lateralized facelike N170 response. Second, Su et al. [39] showed that perceptual learning could shorten theN170 latency only at the left occipital-temporal area. Why is the IFFA more plastic than the rFFA? A possible explanation is that the rFFA in the adult brain has fully developed and its function is fixed after maturation [40], while the IFFA is still open to changes for adapting to the dynamic visual world.

It should be noted that, although we emphasize the importance of the IFFA in perceptual learning of faces, we cannot deny potential contributions from other cortical areas (e.g., rFFA). In future research, it would be interesting to examine whether there is any brain network serving for the face perceptual learning. It is possible that the IFFA is a part of this network and the observed neural effect is most prominent in this area. Also, other kinds of face learning (e.g., identity learning) should be investigated to further clarify the plasticity of the IFFA and other related areas, as well as the brain network for face learning.

### Supplemental Information

Supplemental Information includes Supplemental Experimental Procedures and four figures and can be found with this article online at http://dx.doi.org/ 10.1016/j.cub.2013.12.028.

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