



The subjective experience of emotion: a fearful view

Joseph E LeDoux¹ and Stefan G Hofmann²

We argue that subjective emotional experience, the feeling, is the essence of an emotion, and that objective manifestations in behavior and in body or brain physiology are, at best, indirect indicators of these inner experiences. As a result, the most direct way to assess conscious emotional feelings is through verbal self-report. This creates a methodological barrier to studies of conscious feelings in animals. While the behavioral and physiological responses are not ‘emotions,’ they contribute to emotions indirectly, and sometimes profoundly. Whether non-verbal animals have emotional experiences is a difficult, maybe impossible, question to answer in the positive or negative. But because behavioral and physiological responses are important contributors to emotions, and the circuits underlying these are highly conserved, studies of animals have an important role in understanding how emotions are expressed and regulated in the brain. Conflation of circuits that directly give rise to conscious emotional feelings with circuits that indirectly influences these conscious feelings has hampered progress in efforts to understand emotions, and also to understand and to develop treatments for emotional disorders. Recognition of differences in these circuits will allow research in animals to have a lasting impact on understanding of human emotions as research goes forward.

Addresses

¹ 4 Washington Place, Center for Neural Science, New York University, United States

² Department of Psychological and Brain Sciences, Boston University, United States

Corresponding author: LeDoux, Joseph E (ledoux@cns.nyu.edu)

Current Opinion in Behavioral Sciences 2018, 19:67–72

This review comes from a themed issue on **Emotion-cognition interactions**

Edited by **Mara Mather** and **Michael Fanselow**

For a complete overview see the [Issue](#) and the [Editorial](#)

Available online 16th November 2017

<http://dx.doi.org/10.1016/j.cobeha.2017.09.011>

2352-1546/© 2017 Elsevier Ltd. All rights reserved.

The English word ‘emotion’ is based on the Latin *emovere*, which means “to move away from.” When first applied to psychological events in the 17th century, it pointed to an excited state of mind that causes movement (behavior). This is still the most common meaning of emotion in everyday vernacular speech. But in scientific

discussions, the noun ‘emotion,’ or its adjectival form ‘emotional,’ are variably used to refer to subjective experiences, behavioral movements, physiological responses, and/or cognitions that contribute to any of the above. Given this multiplicity of referents, it is not surprising that there is debate and confusion about the nature of emotions [1–11].

In this article, we argue that restriction of the use of the term ‘emotion’ to subjective experiences, and use of other terminology to describe objective responses that are often correlated with emotional experiences would eliminate much of the conceptual confusion. In making this case, we discuss several different conceptual approaches to subjective emotional experiences and the brain circuits proposed to underlie such experiences in these approaches. Because the emotion *fear* has been studied more than other emotions, especially in relation to brain circuits, and has been the center of much of the controversy about the nature of subjectively experienced emotion, we focus on it in our discussion. Because the argument made in this article applies to both fear and anxiety, we will not distinguish these two terms (for a discussion of the difference see [6]).

Measuring subjective experiences

Before considering different approaches to subjective experiences, it is important to discuss how these unobservable private events are studied. Scientific assessments of inner experiences require some form of self-reporting [12,13]. People can typically give either a verbal or a nonverbal report of information to which they have introspective access, but cannot provide a verbal report of information that is only processed non-consciously [6,14,15]. Fractures between conscious and nonconscious processes by differences between verbal and non-verbal responses have thus played a key role in studies of introspective awareness in humans. While other methods of reporting that do not require verbal report have been proposed [16–18], these also depend on introspection [15].

Verbal self-report remains the gold standard in studies of consciousness. It is most suitable for assessing the content of immediate experiences rather than remembered experiences [14,19] and is less useful for assessing the motivations underlying actions since these are often not consciously available and verbalizable [20,21]. Since non-verbal reporting is the only option in non-verbal (non-human) organisms, determining whether other animals have conscious, subjective experiences is difficult [6].

Contemporary views of subjective emotional experiences in relation to brain circuits

Four contemporary approaches to subjective emotional experiences in the brain, and the historical roots of each, are described below. Included are approaches in the traditions of Charles Darwin, William James, behaviorism, and cognitive psychology.

1. The Neuro-Darwinian Approach: Subjective Fear is an Innate State of Mind Inherited from Animal Ancestors. In his treatise on human and animal emotion, Darwin defined emotions as innate “states of mind” that humans have inherited from animal ancestors, and that, when aroused, cause the expression of so-called emotional behaviors [22]. This is consistent with the original 17th century use of the term emotion mentioned above. It is still the commonsense view of emotion that most people have, and also underlie Ekman’s widely cited basic emotions theory [23]. A contemporary neuroscience proponent of the traditional Darwinian view is Jaak Panksepp, who views emotions as subjective feelings that emerge from a subcortical neural circuit that is highly conserved across mammals, including humans [24,25]. The circuit is centered on the amygdala and related subcortical areas [24]. In Panksepp’s formulation, the amygdala circuit, when activated by a threat to well being, both gives rise to fearful feelings (subjective feelings of fear) and controls innate behaviors and supporting physiological responses that help the organism defend against harm. Cognitive elaboration of subcortical fear by higher-cortical prefrontal circuits makes possible introspection and verbal reports of fear in humans. But the core of fear is the inherited mental state arising from the subcortical circuit.

Problematic for this view is evidence suggesting that the experience of fear is not embodied in the amygdala. Specifically, studies in humans show that the amygdala can respond to threats without the person knowing the threat is present and without feeling fear, and other studies show that fear can be experienced when the amygdala is damaged [26,27]; for review see [6,7]. Also, medications can change behavioral responses to threats without changing subjective feelings of fear [28,29]. Thus, while the amygdala controls behavioral responses to threats it does not seem to be directly responsible for the subjective experience of fear. One could argue that perhaps other subcortical areas are responsible. But the emphasis in the literature has been on the amygdala. Moreover, the other hypothetical subcortical circuits would need to be identified before the role can be evaluated.

2. The Neuro-Jamesian Approach: Subjective Fear is a Consequence of Feedback from Body Responses. William James famously argued that fear and other emotional experiences result by way of feedback from the act of responding [30]. Thus, contrary to the commonsense

view, fear does not cause the responses but instead results as a consequence of the responses. Modern versions of this theory by Antonio Damasio [31,32] and A.D. Craig [33,34] have proposed that fearful and other emotional feelings are the result of activity in body sensing circuits in the neocortex (somato-sensory and/or insula areas) that represent body states, such as those triggered when threats activate amygdala circuits. Initially, Damasio emphasized cortical body sensing circuits but more recently has emphasized brainstem circuits [32]. As in Panksepp’s theory, cognitive elaboration by higher-cortical circuits allows introspection and self-reporting about these states in Damasio’s theory. While these circuits clearly represent body states, convincing evidence that these representations are the main causes of emotional experiences is lacking [6]. We propose below that body feedback makes important contributions to emotional experiences, but as modulators rather than as direct causes.

3. The Neuro-Behaviorist Approach: Subjective Fear is a Folk Psychological Construct that Should be Replaced by a Scientific Explanation. In the early 20th century, behaviorists eliminated mental states from the causal chain between external stimuli and behavioral responses. However, they retained mental state terms when describing the empirical relation between stimuli and responses. For example, ‘fear’ was used to characterize the relation between threats and defensive behaviors [35]. With the rise of physiological approaches to behavior in the middle of the 20th century, fear became a hypothetical physiological state (central state), that connects threats with defensive behaviors [36]. The intended purpose of this approach was to satisfy the behaviorist constraint against using subjective explanations of behavior. Ultimately, the amygdala emerged as the locus of the central fear state [37,38]. The central state model has been popular in behavioral neuroscience [39–42]. Many who call upon the central state view today are ambivalent about the extent to which fear means subjective fear or a non-subjective state. But adherents of a strong version of the position argue that subjective, conscious fear is an inaccurate scientific construct that can be replaced with a more rigorous scientific notion of fear as a non-subjective state of the amygdala-centered circuits [42]. The well-known lack of correlation between verbal reports and amygdala activity in humans [43–45] is, in this view, due to the lack of access to amygdala activity by cognitive-based introspection, and hence verbal report. In short, verbal report of subjective experience is viewed as a less desirable way of assessing fear than simply measuring amygdala-controlled responses.

The mechanistic reason a threat elicits defense responses is because it activates cells and synapses in circuits that control those responses. The fear construct is superfluous in this context [6,7,28,46]. It adds conceptual baggage

that reifies fear and imbues the circuit with the exact phenomenological properties that are trying to be avoided [47]. The way to avoid these problems is by using non-subjective terminology to name the non-subjective central states. Such states, for example, can be referred in terms of neural activity in a defensive survival circuit (rather than in a fear circuit) [4]. This solves part of the problem but leaves open the question of how conscious fear comes about. The neuro-cognitive approach addresses this issue.

4. The Neuro-Cognitive Approach: Subjective Fear Emerges from Higher-Order Processing. Cognitive approaches to emotion arose in the 1960s in an effort to address certain criticisms of the Jamesian approach, and to overcome the behaviorist's restrictions on inner explanations. Stanley Schachter proposed that emotional experiences result when we interpret physiological arousal in the brain and/or body in light of cognitive assessments of social situations, allowing the categorization and labeling of the experience [48,49]. While there have been a number of cognitive approaches to emotion [50–53], recent theorizing has emphasized that emotional experiences are cognitive constructions based on conceptualizations of situations [11,54–56] or higher-order states that emerge as a result of the cognitive integration in working memory of diverse sources of information from within the brain and body [6,7,37,57,58]. Both of these positions reject the idea that emotional experiences arise from subcortical circuits (views of the Neuro-Darwinian and Neuro-Jamesian approaches) and also reject the idea that the subjective experience is a non-scientific construct (Neuro-Behaviorist approach).

The higher-order approach builds on recent developments in the science and philosophy of consciousness (e.g. [7,59–61]), arguing that a general network of cognitions underlies both cognitive and emotional states of consciousness. In higher-order emotion theory, what distinguishes cognitive and emotional states of consciousness, and different kinds of emotional states, is the kind of inputs processed. The subcortical fear circuit of the other models becomes a defensive survival circuit that detects and responds to threats. The consequences of defensive survival circuit activation (brain arousal, body feedback, etc.) contribute indirectly to the experience of fear but do not determine it. Fear is the cognitive assessment that you are in harms way, a view that allows fear to arise from activity in any survival circuits (fear of being harmed by starvation, dehydration, hypothermia, reproductive isolation, and so forth), or by existential concerns (such as fear of the eventuality of death or the meaninglessness of your life), in addition to predatory-related dangers that trigger the defensive survival circuit.

5: Summary of the Four Approaches. The neuro-Darwinian, neuro-Jamesian, and neuro-behaviorist approaches all

imply that if through psychotherapy or pharmaceutical treatment you successfully reduce amygdala activity, as measured by amygdala-mediated behavioral and/or physiological responses, pathological fear should be ameliorated. But different reasons underlie this conclusion in each approach. In the neuro-Darwinian approach, treatments that weaken defensive behavior should treat fear problems by altering amygdala activity underlying both behavior and feelings. In the neuro-Jamesian approach, treatments that weaken defensive behavior by altering amygdala activity should treat fear problems because the behaviors, being weaker, should give rise to less body feedback and thus weaker subjective feelings. In the neuro-behaviorist approach, altering amygdala activity through behavioral (CBT) or pharmaceutical treatments should treat pathological fear because the problems are due to the non-subjective amygdala state. Once this changes, subjective fear might also change because it is an indirect cognitive readout of amygdala activity. However, whether self-reported feelings change is less important than whether defensive behavior changes since the latter is a direct readout of amygdala activity. A major challenge for these approaches is accounting for the failure of studies using defensive behaviors in animals as means of drug-discovery to generate novel agents with clinical efficacy in treating fear and anxiety [6,28,29,62].

Conscious feelings in clinical assessment

A major reason why people seek the help of mental health specialists is because they feel bad and want to feel better. A treatment that reduces behavior (freezing, behavioral timidity, avoidance) and physiology (hyperarousal) but does not diminish subjectively reported fearful feelings is not likely to be viewed as a satisfactory outcome by the afflicted person.

In the contemporary cognitive therapy literature, self-reports, and the subjective experiences that these reports reflect, have not been given much credence. This reflects the influence of behaviorism in the development of both traditional behavioral therapy [63] and cognitive behavioral therapy (CBT) [64], as well as findings suggesting that behavioral physiological responses related to fear or anxiety in humans are poorly correlated [43–45]. A recent argument has been made for more attention to verbal reports of subjective experiences, but mainly to validate non-subjective measures [65]. This reflects the dominance of the amygdala fear center view described above in clinical concepts. If behavioral, physiological and subjective responses are all products of the same circuit, one response is as good as another as a way to judge clinical outcome. This logic is also apparent in the NIHM RDoC initiative, which treats subjective reports as just another measure of fear. Kozak and Cuthbert [66], for example, note that “. . . the RDoC approach accords self-report data no special precedence among different measurement

classes, any of which might contribute to a nomological net (p. 292).”

An important advantage of the neuro-cognitive approach to fear is that it accounts for the two troubling observations mentioned above that are not easily handled by fear-center views: subjective fear does not require the amygdala, and medications that target the amygdala do not necessarily relieve subjective fear. It would seem that the neuro-cognitive approach should mesh well with modern CBT. However, despite the centrality of introspection and cognitive processes in human mental life, as noted above, subjective report never reached a central status in CBT.

Self-report has always been seen as an easy but unreliable way to measure anxiety and fear. For example, Zoellner and Foa [67] agree with Kozak and Cuthbert that self-reports do not have any special status, noting that “self-report is one piece of the net in terms of studying a phenomenon.” But they nevertheless go on to say that self-reports “should be considered potential valid measures of key constructs, some of which cannot be measured in any other way, and sometimes the best measure of the construct of interest (p. 334).”

Today, CBT therapists routinely use subjective patient report as part of treatment. However, its value continues to be debated. Some argue that self-reported fear is an inadequate measure of the patient’s emotional state because reduction in reported fear during exposure sessions does not predict long-term treatment effects [68,69]. While this finding is consistent with the neuro-behaviorist (non-subjective) amygdala fear state model described above, the evidence presented is not especially strong, and other findings directly contradict the conclusion. Specifically, more recent clinical trials examining the augmentation effect of D-cycloserine (DCS), a partial agonist at the glycine recognition site of the NMDA receptor complex, show the DCS enhances exposure therapy primarily in those people who report a clear reduction of reported fear during the exposures [70,71]. This suggests that the reduction in reported fear ratings during an exposure session is, in fact, a crucial factor that determines the success of treatment [72]. Also intriguing is the recent finding that Pavlovian conditioned threat responses can be extinguished without the arousal of conscious fear [73].

We argue that subjective reports of fear and anxiety are not unreliable proxy measures of fear and anxiety, as suggested by the neuro-behaviorist model. Rather, they are the best direct, and, at least so far, the only measures easily accessible for clinicians to assess the efficacy of treatment. If this is correct, therapists need to carefully listen to the clients’ report about their emotional states, further develop reliable assessment instruments of

subjective reports, and utilize verbal strategies to elicit and modify them. However, we suggest that this be done after other approaches, such as CBT or pharmaceutical treatments, to first dampen the amygdala central state, which indirectly affects subjective experience. We also recommend that future research re-examine the relationship between subjective report, behaviors, and physiological responses of fear and anxiety in order to explain the reasons for the dissociation between these measures.

Conflict of interest statement

Nothing declared.

Funding

Funded in part by the James S. McDonnell Foundation 21st Century Science Initiative in Understanding Human Cognition – Special Initiative.

References

1. Candland DK, Fell JP, Keen E, Leshner AI, Tarpay RM: *Emotion*. Belmont, CA: Wadsworth Publishing Company, Inc.; 1977.
2. Ekman P, Davidson R: *The Nature of Emotion: Fundamental Questions*. New York: Oxford University Press; 1994.
3. Scarantino A: **Functional specialization does not require a one-to-one mapping between brain regions and emotions**. *Behav Brain Sci* 2012, **35**:161-162.
4. LeDoux J: **Rethinking the emotional brain**. *Neuron* 2012, **73**: 653-676.
5. LeDoux JE: **Coming to terms with fear**. *Proc Natl Acad Sci U S A* 2014, **111**:2871-2878.
6. LeDoux JE: *Anxious: Using the Brain to Understand and Treat Fear and Anxiety*. New York: Viking; 2015.
7. LeDoux JE, Brown R: **A higher-order theory of emotional consciousness**. *Proc Natl Acad Sci U S A* 2017, **114**: E2016-E2025.
8. Kragel PA, LaBar KS: **Decoding the nature of emotion in the brain**. *Trends Cogn Sci* 2016.
9. Kuppens P, Tuerlinckx F, Yik M, Koval P, Coosemans J, Zeng KJ, Russell JA: **The relation between valence and arousal in subjective experience varies with personality and culture**. *J Pers* 2016.
10. Touroutoglou A, Lindquist KA, Dickerson BC, Barrett LF: **Intrinsic connectivity in the human brain does not reveal networks for ‘basic’ emotions**. *Soc Cogn Affect Neurosci* 2015, **10**:1257-1265.
11. Barrett LF: *How Emotions are Made*. New York: Houghton Mifflin Harcourt; 2017.
12. Weiskrantz L: *Consciousness Lost and Found: A Neuropsychological Exploration*. New York: Oxford University Press; 1997.
13. Overgaard M, Sandberg K: **Kinds of access: different methods for report reveal different kinds of metacognitive access**. In *The Cognitive Neuroscience of Metacognition*. Edited by Fleming SM, Frith CD. Springer-Verlag Berlin Heidelberg; 2014:67-86.
14. Ericsson KA, Simon H: *Protocol Analysis: Verbal Reports as Data*. Cambridge, MA: MIT Press; 1993.
15. Seth AK: **Post-decision wagering measures metacognitive content, not sensory consciousness**. *Conscious Cogn* 2008, **17**:981-983.

16. Persaud N, McLeod P, Cowey A: **Post-decision wagering objectively measures awareness.** *Nat Neurosci* 2007, **10**: 257-261.
17. Lau HC, Passingham RE: **Relative blindsight in normal observers and the neural correlate of visual consciousness.** *Proc Natl Acad Sci U S A* 2006, **103**:18763-18768.
18. Maniscalco B, Lau H: **The signal processing architecture underlying subjective reports of sensory awareness.** *Neurosci Conscious* 2016:2016.
19. Wilson TD: **The proper protocol: validity and completeness of verbal reports.** *Psychol Sci* 1994, **5**:249-252.
20. Nisbett RE, Wilson TD: **Telling more than we can know: verbal reports on mental processes.** *Psychol Rev* 1977, **84**:231-259.
21. Hassin RR, Uleman JS, Bargh JA (Eds): *The New Unconscious*. New York: Oxford University Press; 2005.
22. Darwin C: *The Expression of the Emotions in Man and Animals*. London: Fontana Press; 1872.
23. Ekman P: **Are there basic emotions?** *Psychol Rev* 1992, **99**: 550-553.
24. Panksepp J: *Affective Neuroscience*. New York: Oxford U. Press; 1998.
25. Panksepp J: **The cross-mammalian neurophenomenology of primal emotional affects: from animal feelings to human therapeutics.** *J Comp Neurol* 2016, **524**:1624-1635.
26. Feinstein JS, Buzza C, Hurlmann R, Follmer RL, Dahdaleh NS, Coryell WH, Welsh MJ, Tranel D, Wemmie JA: **Fear and panic in humans with bilateral amygdala damage.** *Nat Neurosci* 2013, **16**:270-272.
27. Anderson AK, Phelps EA: **Is the human amygdala critical for the subjective experience of emotion? Evidence of intact dispositional affect in patients with amygdala lesions.** *J Cogn Neurosci* 2002, **14**:709-720.
28. LeDoux JE, Pine DS: **Using neuroscience to help understand fear and anxiety: a two-system framework.** *Am J Psychiatry* 2016, **173**:1083-1093.
29. Griebel G, Holmes A: **50 years of hurdles and hope in anxiolytic drug discovery.** *Nat Rev Drug Discov* 2013, **12**:667-687.
30. James W: **What is an emotion?** *Mind* 1884, **9**:188-205.
31. Damasio A: *Descartes's Error: Emotion, Reason, and the Human Brain*. New York: Gosset/Putnam; 1994.
32. Damasio A, Carvalho GB: **The nature of feelings: evolutionary and neurobiological origins.** *Nat Rev Neurosci* 2013, **14**:143-152.
33. Craig AD: **How do you feel — now? The anterior insula and human awareness.** *Nat Rev Neurosci* 2009, **10**:59-70.
34. Strigo IA, Craig AD: **Interoception, homeostatic emotions and sympathovagal balance.** *Philos Trans R Soc Lond B Biol Sci* 2016:371.
35. Brown JS, Farber IE: **Emotions conceptualized as intervening variables — with suggestions toward a theory of frustration.** *Psychol Bull* 1951, **48**:465-495.
36. Rescorla RA, Solomon RL: **Two process learning theory: relationships between Pavlovian conditioning and instrumental learning.** *Psychol Rev* 1967, **74**:151-182.
37. LeDoux JE: *The Emotional Brain*. New York: Simon and Schuster; 1996.
38. Davis M: **The role of the amygdala in conditioned fear.** In *The Amygdala: Neurobiological Aspects of Emotion, Memory, and Mental Dysfunction*. Edited by Aggleton JP. Wiley-Liss, Inc; 1992:255-306.
39. Rosen JB, Schulkin J: **From normal fear to pathological anxiety.** *Psychol Rev* 1998, **105**:325-350.
40. Adolphs R: **The biology of fear.** *Curr Biol* 2013, **23**:R79-R93.
41. Anderson DJ, Adolphs R: **A framework for studying emotions across species.** *Cell* 2014, **157**:187-200.
42. Perusini JN, Fanselow MS: **Neurobehavioral perspectives on the distinction between fear and anxiety.** *Learn Mem* 2015, **22**:417-425.
43. Lang PJ: **Fear reduction and fear behavior: problems in treating a construct.** In *Research in Psychotherapy*, vol 3. Edited by Schlien JM. American Psychological Association; 1968:90-103.
44. Hodgson R, Rachman S: **II. Desynchrony in measures of fear.** *Behav Res Ther* 1974, **12**:319-326.
45. Rachman S, Hodgson R: **I. Synchrony and desynchrony in fear and avoidance.** *Behav Res Ther* 1974, **12**:311-318.
46. LeDoux JE: **Semantics, surplus meaning, and the science of fear.** *Trends Cogn Sci* 2017.
47. Marx MH: **Intervening variable or hypothetical construct?** *Psychol Rev* 1951, **58**:235-247.
48. Schachter S: **Cognition and centralist-peripheralist controversies in motivation and emotion.** In *Handbook of Psychobiology*. Edited by Gazzaniga MS, Blakemore CB. Academic Press; 1975:529-564.
49. Schachter S, Singer JE: **Cognitive, social, and physiological determinants of emotional state.** *Psychol Rev* 1962, **69**:379-399.
50. Mandler G: *Mind and Body: The Psychology of Emotion and Stress*. New York: Norton; 1984.
51. Frijda NH, Scherer KR: **Emotion definition (psychological perspectives).** In *Oxford Companion to Emotion and the Affective Sciences*. Edited by Sander D, Scherer KR. Oxford University Press; 2009:142-143.
52. Ortony A, Clore GL: **Emotions, moods, and conscious awareness.** *Cogn Emot* 1989, **3**:125-137.
53. Lewis MD: **Self-organising cognitive appraisals.** *Cogn Emot* 1996, **10**:1-26.
54. Russell JA: **Core affect and the psychological construction of emotion.** *Psychol Rev* 2003, **110**:145-172.
55. Barrett LF: **Solving the emotion paradox: categorization and the experience of emotion.** *Pers Soc Psychol Rev* 2006, **10**:20-46.
56. Barrett LF, Russell JA (Eds): *The Psychological Construction of Emotion*. New York: Guilford Press; 2015.
57. LeDoux JE: *Synaptic Self: How Our Brains Become Who We Are*. New York: Viking; 2002.
58. LeDoux JE: **Emotional colouration of consciousness: how feelings come about.** In *Frontiers of Consciousness: Chichele Lectures*. Edited by Weiskrantz L, Davies M. Oxford University Press; 2008:69-130.
59. Dehaene S: *Consciousness and the Brain: Deciphering how the Brain Codes Our Thoughts*. New York: Penguin Books; 2014.
60. Rosenthal DM: *Consciousness and Mind*. Oxford: Oxford University Press; 2005.
61. Lau H, Rosenthal D: **Empirical support for higher-order theories of conscious awareness.** *Trends Cogn Sci* 2011, **15**:365-373.
62. Miller G: **Is pharma running out of brainy ideas?** *Science* 2010, **329**:502-504.
63. Levis DJ: **The negative impact of the cognitive movement on the continued growth of the behavior therapy movement: a historical perspective.** *Genet Soc Gen Psychol Monogr* 1999, **125**:157-171.
64. Hofmann SG, Asmundson GJ, Beck AT: **The science of cognitive therapy.** *Behav Ther* 2013, **44**:199-212.
65. Hamm AO, Richter J, Pane-Farre C, Westphal D, Wittchen HU, Vossbeck-Elsebusch AN, Gerlach AL, Gloster AT, Strohle A, Lang T et al.: **Panic disorder with agoraphobia from a behavioral neuroscience perspective: applying the research principles formulated by the Research Domain Criteria (RDoC) initiative.** *Psychophysiology* 2016, **53**:312-322.

66. Kozak MJ, Cuthbert BN: **The NIMH Research Domain Criteria Initiative: background, issues, and pragmatics.** *Psychophysiology* 2016, **53**:286-297.
67. Zoellner LA, Foa EB: **Applying Research Domain Criteria (RDoC) to the study of fear and anxiety: a critical comment.** *Psychophysiology* 2016, **53**:332-335.
68. Craske MG, Kircanski K, Zelikowsky M, Mystkowski J, Chowdhury N, Baker A: **Optimizing inhibitory learning during exposure therapy.** *Behav Res Ther* 2008, **46**:5-27.
69. Craske MG, Treanor M, Conway CC, Zbozinek T, Vervliet B: **Maximizing exposure therapy: an inhibitory learning approach.** *Behav Res Ther* 2014, **58**:10-23.
70. Smits JA, Rosenfield D, Otto MW, Marques L, Davis ML, Meuret AE, Simon NM, Pollack MH, Hofmann SG: **D-cycloserine enhancement of exposure therapy for social anxiety disorder depends on the success of exposure sessions.** *J Psychiatr Res* 2013, **47**:1455-1461.
71. Smits JA, Rosenfield D, Otto MW, Powers MB, Hofmann SG, Telch MJ, Pollack MH, Tart CD: **D-cycloserine enhancement of fear extinction is specific to successful exposure sessions: evidence from the treatment of height phobia.** *Biol Psychiatry* 2013, **73**:1054-1058.
72. Hofmann SG: **D-cycloserine for treating anxiety disorders: making good exposures better and bad exposures worse.** *Depress Anxiety* 2014, **31**:175-177.
73. Koizumi A, Amano K, Cortese A, Shibata K, Yoshida W, Seymour B, Kawato M, Lau H: **Fear reduction without fear through reinforcement of neural activity that bypasses conscious exposure.** *Nat Human Behav* 2016, **1**:0006.