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# Editorial: Case reports in anxiety and stress

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#### Editorial on the Research Topic Case reports in anxiety and stress

Anxiety disorders are a heterogenous group of mental illnesses characterized by fear or anxiety that is excessive and difficult to control, and which is associated with specific maladaptive behaviors (1, 2). Anxiety disorders are the most common form of mental illness, affecting almost 46 million people worldwide (3). The global burden of anxiety disorders has increased by around 50% in the past three decades, despite no significant change in their incidence (4). This is due to the chronic nature of these disorders: only about 60–70% of patients respond to treatment, and of these, up to two-thirds experience relapses (5, 6).

Anxiety disorders can often be triggered or exacerbated by stressful life events (7, 8). This is due to the effects of both acute and chronic stress on specific neurotransmitters, neurohormones, and the expression of specific genes (9-11). Stress-related disorders such as post-traumatic stress disorder (PTSD) are highly comorbid with anxiety disorders (12). These groups of disorders share common vulnerability genes and molecular pathways (13), and exposure to traumatic stressors can trigger not just PTSD, but any of the anxiety disorders (14). This was reflected in earlier classifications that placed anxiety and stress-related disorders in the same general category (15). Though recent research has identified key differences between them (16), their overlap remains both conceptually and clinically significant (17).

From a historical perspective, anxiety and stress-related disorders have often been viewed as "neurotic disorders", arising from unresolved psychological conflicts, and requiring resolution through psychodynamic therapies (18). Today, these disorders are viewed as a heterogeneous but interrelated group of conditions, arising from complex interactions between genetic vulnerability, early life experiences, and subsequent stressors (19). The "final common pathway" for these disorders probably involves mislabeling of environmental stimuli as threats (alarms), misinterpretation of these threats and their consequences (beliefs), and subsequent maladaptive behaviors (coping). The neuroanatomical substrates of these events include the prefrontal cortices, amygdala, hippocampus, insula, and basal ganglia (16, 20).

The current Research Topic presents seven papers examining the epidemiology, diagnosis, and management of anxiety and stress-related disorders. These reports reflect the diversity of this group of conditions, the clinical challenges faced in treating them, and their relationship to both individual and collective forms of stress and trauma.

Hallucinations are typically associated with psychotic disorders, but they can be a manifestation of significant anxiety or stress (21, 22). Jiang et al. report the case of an elderly woman with olfactory hallucinations associated with generalized anxiety disorder. No evidence of a psychotic, neurological or nasopharyngeal disorder was found. The patient's hallucinations resolved with pharmacological management of the anxiety disorder.

Culture can significantly shape the presentation of anxiety disorders by modifying interpretations of specific situations, the health-related consequences attached to them, and the specific illnesses they may represent. This can lead to unusual symptom presentations, particularly in Asian and African cultures (23, 24). Religious and spiritual beliefs and practices can also influence anxiety symptomatology (25). Khoe and Gudi describe the case of a patient with recurrent trance and possession episodes, associated with auditory and visual hallucinations. These symptoms were found to be panic attack equivalents, shaped by religious beliefs and guilt related to a past life event. As in the first case, the patient responded to pharmacological treatment for panic disorders. These cases highlight the challenges involved in diagnosing anxiety disorders across cultures.

Collective forms of trauma are associated with significant increases in anxiety and stress-related disorders (26-28). Limone et al. systematically reviewed the literature on anxiety and stress in students in relation to the COVID-19 pandemic and the Russia-Ukraine war. They found high levels of both anxiety symptoms (14-89%) and stress (28-56%) in this population. Risk factors associated with these symptoms included female gender, course of study, social isolation, prior physical or mental illness, and the impact of these disasters on students' families and friends. Kurapov et al. reviewed studies from the first 6 months of the Russia-Ukraine war, and found high levels of both anxiety (36%) and stress (70%) in the Ukrainian general population. Of those with symptoms of anxiety, over one third had severe symptoms suggestive of an anxiety disorder. Risk factors in this population included female gender, older age, financial difficulties, unemployment, forced relocation, and direct experience of war-related trauma. These findings highlight the need for access to adequate mental health care and psychosocial support in conflict and disaster settings.

Anxiety disorders can also be caused by general medical conditions. These "secondary" anxiety disorders should be suspected in patients with atypical symptom presentations (29, 30). Geng et al. describe a patient with Cushing's disease, initially misdiagnosed as generalized anxiety disorder. Zhai et al. describe a patient whose anxiety symptoms were found to be related to a patent foramen ovale. In the first of these cases, the patient experienced a severe adverse reaction to standard doses of an SRI. These cases highlight the biological links between anxiety and factors such as cortisol levels (31) or central perception of suffocation (32), the importance of identifying and treating

## References

1. Stein DJ, Scott KM, de Jonge P, Kessler RC. Epidemiology from anxiety disorders: from surveys to nosology and back. *Dialogues Clin Neurosci.* (2017) 19:127–35. doi: 10.31887/DCNS.2017.19.2/dstein

the underlying medical disorder, and the need for caution when prescribing psychotropics in medically unstable patients.

Response to standard pharmacological or psychological treatments in anxiety and stress-related disorders is often unsatisfactory (5, 6, 33). Non-invasive neurostimulation methods, such as repetitive transcranial magnetic stimulation (rTMS) and transcranial direct current stimulation (tDCS) have shown preliminary evidence of efficacy in anxiety disorders and PTSD (34). Chang et al. describe the successful use of accelerated theta-burst rTMS, applied over the dorsolateral prefrontal cortices (DLPFC), in a patient with treatment-resistant PTSD occurring in the context of emotional and physical abuse. This result is consistent with the existing literature, and highlights the importance of the DLPFC as a key component of the "executive control" network involved in anxiety and stress responses (35, 36).

Overall, the papers included in this Research Topic provide a snapshot of the biological, social, and cultural dimensions of anxiety and stress-related disorders, and will be of interest to clinicians, researchers and public health experts.

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<sup>2.</sup> Lee M, Aggen SH, Otowa T, Castelao E, Preisig M, Grabe HJ, et al. Assessment and characterization of phenotypic heterogeneity of anxiety disorders across five large cohorts. *Int J Methods Psychiatr Res.* (2016) 25:255–66. doi: 10.1002/mpr.1519

3. Yang X, Fang Y, Chen H, Zhang T, Yin X, Man J, et al. Global, regional and national burden of anxiety disorders from 1990 to 2019: results from the Global Burden of Disease Study 2019. *Epidemiol Psychiatr Sci.* (2021) 30:e36. doi: 10.1017/S2045796021000275

4. Xiong P, Liu M, Liu B, Hall BJ. Trends in the incidence and DALYs of anxiety disorders at the global, regional, and national levels: estimates from the Global Burden of Disease Study 2019. *J Affect Disord*. (2022) 297:83–93. doi: 10.1016/j.jad.2021.10.022

5. Ansell EB, Pinto A, Edelen MO, Markowitz JC, Sanislow CA, Yen S, et al. The association of personality disorders with the prospective 7-year course of anxiety disorders. *Psychol Med.* (2011) 41:1019–28. doi: 10.1017/S0033291710001777

6. Solis EC, van Hemert AM, Carlier IVE, Wardenaar KJ, Schoevers RA, et al. The 9year clinical course of depressive and anxiety disorders: New NESDA findings. *J Affect Disord*. (2021) 295:1269–79. doi: 10.1016/j.jad.2021.08.108

7. Moreno-Peral P, Conejo-Ceron S, Motrico E, Rodriguez-Morejon A, Fernandez A, Garcia-Campayo J, et al. Risk factors for the onset of panic and generalised anxiety disorders in the general adult population: a systematic review of cohort studies. *J Affect Disord.* (2014) 168:337–48. doi: 10.1016/j.jad.2014.06.021

8. van Honk J, Bos PA, Terbrug D, Heany S, Stein DJ. Neuroendocrine models of social anxiety disorder. *Dialogues Clin Neurosci.* (2015) 17:287–93. doi: 10.31887/DCNS.2015.17.3/jhonk

9. Knoll AT, Carlezon WA Jr. Dynorphin, stress, and depression. *Brain Res.* (2010) 1314:56–73. doi: 10.1016/j.brainres.2009.09.074

10. Strohle A, Holsboer F. Stress responsive neurohormones in depression and anxiety. *Pharmacopsychiatry*. (2003) 36:S207–14. doi: 10.1055/s-2003-4513

11. Schiele MA, Gottschalk MG, Domschke K. The applied implications of epigenetics in anxiety, affective and stress-related disorders - a review and synthesis on psychosocial stress, psychotherapy and prevention. *Clin Psychol Rev.* (2020) 77:101830. doi: 10.1016/j.cpr.2020.101830

12. Longo MSC, Vilete LMP, Figueira I, Quintana MI, Mello MF, Bressan RA, et al. Comorbidity in post-traumatic stress disorder: a population-based study from the two largest cities in Brazil. J Affect Disord. (2020) 263:715–21. doi: 10.1016/j.jad.2019.11.051

13. Smoller JW. The genetics of stress-related disorders: PTSD, depression, and anxiety disorders. *Neuropsychopharmacology.* (2016) 41:297–319. doi: 10.1038/npp.2015.266

14. Auxemery Y. Post-traumatic psychiatric disorders: PTSD is not the only diagnosis. *Presse Med.* (2018) 47:423–30. doi: 10.1016/j.lpm.2017.12.006

15. Freyberger HJ, Stieglitz RD, Berner P. Neurotic, stress-related and somatoform disorders (section F4) and physiological dysfunction associated with mental or behavioural factors (section F5): results of the ICD-10 field trial. *Pharmacopsychiatry.* (1990) 23:165–9. doi: 10.1055/s-2007-1014558

16. Serra-Blasco M, Radua J, Soriano-Mas C, Gomez-Benlloch A, Porta-Casteras D, Carulla-Roig M, et al. Structural brain correlates in major depression, anxiety disorders and post-traumatic stress disorder: a voxel-based morphometry meta-analysis. *Neurosci Biobehav Rev.* (2021) 129:269–81. doi: 10.1016/j.neubiorev.2021.07.002

17. Barbano AC, van der Mei WF, deRoon-Cassini TA, Grauer E, Lowe SR, Matsuoka YJ, et al. Differentiating PTSD from anxiety and depression: lessons from the ICD-11 PTSD diagnostic criteria. *Depress Anxiety*. (2019) 36:490–8. doi: 10.1002/da.22881

18. Zerbe KJ. Through the storm: psychoanalytic theory in the psychotherapy of the anxiety disorders. *Bull Menninger Clin.* (1990) 54:171–83.

19. Sharma S, Powers A, Bradley B, Ressler KJ. Gene x environment determinants of stress- and anxiety-related disorders. *Annu Rev Psychol.* (2016) 67:239–61. doi: 10.1146/annurev-psych-122414-033408

20. Bystritsky A, Spivak NM, Dang BH, Becerra SA, Distler MG, Jordan SE, et al. Brain circuitry underlying the ABC model of anxiety. *J Psychiatr Res.* (2021) 138:3–14. doi: 10.1016/j.jpsychires.2021.03.030

21. Ohayon MM. Prevalence of hallucinations and their pathological associations in the general population. *Psychiatry Res.* (2000) 97:153–64. doi: 10.1016/s0165-1781(00)00227-4

22. Mertin P, Hartwig S. Auditory hallucinations in nonpsychotic children: diagnostic considerations. *Child Adolesc Ment Health.* (2004) 9:9–14. doi: 10.1046/j.1475-357X.2003.00070.x

23. Hofmann SG, Hinton DE. Cross-cultural aspects of anxiety disorders. Curr Psychiatry Rep. (2014) 16:450. doi: 10.1007/s11920-014-0450-3

24. Hinton DE, Park L, Hsia C, Hofmann S, Pollack MH. Anxiety disorder presentations in Asian populations: a review. CNS Neurosci Ther. (2009) 15:295–303. doi: 10.1111/j.1755-5949.2009.00095.x

25. Agorastos A, Demiralay C, Huber CG. Influence of religious aspects and personal beliefs on psychological behavior: focus on anxiety disorders. *Psychol Res Behav Manag.* (2014) 7:93–101. doi: 10.2147/PRBM.S43666

26. Jain N, Prasad S, Czarth ZC, Chodnekar SY, Mohan S, Savchenko E, et al. War psychiatry: identifying and managing the neuropsychiatric consequences of armed conflicts. *J Prim Care Community Health.* (2022) 13:21501319221106625. doi: 10.1177/21501319221106625

27. Delpino FM, da Silva CN, Jeronimo JS, Mulling ES, da Cunha LL, Weymar MK, et al. Prevalence of anxiety during the COVID-19 pandemic: a systematic review and meta-analysis of over 2 million people. *J Affect Disord.* (2022) 318:272–82. doi: 10.1016/j.jad.2022.09.003

28. Beaglehole B, Mulder RT, Frampton CM, Boden JM, Newton-Howes G, Bell CJ. Psychological distress and psychiatric disorder after natural disasters: systematic review and meta-analysis. *Br J Psychiatry*. (2018) 213:716–22. doi: 10.1192/bjp.2018.210

29. Cassem EH. Depression and anxiety secondary to medical illness. *Psychiatr Clin N Am.* (1990) 13:597–612. doi: 10.1016/S0193-953X(18)30338-1

30. Testa A, Giannuzzi R, Daini S, Bernardini L, Petrongolo L, Silveri NG. Psychiatric emergencies (part III): psychiatric symptoms resulting from organic diseases. *Eur Rev Med Pharmacol Sci.* (2013) 17:86–99. Available online at: https://www.europeanreview. org/article/3087

31. Zorn JV, Schur RR, Boks MP, Kahn RS, Joels M, Vinkers CH. Cortisol stress reactivity across psychiatric disorders: a systematic review and meta-analysis. *Psychoneuroendocrinology*. (2017) 77:25–36. doi: 10.1016/j.psyneuen.2016.11.036

32. Tural U, Iosifescu DV. A systematic review and network meta-analysis of carbon dioxide provocation in psychiatric disorders. *J Psychiatr Res.* (2021) 143:508–15. doi: 10.1016/j.jpsychires.2020.11.032

33. Williams T, Phillips NJ, Stein DJ, Ipser JC. Pharmacotherapy for post traumatic stress disorder (PTSD). *Cochrane Database Syst Rev.* (2022) 3:CD002795. doi: 10.1002/14651858.cd002795.pub3

34. Lee HJ, Stein MB. Update on treatments for anxiety-related disorders. *Curr Opin Psychiatry.* (2023) 36:140–5. doi: 10.1097/YCO.000000000 000841

35. Edinoff AN, Hegefeld TL, Petersen M, Patterson JC, Yossi C, Slizewski J, et al. Transcranial magnetic stimulation for post-traumatic stress disorder. *Front Psychiatry*. (2022) 13:701348. doi: 10.3389/fpsyt.2022.701348

36. White LK, Makhoul W, Teferi M, Sheline YI, Balderston NL. The role of dlPFC laterality in the expression and regulation of anxiety. *Neuropharmacology.* (2023) 224:109355. doi: 10.1016/j.neuropharm.2022.109355