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## **Infections and Emotional Responses**

Alina MAPN da Silva, <sup>1</sup> Luiz Carlos Miller Paiva Nogueira da Silva, <sup>1</sup> Anita L R Saldanha, <sup>1</sup> Ana Paula Pantoja Margeotto, <sup>1</sup> André Luis Valera Gasparoto, <sup>2</sup> and Tania Leme da Rocha Martinez<sup>1,\*</sup>

- 1. Nephrology Department, BP A Beneficência Portuguesa de São Paulo, São Paulo, Brazil
- 2. Intensive Care Unit, BP A Beneficência Portuguesa de São Paulo, São Paulo, Brazil

\*Correspondence: Tania Leme da Rocha Martinez

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#### **Abstract**

Infections can be the cause of emotional disturbances. The psychological aspects related to them have been studied for decades with more emphasis in tuberculosis. On the most recent time scale COVID has been the virus most profoundly and published studies on psychological disorder manifestations. Indications for psychotherapy have multiplied during the COVID pandemic. Also noteworthy are all other infections, such as mPOX, brucellosis, verminosis, influenza etc. Many individuals in contact with virus carriers, such as flu, do not acquire the virus infection while others, per example tuberculosis patients do not become infected by Koch's bacillus. In some patients the symptoms of the infection may disappear in a matter of days, while others even present longer clinical evolution as a neurotropism allergy to brucellosis. This suggests an interrelationship between infections and immune defenses, mainly due to psychic factors influencing the hypothalamic-adrenal axis and producing the relative insufficiency of the adrenal cortex. The attitude of affection and interest adopted by the physician and auxiliary person is of paramount importance, as a form of labor therapy, in an adequate work and subsequent feeling of usefulness. In conclusion, the proper health education related to the illness and treatment plan should be provided to the patients along with their family members. There is a need to improve access to counseling and mental healthcare among patients living with infections.

**Keywords:** Covid; Emotions; Infections; Psychology; Tuberculosis

### Introduction

As from the last decades the emotional impact of infections have been studied more systematically. The citations of tuberculosis, per example, were and still are of the most impact (1-6). On the most recent

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logical disorders.

Indications for psychotherapy have multiplied during the pandemic (7-8). Also noteworthy are all It is a well-known fact that tuberculosis is other infections, such as mPOX, brucellosis, "preferred" by certain families and that certain peoworms, influenza.

Many individuals in contact with carriers of viruses tients themselves, there are frequent comments on (influenza, for example) do not acquire the virus, erroneous observations made by laymen about the others in contact with tuberculosis patients do not prognosis of this or that phthisis: "So-and-so become infected by Koch's bacillus (9-18).

has been described (19).

neurotropism allergic to brucella, which suggests passively receive affection. an interrelationship between infections and immune gland there is a hormone, etiocholanone, which important, in a multiform etiology. It is observed produces hyperthermia, which can explain certain that behind each constitution there is another factor, mias", the first resulting from a need to be sick to conscious is what governs the individual's immune receive attention, affection and affection due to a state; their conflicts can weaken defenses against disgust of sensual origin.

time scale it has been the most profound and wide- Saul (20) in 15 individuals who frequently suffered ly studied COVID in the manifestations of psycho- from pulmonary flu with catarrhal bronchitis, psychoanalytic therapy made susceptibility to colds disappear.

ple (long-terminian-asthenic biotype) are more prone to this disease. Among the tuberculosis pa-(strong type, obese individual) will be cured soon"; a few months later he died. Another said: "so-and-Another infectious state in which the emotional so won't take long" (asthenic, long-haired and thin component enters in large part is in chronic brucel- type); he was wrong too. Dunbar (21) tried to delosis, whose diagnosis by the positivity of blood scribe the "psychological profile of the tuberculosis agglutination is easy, but the persistence of its in- patient". An individual in good previous health, fection, with symptoms very similar to asthenic with few surgical interventions but with a marked neurosis, exacerbated in periods of emotional or tendency to suffer accidents (which shows a desire physical "stress", shows that the body's defenses to self-destruct) being in general, effective, indeare weakened, not by food deficiency, exhaustion, pendent, with good adaptation in the sexual sphere but rather by unconscious psychological factors, as (it is a popular saying that so-and-so became tubercular due to masturbatory or sexual excess, which is not true, because any excess is a self-destructive In some individuals, the symptoms of the infection manifestation of neurosis) in the social and ecodisappear within 60 days, and others even present a nomic sphere, but with an intense childish need to

defenses, mainly due to psychic factors influencing Psychiatric studies have failed to establish personthe hypothalamic-adrenal axis and producing the ality uniformity in tuberculosis patients, but have relative insufficiency of the adrenal cortex. In this shown that in this disease psychological factors are "incomprehensible fevers" or "hysterical hyperther- difficult to observe: unconscious feelings. This uninfections; it is not yet known by what mechanism. Feelings of guilt from childhood, especially from destruction through tuberculosis.

the mold period, are the main responsible for self- tration of Mycobacterium tuberculosis.

Nowadays tuberculosis is included in the group of chic factors on the tuberculosis process; among diseases in which psychosomatic factors may play other signs, he verified the fluctuations of the opa role in the individual's predisposition to infec- sonic index, finding that emotional conflicts detions, probably through the immune system and the creased the value of the index and delayed the reautonomic nervous system. Rich (22) emphasized turn to normality; these changes predisposed the the role of psyche and endocrine disorders in the body to tuberculosis. Eyre (25) stated that the enerproduction and maintenance of resistance to the gy consumed by the state of emotional tension is clinical manifestation of tuberculosis. Kuntz (23) lost and ceases to assist other physiological funcrelated the vegetative nervous system to tuberculo- tions; tuberculosis is especially suitable for causing sis for the following reasons: 1) lesions in the gan-emotional conflicts because of its long duration of glia of the vegetative nervous system are found in the disorder of social and economic relationships autopsies of tuberculosis patients; 2) the autonomic and the isolation it entails. Hartz (26) became tunervous system influences the local character and berculosis and could, as a patient, observe, in the progress of the infection; 3) the section of the va-sanatorium, that the tuberculosis patients kept a gus slows down the unilateral tuberculous process second history different from that referred to in the in rabbits; 4) greater irritability of the tissues (in entrance, that is, there were strong emotional reacwhich the vegetative system plays a predominant tions in the face of certain vital situations that could role), greater exudative tendency; 5) alteration in have prevented a healthy life; he ended up thinking the reactions of blood vessels to toxins and tissue that tuberculosis was another personality disease. acidity also depend on the vegetative nervous sys- Berle (27) in his review says: the clinical evolution tem.

drome. On the other hand, the formation of anti- mation of an obsessive compulsive character. bodies would be related to vegetative tone, being

Ishigami (24) in 1919 studied the influence of psyruns in parallel with the emotional life of the patient and this is one of the factors that influences In Selye's alarm reaction, the stimulation of adrenothe growth of the bacillus in the organism and no corticotropic hormone (ACTH) and these on corti- progress will be made if stressful emotions are not costeroids would act on the lymphoid tissue by in-related to the study of the problem of its virulence, creasing globulins, vectors of antibodies to the which has been recognized since the third century. body's immune defense against infections. In the Merril (28) also thought that these patients had an exhaustion of the body, this mechanism would be intense infantile attachment in the sense of being sacrificed, as in acute adrenal insufficiency syn- passive receptors of affection and the reactive for-

greater when the parasympathetic tone predomi- Hayes (29) understood that tuberculosis was one of nates in the production by the body of agglutinins, the easiest curable chronic diseases, but its prognolysines and precipitins, and, in sympathetic excita- sis became uncertain because of many factors, intion, alexins and opsonins would appear, along cluding the psychology of the patient. Muhl (30) with greater capillary permeability and easier pene- had long described the "suicidal attitudes" of cerverted homicidal desires, and the patient uncon- harmful consequences), psychoanalysis should be sciously identified himself with the one he wished advised, especially if there are cases in the family, to destroy, due to sentimental factors of guilt.

The psychosomatic conception of tuberculosis nei- sis. ther denies nor underestimates the influence of the germ - for without germ there is no tuberculosis - Immunocompetence but considers the problem in all its aspects, from (psychoneuroimmunology) the cellular to the social plane.

progress of the treatment. Thomas Mann in the when they do, they speak different languages". voice of Hans Castorp said: "These pneumatic mothorax, favorably).

The attitude of affection and interest adopted by the that is, that it is not a total incapacitated.

sults in the treatment of tuberculosis are obtained rats. when the patient is considered to have not only diseased organs, but also a diseased personality".

primary cause, which is the unconscious conflict of (35). self-destruction. Therefore, if an individual begins

tain tuberculosis patients, believing them to be con-self-aggression because he knows that it will have because we know that it is a genetic predisposition to acquire infection by Mycrobacterium tuberculo-

> psychosomatics and

The editorial article in the Lancet (31) says: The mental state has a great influence on the good "psychiatrists and immunologists hardly meet, and

friends of yours seem to be in a very good mood Experimental stress in animals produces a decrease too" (referring to a group that submitted to pneu- in immune defenses, low levels of interferon, T lymphocytes and opsonins, which favors infections (32,33).

physician and auxiliary person is of paramount im- Experimental polyarthritis can be produced by inportance, as is essential to labor therapy, but an ad- jecting "mycobacterium butyrucon", but if the coloequate work and subsequent feeling of usefulness, ny is overflowing with rats, there will be an increase in the arthritic lesion (34).

Finally, as Portland, Young and Williams said in Three groups of rats: 1. those that live without disthe name of the "National Association for The Pre-turbances; 2. those exposed to a cat chase for 10 ventions of Tuberculosis" of England: "the best reminutes, 4 times a day; 3. and finally, very handled

In the rats without disorders, 40% developed arthritis, 13% in the rats exposed to the cats and 10% in All these facts lead us to advocate individual or the ones that were handled. It is concluded that group psychoanalysis as the most powerful thera- chronic stress increases the incidence of arthritis peutic weapon in the prophylaxis of tuberculosis and that acute stress, in a sense, protects you from and chronic infections, as it directly combats the arthritis. Gastrectomy impairs immune defenses

to overwork in entertainment, without eating Monkeys separated from their mother during the enough (often this hyponutrition is a symptom of first five years of life, once adults, decrease immune responses (36).

as it confirms that the periods of mold (0 to 7 years methasone, fenfluramine (prolactin does not rise of age), periods in which unconscious fantasies de- after the application of fenfluramine), growth horvelop. Traumas in this period are indeed cofactors mone (increases after ingestion of imipramine). In in the production of most psychosomatic and men- asthenia in psychotic depression, there is an elevatal illnesses, including the etiopathogenesis of cer- tion of dehydroisoandrosterone and loss of phostain types of cancer.

Mertz et al. (37), after having used oral acyclovir Reduced monocytes and macrophages (mature found that the duration of the lesions was shortened ital herpes.

as chlorpromazine and benzodiazepinema in the bands (46). face of the antibody response to the typhoid antigen H(39).

sackie B2 virus, Epstem-Bart virus, killer cells erative period. (NK), plasma IgA (40), decrease in T lymphocytes by phytohemoglutinin and cortisol.

frequent in children with family conflicts (40,41).

The distinction between postviral fatigue and neurocirculatory asthenia is not only by the dosage of In conclusion, the proper health education related the IgM antibody, but also by the measurements of to the illness and treatment plan should be provided

blood lactic acid, before and after exercise (in asthenia the lacticidemia increases), triiodothyronine This fact is extremely important in psychosomatics, (T3), tetraiodothyronine (T4) and the tests of dexaphates in the urine (42,43).

for one year in 227 patients with herpes simplex, monocytes) predict susceptibility to infections (44).

in relation to controls. Hoon et al. (38) in 153 uni- The unemployed and divorced individual has deversity students (67% women) found that a high creased immune defenses (decreased lymphocyte level of positive thymosin-alpha-1 was related to reactivity to phytohemoglutinin PHA and the tubervulnerability to diseases in general, not only to gen- culin-derived protein PPD), anxiety, depression, decreased mental concentration, tobacco and alcohol abuse, and suicidal ideation (45), especially in Certain drugs can modify immune responses, such women who had a good attachment with their hus-

Preoperative stress tests that were altered, i.e., with high response to cold pressor and low lymphocyte The most sensitive tests to verify immunocompe- response to phytoagglutinin and concanavalin, the tence are: infection by herpes simplex virus, Cox- patients had infectious complications in the postop-

Psychoimmunology has demonstrated the influence of stress on lymphocytes (which have receptors for Post-viral fatigue or dysphoric fatigue syndrome or the neurotransmitters T and K - killer cell) causing epidemic of neuromyasthenia or chronic fatigue depressive states. They found once again that the syndrome or neurocirculatory asthenia is measured anxieties in the face of the Acquired Immunodefiby immunoglobulin M (IgM) antibody, and is more ciency Syndrome (AIDS) are overwhelming due to the feeling of abandonment, especially in homosexuals with overprotective.

to the patients along with their family members. There is a need to improve access to counseling and mental healthcare among patients living with 6. infections.

## **Acknowledgments**

In memoriam: Luiz Miller de Paiva.

### **Conflict of interest**

None.

### References

- 1. Nasir A, Hassan II, Ma'ruf A, et al. Coping efforts made: psychological burden of people liv- 8. ing with tuberculosis due to social stigma in society. A qualitative phenomenology study. PLoS One. 2024; 19(7):e0303331. 10.1371/journal.pone.0303331
- 2. Numpong S, Kengganpanich M, Kaewkungwal J, et al. Confronting and coping with multidrug- 9. resistant tuberculosis: life experiences in Thailand. Qual Health Res. 2022; 32(1):159-167. doi: 10.1177/10497323211049777
- 3. Tornu E, Dzansi G, Wilson DM, et al. Mothers' perspectives of physical and psychological issues associated with caring for Ghanaian chil- 10. Walle KM, Askeland RB, Gustavson K, et al. dren living with tuberculosis: a qualitative study. Nurs Open. 2023; 10(4):2439-2448. doi: 10.1002/nop2.1499
- 4. Lu Y, Wang H, Zhu J, et al. Effect of targeted nursing intervention plus psychological coun- 11. Persell SD, Friedberg MW, Meeker D, et al. seling on quality of life, negative emotions, and complications in patients with extensively drug -resistant tuberculosis. Am J Transl Res. 2021; 13(12):13950-13958. https:// pubmed.ncbi.nlm.nih.gov/35035736/
- 5. Morse RM, Myburgh H, Reubi D, et al. Opportunities for mobile app-based adherence support for children with tuberculosis in South Af-

- rica. **JMIR** Mhealth Uhealth. 2020; (11):e19154. doi: 10.2196/19154
- Monistrol-Mula A, Felez-Nobrega M, Oh H, et al. Association between tuberculosis and psychotic experiences: mediating factors and implications for patient care in low- and middleincome countries. J Glob Health. 2024; 14:04005. doi: 10.7189/jogh.14.04005
- 7. de Almondes KM, Bizarro L, Miyazaki MCOS, et al. Comparative analysis of psychology responding to COVID-19 pandemic in Brics nations. Front Psychol. 2021; 12:567585. doi: 10.3389/fpsyg.2021.567585
  - Nel KA, Govender S. Existential Positive Psychology (EPP): a positive tool for healing existential anxieties in South Africa during, and after, the COVID-19 pandemic. Int J Environ Res Public Health. 2022; 19(16):10248. doi: 10.3390/ijerph191610248
  - Ulset VS, Czajkowski NO, Kraft B, et al. Are unpopular children more likely to get sick? Longitudinal links between popularity and infectious diseases in early childhood. PLoS One. 2019; 14(9):e0222222. doi: 10.1371/ journal.pone.0222222
- Risk of attention-deficit hyperactivity disorder in offspring of mothers with infections during pregnancy. JCPP Adv. 2022; 2(2):e12070. doi: 10.1002/jcv2.12070
- Use of behavioral economics and social psychology to improve treatment of acute respiratory infections (BEARI): rationale and design of a cluster randomized controlled trial [1RC4AG039115-01]--study protocol and baseline practice and provider characteristics. Infect Dis. BMC 2013; 13:290. doi: 10.1186/1471-2334-13-290

- 12. Suleri A, Rommel AS, Neumann A, et al. Exposure to prenatal infection and the development of internalizing and externalizing problems in children: a longitudinal population-based study. 19. Harris HJ, Kemple C. Chronic brucellosis and J Child Psychol Psychiatry. 2024; 65(7):874-886. doi: 10.1111/jcpp.13923
- 13. Harris L, Griem J, Gummery A, et al. Neurocephalitis: A multi-centre case-control study. 2020; PLoS One. 15(3):e0230436. doi: 10.1371/journal.pone.0230436
- 14. Rzeszutek M, Oniszczenko W, Firlag-Burkacka E. Social support, stress coping strategies, resil- 21. Dunbar ience and posttraumatic growth in a Polish sample of HIV-infected individuals: results of a 1 year longitudinal study. J Behav Med. 2017; 40 (6):942-954. doi: 10.1007/s10865-017-9861-z
- 15. Sipilä PN, Heikkilä N, Lindbohm JV, et al. Hospital-treated infectious diseases and the risk 22. Rich AR. The pathogenesis of tuberculosis. of dementia: a large, multicohort, observational study with a replication cohort. Lancet Infect Dis. 2021; 21(11):1557-1567. doi: 10.1016/ S1473-3099(21)00144-4
- 16. Lampard-Scotford AR, McCauley A, Kuebel JA, et al. Impact of parasitic infection on mental health and illness in humans in Africa: a systematic review. Parasitology. 2022; 149(8):1003 -1018. doi: 10.1017/S0031182022000166
- 17. Samsudin NA, Karim N, Othman H, et al. Exploring community behaviours and stakeholder 24. Ishigami T. The influence of psychic acts on the challenges in engaging communities with dengue prevention behaviour in Malaysia: implementation research for a qualitative study with a community-based participatory research design. BMJ Open. 2024; 14(3):e074222. doi: 25. Eyre MB. The rôle of emotion in tuberculosis. 10.1136/bmjopen-2023-074222
- 18. Doan DA, Nguyen TTX, Le GB, et al. macy university students regarding monkeypox:

- a multicenter, cross-sectional study in Vietnam. BMC Med Educ. 2024; 24(1):807. 10.1186/s12909-024-05805-4
- psychoneurosis. Psychosom Med. 1954; 16 (5):414-425.doi: 10.1097/00006842-195409000-00007
- psychological and psychiatric outcomes in en- 20. Saul LJ. Psychogenic factors in the etiology of the common cold and related symptoms. Int J Psychoanal. 1938; 19:451-470. https://pepweb.org/browse/IJP/volumes/19? preview=IJP.019.0451A
  - F. Diagnóstico tratamiento psicosomáticos. 1st ed. Barcelona: Ed. José Janés 1950; 279-386. https://www.google.com.br/ books/edition/Diagn%C3% B3stico y tratamiento psicosom%C3%A1tic/ urXBMwEACAAJ?hl=pt-BR
  - Springfield, Baltimore: Charles C Thomas Ed https://www.science.org/doi/10.1126/ 1944. science.100.2601.407
  - 23. Kuntz A. The autonomic nervous system. Philadelphia: Lea & Febiger 1945. https:// scholar.google.com/scholar lookup? &title=The%20Autonomic%20Nervous% 20System&publication year=1945&author=Kuntz% 2CAlbert
    - progress of pulmonary tuberculosis. Am Rev 1919; 2(8):470-484. Tuberc. https:// www.atsjournals.org/doi/abs/10.1164/ art.1918.2.8.470?journalCode=art
    - Tubercle. 1933; 14(7):300-311. https:// doi.org/10.1016/S0041-3879(33)80065-1
- Knowledge and attitudes of medical and phar- 26. Hartz J. Tuberculosis and personality conflicts. Psychosom Med. 1944; 6(1):17-22. https://

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- journals.lww.com/psychosomaticmedicine/ Citation/1944/01000/
- Tuberculosis and Personality Conflicts.3.aspx
- 27. Berle BB. Emotional factors and tuberculosis; a critical review of the literature. Psychosom Med. 1948; 10(6):366-373. doi: 10.1097/00006842-194811000-00010
- 28. Merrill BR. Some psychosomatic aspects of pulmonary tuberculosis; a review of the English language literature. J Nerv Ment Dis. 1953; 10.1097/00005053-117(1):9-28. doi: 195301000-00002
- 29. Hayes EW. The prognosis in tuberculosis with especial reference to the psychological aspects. Med. 1931; 4:1183-1187. Ann Intern doi:10.7326/0003-4819-4-9-1183
- 30. Muhl AM. Tuberculosis from the psychiatric 16(4):397-403. https://pep-web.org/browse/ document/psar.016.0397a
- 31. Emotion and immunity. Lancet. 1985; 2 6736(85)90232-6
- 32. Hamilton DR. Immunosuppressive effects of predator induced stress in mice with acquired immunity to Hymenolepis nana. J Psychosom 3999(74)90014-2
- 33. Palmblad J, Cantell K, Strander H, et al. Stressor exposure and immunological response in man: interferon-producing capacity and phagocytosis. J Psychosom Res. 1976; 20(3):193-199. doi: 10.1016/0022-3999(76)90020-9
- 34. Amkraut AA, Solomon GF, Kraemer HC. Stress, early experience and adjuvant-induced arthritis in the rat. Psychosom Med. 1971; 33 doi: (3):203-214.197105000-00002
- 35. Gryglewski A, Marcinkiewicz J, Popiela T, et

- al. Effect of surgical trauma (gastrectomy) on cell-mediated and humoral responses in mice. Clin Exp Immunol. 1985; 59(1):50-54. https:// www.ncbi.nlm.nih.gov/pmc/articles/ PMC1577173/pdf/clinexpimmunol00136-0059.pdf
- 36. Laudenslager M, Capitanio JP, Reite M. Possible effects of early separation experiences on subsequent immune function in adult macaque monkeys. Am J Psychiatry. 1985; 142(7):862-864. doi: 10.1176/ajp.142.7.862
- 37. Mertz GJ, Jones CC, Mills J, et al. Long-term acyclovir suppression of frequently recurring genital herpes simplex virus infection. A multicenter double-blind trial. JAMA. 1988; 260 doi:10.1001/ (2):201-206. jama.1988.03410020067030
- approach. A follow-up. Psychoanal Rev. 1929; 38. Hoon EF, Hoon PW, Rand KH, et al. A psychobehavioral model of genital herpes recurrence. J Psychosom Res. 1991; 35(1):25-36. doi: 10.1016/0022-3999(91)90004-8
- (8447):133-134. https://doi.org/10.1016/S0140- 39. Dadhich AP, Sharma VN, Godhwani JL. Effect of restraint stress on immune response & its modification by chlorpromazine, Diazepam & pentobarbitone. Indian J Exp Biol. 1980; 18 (7):756-757. PMID: 7191830
- Res. 1974; 18(3):143-153. doi: 10.1016/0022- 40. Wilson PM, Kusumakar V, McCartney RA, et al. Features of Coxsackie B virus (CBV) infection in children with prolonged physical and psychological morbidity. J Psychosom Res. 1989; 33(1):29-36. doi: 10.1016/0022-3999(89) 90103-7
  - 41. Taylor GR, Neale LS, Dardano JR. Immunological analyses of U.S. Space Shuttle crewmembers. Aviat Space Environ Med. 1986; 57 (3):213-217. PMID: 3485967
  - 10.1097/00006842- 42. Miller de Paiva L. Técnica de psicanálise: bricolage e filigranas [Psychoanalysis technic: bricolage and filigrees]. Rio de Janeiro, Brazil:

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- Ed. Imago 1987; 231-260.
- 43. Miller de Paiva L. Psychosomatic psychiatry. São Paulo, Brazil: Garatuja Press 1990.
- 44. Moldofsky H, Tullis C, Lue FA, et al. Sleeprelated myoclonus in rheumatic pain modula- 46. Kiecolt-Glaser JK, Fisher LD, Ogrocki P, et al. tion disorder (fibrositis syndrome) and in excessive daytime somnolence. Psychosom Med. 1984; 46(2):145-151. doi: 10.1097/00006842-198403000-00006
- 45. Arnetz BB, Wasserman J, Petrini B, et al. Immune function in unemployed women. Psycho-Med. 1987; 49(1):3-12. doi: som 10.1097/00006842-198701000-00001
  - Marital quality, marital disruption, and immune function. Psychosom Med. 1987; 49 (1):13-34. doi: 10.1097/00006842-198701000-00002

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