Re: Query publication [201220-002101] [210218-006595] [210508-005905]

From: Sci Tech Jrnls CS < JournalCustomerService-usa@elsevier.com>

To: helmy.m@protonmail.com <helmy.m@protonmail.com>

Date: Saturday, May 8th, 2021 at 5:19 PM

Dear Mohamed Helmy,

Thank you for contacting Elsevier Customer Service.

This is an automated acknowledgement to confirm we have received your query.

Ticket number 210508-005905 has been opened on your behalf and you can expect to receive a response within 48 hours (excluding weekends).

For immediate assistance, please visit the Elsevier Support Center.

Regards,

Elsevier Customer Service

From: Mohamed Helmy Date: 08/05/2021 09.19 AM

Dear Mr. McDowell,

According to *Heliyon* Ethics and Editorial Policies, the evidence I submitted concerning the article by Lee *et al.* (2020) in my email to you on 20 December 2020 (see correspondence with references below) indicates that several policies may have been breached. This includes *Heliyon Reporting Standards*, such as accuracy of the information on data provided by the authors. Indeed, according to the *Elsevier Publishing Ethics Resource Kit for editors*, the evidence presented indicates gross inaccuracies and/or errors in the article, and may amount to fraud. To take one example, the issues concerning statistical manipulation of the data by the authors is in contradiction to ARRIVE guidelines (2.0).

In light of the evidence presented showing irregularities putatively amounting to research misconduct in the article by Lee *et al.* (2020), your statement that "...we have concluded that the points you have raised are not issues...", and which you made without presenting any evidence whatsoever, is simply dismissive and inadequate.

I am also puzzled by your choice of words in thanking me "...for taking the time to bring forward *your* issues to us..." (emphasis added). The issues are mine insofar that I am a member of the academic community. Any *issues* as such belong to *Heliyon* and the authors.

May I suggest that you please address the points 1 to 4 mentioned in my message of 20 December 2020. Since these points are 'not issues', and since the editorial review and discussion with the authors is complete, it should be a straight-forward matter to clarify why the evidence presented does not constitute issues. Failing that, I do not see why I should not publish our correspondence on <u>www.nanyangscandal.com</u>. Confidentiality is assumed during

Publications | helmy.m@protonmail.com | ProtonMail

an investigative process, and was honored. Now that the investigation is complete, what is your objection to publication of non-issues?

Kind regards, Mohamed Helmy MD, PhD

www.nanyangscandal.com helmy.m@protonmail.com +65 83 555 817 10 Jurong Lake Link, #15-39 Singapore, 648131

------ Original Message ------On Tuesday, March 30, 2021 4:00 AM, Schulz, Christian (ELS-AMS) <<u>c.schulz@cell.com</u>> wrote:

Dear Dr. Helmy,

The authors have requested additional time to respond to this. We have allowed them a new deadline for 8.4.21.

With kind regards,

Christian Schulz



F

Dr. Christian Schulz | Cell Press | Lead Editor Heliyon | c.schulz@cell.com

The name Heliyon comes from "Helios," the name for the Greek god of the sun. This name

gave us inspiration as we aspire to illuminate knowledge across a broad spectrum of research.

From: Helmy, M. <<u>helmy.m@protonmail.com</u>>
Sent: Monday, March 29, 2021 06:35
To: Schulz, Christian (ELS-AMS) <<u>c.schulz@cell.com</u>>; JournalCustomerService-usa@elsevier.com>; elvira.brattico@clin.au.dk;
elvira.brattico@uniba.it; janice.pluth@unlv.edu; wim.crusio@u-bordeaux.fr
Subject: RE: : Query publication [201220-002101] [210218-006595]

Dear Heliyon,

Regarding the article published in Heliyon in 2020, titled 'Female rats are resilient to the behavioral effects of maternal separation stress and exhibit stress-induced neurogenesis', by Lee, Koe, Ashokan, and Mitra, details below and correspondence reference [201220-002101] [210218-006595]:

I believe it is unethical to publicly comment on an ongoing-investigation. However, if the investigation is closed or there is no response by the authors during a reasonable period of time and no expression of concern was made on the article, then it is my duty to the academic community to raise the issue in the public domain. Please let me know the status of the current investigation.

In this particular incident, relevant to this article, any biologist with some knowledge of neurophysiology may understand the irregularities and invalidity of the article quite comprehensibly. Please see <u>www.nanyangscandal.com</u> for particulars of misconduct by Mitra and others.

I look forward to your reply.

Kind regards,

Mohamed Helmy

MD, PhD

www.nanyangscandal.com

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

----- Original Message ------

On Thursday, February 18, 2021 4:33 PM, Schulz, Christian (ELS-AMS) <<u>c.schulz@cell.com</u>> wrote:

Dear Dr. helmy,

Thank you for sharing your concerns. Following your message from December to the editors of the *Heliyon Neuroscience* section we are already investigating this case and will get back to you.

With kind regards,

Christian Schulz



P

Dr. Christian Schulz | Cell Press | Lead Editor Heliyon | c.schulz@cell.com

The name Heliyon comes from "Helios," the name for the Greek god of the sun. This

name gave us inspiration as we aspire to illuminate knowledge across a broad spectrum of research.

Notice of confidentiality:

This email and any information and files transmitted with it are confidential and intended solely for the private use of the individual or entity to whom they are addressed. Public posting or distribution of all or part of this communication without the express permission of the sender is not permitted. If you are not the intended recipient, any use of this communication is unlawful. If you have received this transmission in error, please immediately notify us by return e-mail and destroy the original message.

From: Heliyon <<u>info@heliyon.com</u>>

Sent: Thursday, February 18, 2021 05:13

To: Schulz, Christian (ELS-AMS) <<u>c.schulz@cell.com</u>>

Subject: Re: : Query publication [201220-002101] [210218-006595]

Dear Christian,

Hope you are well.

Dr Mohamed Helmy contacted regarding the published article in Heliyon entitled "Female rats are resilient to the behavioral effects of maternal separation stress and exhibit stress-induced neurogenesis" (HLY4753).

He mentioned a major finding of the article is a misinterpretation of erratic statistical method and/or reporting. Please refer to his email below.

Kindly look into this and advise him at <u>helmy.m@protonmail.com</u>

Thank you in advance.

Kind regards

Sharon Joy Lintao Researcher Support ELSEVIER

Make all your research work discoverable, comprehensible, citable and reproducible. Find out more
about Elsevier's open access, multidisciplinary Research Elements journals

From: Sharon Joy Lintao Date: Thursday, February 18, 2021 04:02 AM GMT

Dear Dr Helmy,

Thank you for your email that has been forwarded to us regarding the published article in Heliyon entitled "Female rats are resilient to the behavioral effects of maternal separation stress and exhibit stress-induced neurogenesis.".

I wish to inform you that I have contacted the Publisher about your concern and requested to contact you directly via email.

Please feel free to contact me if you need further assistance.

Kind regards

Sharon Joy Lintao Researcher Support ELSEVIER

Make all your research work discoverable, comprehensible, citable and reproducible. Find out more about Elsevier's open access, multidisciplinary <u>Research Elements journals</u>

From: Sharon Joy Lintao Date: Thursday, February 18, 2021 04:02 AM GMT

Dear Rhean,

I beg your pardon. What do you mean by 'we only accept publication for authors'? I am reporting putative research misconduct in a publication from your office. According to your guidelines you are required to investigate and correct the scientific record if necessary.

Please acknowledge and inform me of corrective action to be taken.

Kind regards,

Mohamed Helmy

MD, PhD

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

----- Original Message ------

On Wednesday, February 17, 2021 6:43 AM, Sci Tech Jrnls CS <<u>JournalCustomerService</u><u>usa@elsevier.com</u>> wrote:

Dear Mohamed Helmy,

Thank you for your e-mail regarding publication.

Please be advised that we only accept publication for authors.

You may contact them directly.

If I can be of further assistance, do not hesitate to contact me.

Important:

This e-mail contains personal data which should only be used and retained for legitimate business purposes. It should be deleted in accordance with **Elsevier's GDPR (General Data Protection Regulation)** policy.

Kind Regards,

Rhean Dee Elnar Customer Service Representative ELSEVIER | Operations, Global Customer Service www.elsevier.com

Twitter| Facebook| LinkedIn| Google+

Did you know? You can visit our customer support website and view our Frequently Asked Questions at <u>Elsevier Support Center</u>.

From: Administrator

Date: Sunday, February 14, 2021 08:18 PM GMT

Dear Customer,

Thank you for contacting Elsevier Customer Service.

This is an automated acknowledgement to confirm we have received your query.

Ticket number 210214-009528 has been opened on your behalf and you can expect to receive a response within 48 hours (excluding weekends).

For immediate assistance, please visit our Elsevier Support Center.

Regards,

Elsevier Customer Service

From: Mohamed Helmy

Date: Sunday, February 14, 2021 08:18 PM GMT

Dear Heliyon,

I was wondering if you had time to consider my message below?

Kind regards,

Mohamed Helmy

MD, PhD

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

----- Original Message ------

On Sunday, December 20, 2020 2:54 PM, Sci Tech Jrnls CS <<u>JournalCustomerService</u><u>usa@elsevier.com</u>> wrote:

/span>

This email is for use by the intended recipient and contains information that may be confidential. If you are not the intended recipient, please notify the sender by return email and delete this email from your inbox. Any unauthorized use or distribution of this email, in whole or in part, is strictly prohibited and may be unlawful. Any price quotes contained in this email are merely indicative

and will not result in any legally binding or enforceable obligation. Unless explicitly designated as an intended e-contract, this email does not constitute a contract offer, a contract amendment, or an acceptance of a contract offer.

Elsevier Limited. Registered Office: The Boulevard, Langford Lane, Kidlington, Oxford, OX5 1GB, United Kingdom, Registration No. 1982084, Registered in England and Wales. <u>Privacy Policy</u>

----- Original Message ------

On Monday, February 15, 2021 4:17 AM, Helmy, M. <<u>helmy.m@protonmail.com</u>> wrote:

Dear Heliyon,

I was wondering if you had time to consider my message below?

Kind regards,

Mohamed Helmy

MD, PhD

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

----- Original Message ------

On Sunday, December 20, 2020 2:54 PM, Sci Tech Jrnls CS <<u>JournalCustomerService</u><u>usa@elsevier.com</u>> wrote:

Dear Customer,

Thank you for contacting Elsevier Customer Service.

This is an automated acknowledgement to confirm we have received your query.

Publications | helmy.m@protonmail.com | ProtonMail

Ticket number 201220-002101 has been opened on your behalf and you can expect to receive a response within 48 hours (excluding weekends).

For immediate assistance, please visit our Elsevier Support Center.

Regards,

Elsevier Customer Service

From: Mohamed Helmy

Date: 20/12/2020 06.54 AM

Dear Professor Brattico, Professor Pluth, and Professor Crusio,

Regarding the article published in *Heliyon* [1] in 2020, titled 'Female rats are resilient to the behavioral effects of maternal separation stress and exhibit stress-

induced neurogenesis', by Lee, Koe, Ashokan, and Mitra:

1. A major finding of the article is a misinterpretation of erratic statistical method and/or reporting:

- a. In [1], page 2 it states: "...Inter-group differences for behavioral and dendritic parameters were analyzed using an independent sample t-test. An analysis of variance was used for analyzing data for neurogenesis, with dentate gyrus of ventral/dorsal hippocampus serving as within-subject and experimental treatment serving as a between-subject source of variance...". What is the justification for analyzing neurogenesis data differently to other tests in the same experiment, given that variables are equal?
- b. Subsequently, "...An analysis of variance was conducted with [control or AFR] or [maternally separated or MS] as a between-subject source of variance and dorsal or ventral hippocampal aspect of the dentate gyrus as a within-subject source of variance. Experimental treatment significantly affected dentate gyrus neurogenesis...Neurogenesis was not significantly different between dentate gyrus in dorsal and ventral hippocampus...Similarly, interaction between experimental treatment and brain regions did not reach statistical significance...<u>Thus, maternal separation increased neurogenesis in the dentate gyrus</u>..." emphasis added, [1], page 3. Did maternal separation have a *significant effector* a *main effecton* hippocampal neurogenesis? Whether maternal separation significantly increased neurogenesis in hippocampus or not, this is not represented in Figure 3. with a *p* value or otherwise above the data as was done in Figures 1. and 2. May the statistical method be clarified and significance representation in Figures be corrected to match the text?
- 2. Given the vital roles of amygdala and neurostructural plasticity in fear processing and the stress response, and the understudied nature of gender dimorphism of the same, it is confusing that a large number of **reference citations in the article [1] are incorrect, misleading, or faulty**, for example:
- a. "...chronic stressors during adulthood cause a long-lasting dendritic expansion in principal neurons of the basolateral amygdala (BLA) in males [17]..." [1], page 1. The reference [17] is probably a conference abstract and links to it are faulty, they do not lead to the material. **May the reference to the conference abstract be corrected?** It would be interesting to know why it was necessary to refer to the conference abstract cited in the presence of abundant publications on long-lasting dendritic expansion of basolateral amygdala neurons after various forms of stress in adulthood including chronic stress [3-10].
- b. "...This is in agreement with earlier reports showing a lack of anxiogenesis in female rodents after exposure to chronic stressors in adulthood [11, 12, 21]..." [1], page 3. None of the references [11, 12, 21] support the claim that there is a 'lack' or even attenuated anxiogenesis in female rodents after exposure to stress, may the citations in [1] be verified? To support this statement I recommend [14] and to balance it as well as to address opposing findings in neuroplasticity [15] and perhaps [16,17].

- 3. Regarding the representative drawings and photomicrographs shown in [1]:
- a. May it be clarified from which experimental group photomicrographs in Figure 3.(C) are?
- b. What is the justification for using a drawing tube, or *camera lucida*, to draw neurons by hand, then scan them, then digitally process them when the process could have been done using the same apparatus and software already used in the experiments as reported in [1]?
- 4. About half the text in the Discussion is about a putative estrogenic neuroprotective effect, and estrus is ducussed. However, no efforts were made to assess estrus in the experiment, which may highlight the robust nature of the findings. Why was estrus not assessed in the experimental animals using methods well-established by the corresponding author's co-authors and collaborators [18,19]?

Unfortunately the corresponding author did not respond to my query.

Kind regards,

Mohamed Helmy

MD, PhD

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

1. Lee YJ, Koe AS, Ashokan A, Mitra R. (2020). Female rats are resilient to the behavioral effects of maternal separation stress and exhibit stress-induced neurogenesis. *Heliyon* **6** (8):e04753-e04753. doi:10.1016/j.heliyon.2020.e04753

17. Vyas A, Mitra R, Chattarji S. (2003). Enhanced anxiety and hypertrophy in basolateral amygdala neurons following chronic stress in rats. *Annals of the New York Academy of Sciences* **985** (1):554-555. doi:10.1111/j.1749-6632.2003.tb07127.x

3. Cui H, Sakamoto H, Higashi S, Kawata M. (2008). Effects of singleprolonged stress on neurons and their afferent inputs in the amygdala. *Neuroscience* **152** (3):703-712. doi:<u>10.1016/j.neuroscience.2007.12.028</u>

4. Heinrichs SC, Leite-Morris KA, Guy MD, Goldberg LR, Young AJ, Kaplan GB. (2013). Dendritic structural plasticity in the basolateral amygdala after fear conditioning and its extinction in mice. *Behav. Brain Res.* **248**:80-84. doi:10.1016/j.bbr.2013.03.048

5. Colyn L, Venzala E, Marco S, Perez-Otaño I, Tordera RM. (2019). Chronic social defeat stress induces sustained synaptic structural changes in the prefrontal cortex and amygdala. *Behav. Brain Res.* **373**:112079. doi:10.1016/j.bbr.2019.112079

6. Cohen H, Kozlovsky N, Matar MA, Zohar J, Kaplan Z. (2014). Distinctive hippocampal and amygdalar cytoarchitectural changes underlie specific patterns of behavioral disruption following stress exposure in an animal model of PTSD. *Eur. Neuropsychopharmacol.* **24** (12):1925-1944. doi:<u>10.1016/j.euroneuro.2014.09.009</u>

7. Hill MN, Hillard CJ, Mcewen BS. (2011). Alterations in Corticolimbic Dendritic Morphology and Emotional Behavior in Cannabinoid CB1 Receptor– Deficient Mice Parallel the Effects of Chronic Stress. *Cereb. Cortex* **21** (9):2056-2064. doi:10.1093/cercor/bhq280

8. Qin M, Xia Z, Huang T, Smith CB. (2011). Effects of chronic immobilization stress on anxiety-like behavior and basolateral amygdala morphology in Fmr1 knockout mice. *Neuroscience* **194**:282-290. doi:10.1016/j.neuroscience.2011.06.047

9. Wang Y-C, Ho U-C, Ko M-C, Liao C-C, Lee L-J. (2012). Differential neuronal changes in medial prefrontal cortex, basolateral amygdala and nucleus accumbens after postweaning social isolation. *Brain Structure and Function* **217** (2):337-351.

10. Adamec R, Hebert M, Blundell J, Mervis RF. (2012). Dendritic morphology of amygdala and hippocampal neurons in more and less predator stress responsive rats and more and less spontaneously anxious handled controls. *Behav. Brain Res.* **226** (1):133-146. doi:<u>10.1016/j.bbr.2011.09.009</u>

11. Farinetti A, Aspesi D, Marraudino M, Marzola E, Amianto F, Abbate-Daga G, Gotti S. (2020). Sexually dimorphic behavioral effects of maternal separation in anorexic rats. *Developmental Psychobiology* **62** (3):297-309. doi:10.1002/dev.21909

12. Slotten HA, Kalinichev M, Hagan JJ, Marsden CA, Fone KC. (2006). Longlasting changes in behavioural and neuroendocrine indices in the rat following neonatal maternal separation: gender-dependent effects. *Brain research* **1097** (1):123-132. doi:10.1016/j.brainres.2006.04.066

21. Mitra R, Vyas A, Chatterjee G, Chattarji S. (2005). Chronic-stress induced modulation of different states of anxiety-like behavior in female rats. *Neuroscience letters* **383** (3):278-283. doi:10.1016/j.neulet.2005.04.037

14. Guadagno A, Wong TP, Walker C-D. (2018). Morphological and functional changes in the preweaning basolateral amygdala induced by early chronic stress associate with anxiety and fear behavior in adult male, but not female rats. *Progress in Neuro-Psychopharmacology and Biological Psychiatry* **81**:25-37. doi:10.1016/j.pnpbp.2017.09.025

15. Koss WA, Belden CE, Hristov AD, Juraska JM. (2014). Dendritic remodeling in the adolescent medial prefrontal cortex and the basolateral amygdala of male and female rats. *Synapse* **68** (2):61-72. doi:10.1002/syn.21716

16. Eiland L, Ramroop J, Hill MN, Manley J, Mcewen BS. (2012). Chronic juvenile stress produces corticolimbic dendritic architectural remodeling and modulates emotional behavior in male and female rats. *Psychoneuroendocrinology* **37** (1):39-47. doi:10.1016/j.psyneuen.2011.04.015

17. Ehrlich DE, Ryan SJ, Rainnie DG. (2012). Postnatal development of electrophysiological properties of principal neurons in the rat basolateral amygdala. *The Journal of Physiology* **590** (19):4819-4838. doi:10.1113/jphysiol.2012.237453

18. Kumar V, Vasudevan A, Soh LJT, Le Min C, Vyas A, Zewail-Foote M, Guarraci FA. (2014). Sexual Attractiveness in Male Rats Is Associated with Greater Concentration of Major Urinary Proteins. *Biology of Reproduction* **91** (6). doi:10.1095/biolreprod.114.117903

19. Abdulai-Saiku S, Hegde A, Vyas A, Mitra R. (2017). Effects of stress or infection on rat behavior show robust reversals due to environmental disturbance. *F1000Research* **6**. doi:10.12688/f1000research.13171.1

Publications | helmy.m@protonmail.com | ProtonMail

This email is for use by the intended recipient and contains information that may be confidential. If you are not the intended recipient, please notify the sender by return email and delete this email from your inbox. Any unauthorized use or distribution of this email, in whole or in part, is strictly prohibited and may be unlawful. Any price quotes contained in this email are merely indicative and will not result in any legally binding or enforceable obligation. Unless explicitly designated as an intended e-contract, this email does not constitute a contract offer, a contract amendment, or an acceptance of a contract offer.

Elsevier Limited. Registered Office: The Boulevard, Langford Lane, Kidlington, Oxford, OX5 1GB, United Kingdom, Registration No. 1982084, Registered in England and Wales. <u>Privacy Policy</u>

From: Sharon Joy Lintao Date: Thursday, February 18, 2021 04:02 AM GMT

Dear Customer,

Thank you for contacting Elsevier Customer Service.

This is an automated acknowledgement to confirm we have received your query.

Ticket number 210214-009528 has been opened on your behalf and you can expect to receive a response within 48 hours (excluding weekends).

For immediate assistance, please visit our Elsevier Support Center.

Regards,

Elsevier Customer Service

From: Sharon Joy Lintao Date: Thursday, February 18, 2021 04:02 AM GMT

Dear Heliyon,

I was wondering if you had time to consider my message below?

Kind regards,

Mohamed Helmy

MD, PhD

helmy.m@protonmail.com

+65 83 555 815

10 Jurong Lake Link, #15-39

Singapore 648131

----- Original Message ------

On Sunday, December 20, 2020 2:54 PM, Sci Tech Jrnls CS <<u>JournalCustomerService</u>-<u>usa@elsevier.com</u>> wrote:

/span>

This email is for use by the intended recipient and contains information that may be confidential. If you are not the intended recipient, please notify the sender by return email and delete this email from your inbox. Any unauthorized use or distribution of this email, in whole or in part, is strictly prohibited and may be unlawful. Any price quotes contained in this email are merely indicative and will not result in any legally binding or enforceable obligation. Unless explicitly designated as an intended e-contract, this email does not constitute a contract offer, a contract amendment, or an acceptance of a contract offer.

Elsevier Limited. Registered Office: The Boulevard, Langford Lane, Kidlington, Oxford, OX5 1GB, United Kingdom, Registration No. 1982084, Registered in England and Wales. <u>Privacy Policy</u>

This email is for use by the intended recipient and contains information that may be confidential. If you are not the intended recipient, please notify the sender by return email and delete this email from your inbox. Any unauthorized use or distribution of this email, in whole or in part, is strictly prohibited and may be unlawful. Any price quotes contained in this email are merely indicative and will not result in any legally binding or enforceable obligation. Unless explicitly designated as an intended e-contract, this email does not constitute a contract offer, a contract amendment, or an acceptance of a contract offer.

Elsevier Limited. Registered Office: The Boulevard, Langford Lane, Kidlington, Oxford, OX5 1GB, United Kingdom, Registration No. 1982084, Registered in England and Wales. Privacy Policy