- 1 TITLE: Reproducible and Transparent Research Practices in Published Neurology Research
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Abstract Background The objective of this study was to evaluate the nature and extent of reproducible and transparent research practices in neurology research. Methods The NLM catalog was used to identify MEDLINE-indexed neurology journals. A PubMed search of these journals was conducted to retrieve publications over a 5-year period from 2014 to 2018. A random sample of publications was extracted. Two authors conducted data extraction in a blinded, duplicate fashion using a pilot-tested Google form. This form prompted data extractors to determine whether publications provided access to items such as study materials, raw data, analysis scripts, and protocols. In addition, we determined if the publication was included in a replication study or systematic review, was preregistered, had a conflict of interest declaration, specified funding sources, and was open access. Results Our search identified 223,932 publications meeting the inclusion criteria, from which 300 were randomly sampled. Only 290 articles were accessible, yielding 202 publications with empirical data for analysis. Our results indicate that 8.99% provided access to materials, 9.41% provided access to raw data, 0.50% provided access to the analysis scripts, 0.99% linked the protocol, and 3.47% were preregistered. A third of sampled publications lacked funding or conflict of interest statements. No publications from our sample were included in replication studies, but a fifth were cited in a systematic review or meta-analysis. **Conclusions** Current research in the field of neurology does not consistently provide information needed for reproducibility. The implications of poor research reporting can both affect patient care and increase research waste. Collaborative intervention by authors, peer reviewers, journals, and funding sources is needed to mitigate this problem.

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Keywords Reproducibility, transparency, neurology, cross-sectional **Background** Scientific advancement is hampered by potential research flaws, such as the lack of replication; poor reporting; selective reporting bias; low statistical power; and inadequate access to materials, protocols, analysis scripts, and experimental data.[1-3] These factors may undermine the rigor and reproducibility of published research. Substantial evidence suggests that a large proportion of scientific evidence may be false, unreliable, or irreproducible.[4-8] Estimates of irreproducible research range from 50% to 90% in preclinical sciences[9] and substantiated in a recent survey of scientists. Prior survey studies reported that roughly 70% of scientists were unable to replicate another scientist's experiment, and 90% agreed that scientific research is currently experiencing a "reproducibility crisis."[7] Reproducibility is vital for scientific advancement as it aids in enhancing the credibility of novel scientific discoveries and mitigates erroneous findings. One review discussed potential pitfalls in fMRI reproducibility, such as scanner settings, consistency of cognitive tasks, and analysis methods.[10] Boekel et al. replicated five fMRI studies measuring a total of 17 structural brain-behavior correlations. After reanalysis, only one of the 17 was successfully replicated.[11] Thus, practices related to transparency and reproducibility can be improved within fMRI and other neurology research. Adopting open science in neurology would help mitigate irreproducible research, such as the studies on brain-behavior correlation. Open science practices – such as data sharing, open access articles, sharing protocols and methods, and study preregistration – promote transparency and reproducibility.[12] For example, preregistering a study helps guard against selective outcome reporting.[13] Selective outcome

reporting occurs when discrepancies exist between outcome measures prespecified in trial registries or

research protocols and the outcomes listed in the published report. [14] In neurology, an audit of

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randomized clinical trials published in neurology journals found 180 outcome inconsistencies across 180 trials, with most inconsistencies favoring changes in accordance with statistically significant results. Additionally, only 55% of neurology trials were prospectively registered [15], providing indications that neurology researchers are not adhering to transparency and reproducibility practices early in research planning. Reproducible research and open science practices are widely endorsed by a large proportion of authors. Despite this support, evidence suggests that authors infrequently implement them.[16–18] Given the recent attention to the reproducibility crisis in science, further investigation is warranted to ensure the existence of reproducible and transparent research in the field of neurology. Here, we examine key transparency- and reproducibility-related research practices in the published neurology literature. Our findings from this investigation may serve as a baseline to measure future progress regarding transparency and reproducibility-related practices. Methods This observational, cross-sectional study used the methodology proposed by Hardwicke et. al. [3], with modifications. We reported this study in accordance with the guidelines for meta-epidemiological methodology research[19] and, when pertinent, the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA).[20] Our study did not use any human subjects or patient data and, as such, was not required to be approved by an institutional review board prior to initiation. We have used The Open Science Framework to host our protocol, materials, training video, and study data in a publically available database (https://osf.io/n4yh5/). Journal and Publication Selection On June 25, 2019, one investigator (D.T.) searched the National Library of Medicine (NLM) catalog for all journals using the subject terms tag "Neurology[ST]." The inclusion criteria required that all journals publish English, full-text manuscripts and be indexed in the MEDLINE database. The final list of

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included journals was created by extracting the electronic international standard serial number (ISSN) or the linking ISSN, if necessary. PubMed was searched with the list of journal ISSNs on June 25, 2019 to identify all publications. We then limited our publication sample to those between January 1, 2014 and December 31, 2018. Three hundred publications within the time period were randomly sampled for data extraction. The rest were available, but not needed (https://osf.io/wvkgc/). Extraction Training Prior to data extraction, two investigators (S.R. and J.P.) completed in-person training designed and led by another investigator (D.T.). The training sessions included reviewing the protocol, study design, data extraction form, and likely locations of necessary information within example publications. The two authors being trained received two sample publications to extract data from. This example data extraction was performed in the same duplicate and blinded fashion used for data acquisition for this study. The two investigators then met to reconcile any discrepancies. After the two sample publications were completed, investigators extracted data and reconciled differences from the first 10 of the included 300 neurology publications. This process insured interrater reliability prior to analyzing the remaining 290 publications. A final reconciliation meeting was conducted, with a third investigator (D.T.) available for disputes but not needed. Data Extraction After completing training, the same two investigators extracted data from the included list of randomly sampled publications between June 3, 2019 and June 10, 2019 using a pilot-tested Google form. This Google form was based on the one used by Hardwicke et al., but including modifications.[3] We specified the 5-year impact factor and that for the most recent year as opposed to the impact factor of a specific year. The available types of study designs were expanded to include case series, cohort studies, secondary analyses, chart reviews, and cross-sectional analyses. Last, we specified funding sources, such as hospital,

private/industry, non-profit, university, or mixed, instead of restricting the criteria to public or private.

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Assessment of Reproducibility and Transparency Characteristics This study used the methodology by Hardwicke et al.[3] for analyses of transparency and reproducibility of research, with modifications. Full publications were examined for funding disclosures, conflicts of interest, available materials, data, protocols, and analysis scripts. Publications were coded to fit two criteria: those with and those without empirical data. Publications without empirical data (e.g., editorials, reviews, news, simulations, or commentaries without reanalysis) were analyzed for conflict of interest statements, open access, and funding. Given that protocols, data sets, and reproducibility were not relevant, these were omitted. Case studies and case series were listed as empirical studies; however, questions pertaining to the availability of materials, data, protocol, and registration were excluded due to previous study recommendations.[18] Data extraction criteria for each study design is outlined in Table 1. Publication Citations Included in Research Synthesis and Replication For both empirical and nonempirical studies, we measured the impact factor of each journal by searching for the publication title on the Web of Science (https://webofknowledge.com). For empirical studies, we used the Web of Science to determine whether our sample of studies were cited in either a meta-analysis, systematic review, or a replication study. The Web of Science provided access to studies that cited the queried publication and provided the title, abstract, and link to the full-text article. This permitted evaluation of the inclusion of the queried article in data synthesis. Extraction was performed by both investigators in a duplicate, blinded fashion. Assessment of Open Access Important core components of publications necessary for reproducibility are only available within the full text of a manuscript. To determine the public's access to each publication's full text, we systematically searched the Open Access Button (https://openaccessbutton.org), Google, and PubMed. First, we searched the title and DOI using the Open Access Button to determine if the publication was available for public

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access. If this search returned no results or had an error, then we searched the publication title on Google or PubMed and reviewed the journal website to determine if the publication was available without a paywall. Statistical Analysis Microsoft Excel was used to report statistics for each category of our analysis. In particular, we used Excel functions to calculate our study characteristics, results, and 95% confidence intervals. **Results** Journal and Publication Selection After searching the National Library of Medicine catalog, 490 neurology journals were eligible for analysis. After screening for inclusion criteria, 299 journals remained for analysis, yielding 223,932 publications. Of the 223,932 publications, we randomly sampled 300 (https://osf.io/qfy7u/). Ten publications were inaccessible, which left 290 publications for analysis. Of the 290 eligible publications, 218 provided analyzable empirical data, and 72 articles were excluded because they did not contain characteristics measurable for reproducibility. Of the 218 publications eligible for analysis, an additional 16 case studies and case series were excluded, as they are irreproducible. Our final analysis was based on 202 publications with measurable reproducibility characteristics (Figure 1 and Table 1). Sample Characteristics Of the eligible publications, the median 5-year impact factor was 3.555 (Interquartile range (IQR): 2.421-4.745), although 20 publications had inaccessible impact factors. The United States was the location of most of the primary authors (30.69%, 89/290) and the country of most publications (56.55%, 164/290). Of the 290 publications that were accessible, 33.10% did not report a funding source (96/290), and 27.93% reported funding from mixed sources (81/290; Table 2).

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Of the randomly sampled 300 publications that were findable, 61.38% were not accessible to the public without a paywall (178/290), and only 40.34% were available to the public via the Open Access Button (117/290). Approximately half of analyzed publications stated that they did not have any conflicts of interest (53.10%, 154/290), and 33.10% did not report whether or not conflicts of interest existed (96/290). Humans were the focus of 51.72% of the analyzed publications (150/290). Additional sample characteristics are viewable in Supplemental Tables 1, 2, and 3. Reproducibility-Related Characteristics Among the 202 publications with empirical data that were analyzed, a mere 3.47% provided preregistration statements or claimed to be preregistered (7/202). Of the 202 publications, just 0.99% provided access to the protocol (2/202). Interestingly, only 8.99% provided access to the materials list (17/189), 9.41% provided access to the raw data (19/202), and just a single article provided the analysis script (0.50%, 1/202). Not a single publication claimed to be a replication study. Additional characteristics are viewable in Supplemental Tables 1, 2, and 3. **Discussion** Our analysis demonstrates inadequate reproducibility practices within neurology and neuroscience research. We found that few publications contained data or materials availability statements and even fewer contained a preregistration statement, made the protocol available, or included an analysis script. Our overall finding – that a majority of neurology publications lack the information necessary to be reproduced and transparent – is comparable to findings in the social and preclinical sciences.[3, 5, 21–24] Here, we present a discussion on prominent reproducibility and transparency indicators that were lacking in our study while presenting recommendations and practices to help improve neurology research. First, data and materials availability is essential for reproducing research. Without source data, corroborating the results is nearly impossible. Without a detailed description of materials, conducting the

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experiment becomes a guessing game. Less than 10% of publications in our sample reported either a data or a materials availability statement. Efforts toward data sharing in neurological research originated with brain mapping and neuroimaging, but has spread to other areas within the specialty to improve reproducibility, transparency, and data aggregation. [25] Although data sharing poses challenges, steps have been taken in fMRI studies. [26, 27] fMRI data are complex and cumbersome to handle, but can be managed with software, such as Automatic Analysis [28], C-BRAIN [29], and Neuro Imaging Analysis Kit.[30] Furthermore, these data can be hosted on online repositories, such as The National Institute of Mental Health Data Archive[31], Figshare[32], and other National Institutes of Health repositories.[33] Although researchers may take these steps voluntarily, journals – the final arbiters of research publications – can require such practices. Our study found that less than half of the sampled journals had a data availability policies, with approximately 20% of articles from these journals reporting source data. [34] Another study in PLOS ONE found that only 20% of nearly 50,000 publications included a data sharing statement and found that once a data sharing policy was enacted, open access to raw data increased.[35] Based on this evidence, journals and funders should consider implementing and enforcing data sharing policies that, at minimum, require a statement detailing whether data are available and where data are located. For example, the journal Neurology has endorsed the International Committee of Medical Journal Editors policy of requiring a data sharing statement and encourages open access.[36–38] If other neurology journals follow suit, an environment of transparency and reproducibility may be established. Second, preregistration practices were uncommon among neurology researchers. Preregistration prior to conducting an experiment safeguards against selective outcome reporting. This form of bias affects the quality of research in neurology. For example, when a randomized controlled trial (RCT) contains an outcome deemed "not significant" and is selectively removed from a trial, the validity of the RCT may be questioned. Previous studies have already established outcome reporting bias as an issue within neurology, noting that only 40% of analyzed RCTs were preregistered and, therefore, prespecified their

analysis.[15] This same study found outcome reporting inconsistencies that often favored statistically significant results.[15] *JAMA Neurology, The Lancet Neurology*, and *Neurology* all requiring the preregistration of clinical trials prior to study commencement in accordance with the International Committee of Medical Journal Editors (ICJME).[39] Only *The Lancet Neurology* mentions registration of other study designs, such as observational studies, and only "encourages the registration of all observational studies on a WHO-compliant registry."[40–42] The ICJME notes that although non-trial study designs lack a researcher prespecified intervention, it is recommended to preregister all study types to discourage selective reporting and selective publication of results [39]. On ClinicalTrials.gov alone, almost 65,000 observational study designs have been preregistered, comprising 21% of all registered studies [43]. Encouraging the preregistration of clinical trials and observational studies, alike, will increase transparency, increase the evidence available for systematic reviews and meta-analyses, and improve reproducibility [44, 45].

Moving Forward

We propose the following solutions to promote reproducible and transparent research practices in neurology. With regards to journals, we recommend requiring open data sharing upon submission, or, at least, a statement from the authors signifying why open data sharing does not apply to their study. There are many open data repositories available, including the Open Science Framework (https://osf.io/), opendatarepository.org, and others listed at re3data.org. Second, we recommend journals and funding providers consider incentivizing reproducible research practices. For example, the Open Science Framework awards "badges" for open research practices, such as open data sharing, materials availability, and preregistration.[46] If one or more of these reproducible research practices do not apply to a particular study, a statement as to such should still qualify for the award. One Neuroscience journal, *Journal of Neurochemistry*, has already implemented open science badges with considerable success.[47] With regards to researchers, better awareness and education is necessary to encourage transparent and reproducible practices. Organizations, such as the *Global Biological Standards Institute*, have committed

to improving the reproducibility of life sciences research through multiple methods, including training and educating researchers in effective trial design.[48, 49] The institute's president has called for and implemented training programs aimed at teaching students, postdoctoral fellows, and principal investigators the importance of robust study design.[48] Additionally, we propose that medical schools and residency programs incorporate classes and didactic programs detailing proper experimental design with an emphasis on reproducible scientific practices. Research education should be a pillar of medical education, as physicians play an important role in guiding evidence-based healthcare. We anticipate that these recommendations, if implemented, will improve reproducibility within neurology and, as a result, the quality of research produced within this specialty.

Strengths and Limitations

We feel that our methodology is robust and has many strengths, including blind and duplicate data extraction. Additionally, our protocol and data are available online to encourage reproducibility and transparency. However, we acknowledge a few limitations. First, we recognize that not all publications (clinical trials and protected patient data) are readily able to share their data and materials, although we feel a statement should still be reported. Second, we did not contact authors to obtain data, materials or analysis scripts and only used published materials for extraction. Had we contacted the authors, then source data, materials, and protocols may have been available.

Conclusions

In summary, improvement is needed to incorporate reproducibility factors in neurology research. Such necessary improvement is attainable. Authors, journals, and peer-reviewers all have a part to play in developing an improved community of patient-centered neurology researchers. Reproducibility is paramount in evidence-based medicine to corroborate findings and ensure physicians have the highest quality evidence upon which to base patient care.

List of Abbreviations

- Randomized Control Trial (RCT)
- 279 Declarations

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- 280 Ethics Approval and Consent to Participate Not Applicable
- 281 Consent for Publication Not Applicable
- 282 Availability of Data and Materials All protocols, materials, and raw data are available online via
- 283 bioRxiv (BIOARKIV/2019/763730).
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- 289 Acknowledgements Not Applicable
- 290 Author's Information Not Applicable
- 292 References

- 293 1. Wicherts JM, Borsboom D, Kats J, Molenaar D. The poor availability of psychological research data
- 294 for reanalysis. Am Psychol. 2006;61:726–8.
- 295 2. Iqbal SA, Wallach JD, Khoury MJ, Schully SD, Ioannidis JPA. Reproducible Research Practices and
- Transparency across the Biomedical Literature. PLoS Biol. 2016;14:e1002333.
- 3. Hardwicke TE, Wallach JD, Kidwell M, Ioannidis J. An empirical assessment of transparency and
- reproducibility-related research practices in the social sciences (2014-2017). 2019.
- 299 doi:10.31222/osf.io/6uhg5.

- 300 4. Ioannidis JPA. Why most published research findings are false. PLoS Med. 2005;2:e124.
- 5. Button KS, Ioannidis JPA, Mokrysz C, Nosek BA, Flint J, Robinson ESJ, et al. Power failure: why
- small sample size undermines the reliability of neuroscience. Nat Rev Neurosci. 2013;14:365–76.
- 303 6. Peng R. The reproducibility crisis in science: A statistical counterattack. Significance. 2015;12:30–2.
- 304 7. Baker M. 1,500 scientists lift the lid on reproducibility. Nature. 2016;533:452–4.
- 8. Begley CG, Glenn Begley C, Ioannidis JPA. Reproducibility in Science. Circulation Research.
- 306 2015;116:116–26. doi:10.1161/circresaha.114.303819.
- 9. Freedman LP, Cockburn IM, Simcoe TS. The Economics of Reproducibility in Preclinical Research.
- 308 PLoS Biol. 2015;13:e1002165.
- 309 10. Bennett CM, Miller MB. How reliable are the results from functional magnetic resonance imaging?
- 310 Ann N Y Acad Sci. 2010;1191:133–55.
- 311 11. Boekel W, Wagenmakers E-J, Belay L, Verhagen J, Brown S, Forstmann BU. A purely confirmatory
- 312 replication study of structural brain-behavior correlations. Cortex. 2015;66:115–33.
- 313 12. Banks GC, Field JG, Oswald FL, O'Boyle EH, Landis RS, Rupp DE, et al. Answers to 18 Questions
- 314 About Open Science Practices. J Bus Psychol. 2019;34:257–70.
- 315 13. Nosek BA, Ebersole CR, DeHaven AC, Mellor DT. The preregistration revolution. Proc Natl Acad
- 316 Sci U S A. 2018;115:2600-6.
- 317 14. Mathieu S, Boutron I, Moher D, Altman DG, Ravaud P. Comparison of registered and published
- primary outcomes in randomized controlled trials. JAMA. 2009;302:977–84.
- 319 15. Howard B, Scott JT, Blubaugh M, Roepke B, Scheckel C, Vassar M. Systematic review: Outcome
- reporting bias is a problem in high impact factor neurology journals. PLoS One. 2017;12:e0180986.

- 321 16. Harris JK, Johnson KJ, Carothers BJ, Combs TB, Luke DA, Wang X. Use of reproducible research
- practices in public health: A survey of public health analysts. PLoS One. 2018;13:e0202447.
- 323 17. Stupple A, Singerman D, Celi LA. The reproducibility crisis in the age of digital medicine. npj Digital
- 324 Medicine. 2019;2:2.
- 325 18. Wallach JD, Boyack KW, Ioannidis JPA. Reproducible research practices, transparency, and open
- access data in the biomedical literature, 2015–2017. PLoS Biol. 2018;16:e2006930.
- 327 19. Murad MH, Wang Z. Guidelines for reporting meta-epidemiological methodology research. Evid
- 328 Based Med. 2017;22:139–42.
- 329 20. Liberati A, Altman DG, Tetzlaff J, Mulrow C, Gøtzsche PC, Ioannidis JPA, et al. The PRISMA
- 330 statement for reporting systematic reviews and meta-analyses of studies that evaluate health care
- interventions: explanation and elaboration. J Clin Epidemiol. 2009;62:e1–34.
- 332 21. Prinz F, Schlange T, Asadullah K. Believe it or not: how much can we rely on published data on
- potential drug targets? Nat Rev Drug Discov. 2011;10:712.
- 22. Begley CG, Glenn Begley C, Ellis LM. Raise standards for preclinical cancer research. Nature.
- 335 2012;483:531–3. doi:10.1038/483531a.
- 336 23. Tsilidis KK, Panagiotou OA, Sena ES, Aretouli E, Evangelou E, Howells DW, et al. Evaluation of
- excess significance bias in animal studies of neurological diseases. PLoS Biol. 2013;11:e1001609.
- 338 24. Begley CG, Ioannidis JPA. Reproducibility in science: improving the standard for basic and
- preclinical research. Circ Res. 2015;116:116–26.
- 340 25. Ferguson AR, Nielson JL, Cragin MH, Bandrowski AE, Martone ME. Big data from small data: data-
- sharing in the "long tail" of neuroscience. Nature Neuroscience. 2014;17:1442–7. doi:10.1038/nn.3838.

- 342 26. Nichols TE, Das S, Eickhoff SB, Evans AC, Glatard T, Hanke M, et al. Best practices in data analysis
- and sharing in neuroimaging using MRI. Nat Neurosci. 2017;20:299–303.
- 344 27. Borghi JA, Van Gulick AE. Data management and sharing in neuroimaging: Practices and perceptions
- 345 of MRI researchers. PLoS One. 2018;13:e0200562.
- 28. Cusack R, Vicente-Grabovetsky A, Mitchell DJ, Wild CJ, Auer T, Linke AC, et al. Automatic
- analysis (aa): efficient neuroimaging workflows and parallel processing using Matlab and XML. Front
- 348 Neuroinform. 2014;8:90.
- 29. CBRAIN | Home. http://www.cbrain.ca/. Accessed 14 Aug 2019.
- 350 30. NITRC: NeuroImaging Analysis Kit (NIAK): Tool/Resource Info.
- 351 https://www.nitrc.org/projects/niak. Accessed 14 Aug 2019.
- 352 31. NDA. https://nda.nih.gov/. Accessed 14 Aug 2019.
- 353 32. Kraker P, Lex E, Gorraiz J, Gumpenberger C, Peters I. Research Data Explored II: the Anatomy and
- Reception of figshare. arXiv [cs.DL]. 2015. http://arxiv.org/abs/1503.01298.
- 355 33. NIH Data Sharing Repositories. 2013.
- 356 https://www.nlm.nih.gov/NIHbmic/nih_data_sharing_repositories.html. Accessed 14 Aug 2019.
- 34. Johnson JN, Hanson KA, Jones CA, Grandhi R, Guerrero J, Rodriguez JS. Data Sharing in
- Neurosurgery and Neurology Journals. Cureus. 2018;10:e2680.
- 35. Federer LM, Belter CW, Joubert DJ, Livinski A, Lu Y-L, Snyders LN, et al. Data sharing in PLOS
- ONE: An analysis of Data Availability Statements. PLoS One. 2018;13:e0194768.
- 36. Taichman DB, Sahni P, Pinborg A, Peiperl L, Laine C, James A, et al. Data Sharing Statements for
- 362 Clinical Trials: A Requirement of the International Committee of Medical Journal Editors. Ethiopian

- 363 journal of health sciences. 2017;27:315–8.
- 364 37. Baskin PK, Gross RA. The new Neurology: Redesigns, short articles for print, full articles online, and
- data availability policies. Neurology. 2017;89:2026–8.
- 38. Research Policies and Guidelines | American Academy of Neurology Journals.
- 367 https://www.neurology.org/research-policies-and-guidelines. Accessed 14 Aug 2019.
- 368 39. of Medical Journal Editors IC, Others. Recommendations for the conduct, reporting, editing, and
- publication of scholarly work in medical journals. 2016.
- 370 40. Instructions for Authors | JAMA Neurology | JAMA Network.
- 371 https://jamanetwork.com/journals/jamaneurology/pages/instructions-for-authors. Accessed 15 Aug 2019.
- 372 41. Research Policies and Guidelines | American Academy of Neurology Journals.
- 373 https://www.neurology.org/research-policies-and-guidelines. Accessed 15 Aug 2019.
- 374 42. Information for Authors. The Lancet Neurology. https://els-jbs-prod-
- 375 cdn.literatumonline.com/pb/assets/raw/Lancet/authors/tln-info-for-authors-1564564413020.pdf. Accessed
- 376 15 Aug 2019.
- 377 43. Trends, Charts, and Maps ClinicalTrials.gov. https://clinicaltrials.gov/ct2/resources/trends. Accessed
- 378 15 Aug 2019.
- 379 44. Dal-Ré R, Ioannidis JP, Bracken MB, Buffler PA, Chan A-W, Franco EL, et al. Making prospective
- registration of observational research a reality. Sci Transl Med. 2014;6:224cm1.
- 381 45. Ioannidis JPA. The importance of potential studies that have not existed and registration of
- 382 observational data sets. JAMA. 2012;308:575–6.
- 383 46. Blohowiak BB, Cohoon J, de-Wit L, Eich E, Farach FJ, Hasselman F, et al. Badges to acknowledge

open practices. 2013. https://osf.io/tvyxz/wiki/home/.

47. Jones J, More Content by. Connecting Research with Results: Open Science Badges. 2018.

https://www.wiley.com/network/researchers/licensing-and-open-access/connecting-research-with-results-open-science-badges. Accessed 19 Jul 2019.

48. One Way to Fix Reproducibility Problems: Train Scientists Better. The Scientist Magazine®.

https://www.the-scientist.com/news-opinion/one-way-to-fix-reproducibility-problems-train-scientists-better-30577. Accessed 19 Jul 2019.

49. Our Mission - Global Biological Standards Institute. Global Biological Standards Institute.

https://www.gbsi.org/about/who-we-are/. Accessed 14 Aug 2019.

Figure 1: Flow Diagram of Included and Excluded Studies for the Reproducibility Analysis

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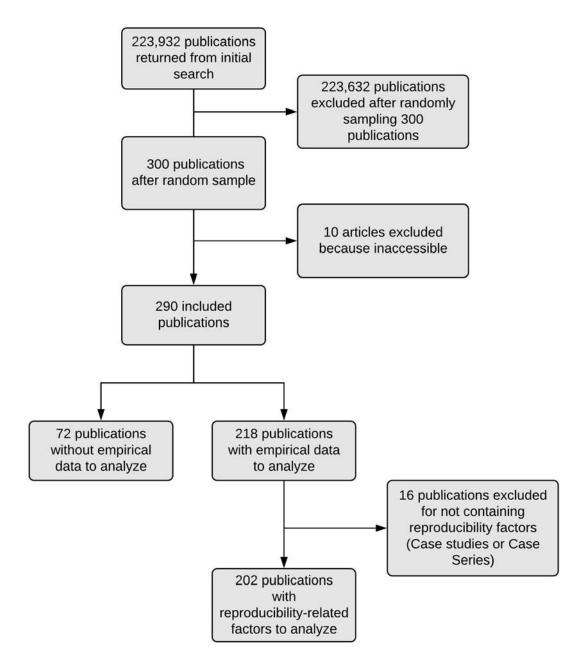


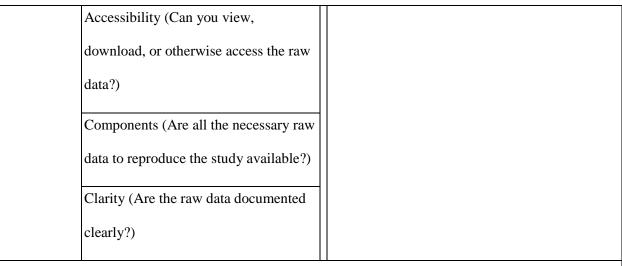
Table 1: Reproducibility related characteristics. Variable numbers (N) are dependent upon study design. Full detailed protocol pertaining to our measured variables is available online

(https://osf.io/x24n3/)

Indicators of Reproducibility Included in Present		Significance of measure variable for		
Study		transparency and reproducibility.		
Publications				
All (N=300)	Publication accessibility (Is the publication open access to the general public or accessible through a paywall?)	The general public's ability to access scientific research may increase transparency of results and improve the ability for others to critically assess studies, potentially resulting in more replication studies		
Funding				
Included studies	Funding statement (Does the publication state their funding	Explicitly providing source of funding may help mitigate bias and potential conflicts of		
(N=290)	sources?)	interest		
Conflict of I	nterest			
Included studies (N=290) Publication	Conflict of interest statement (Does the publication state whether or not the authors had a conflict of interest?) Citations	Explicitly providing conflicts of interest may allow for full disclosure of factors that may promote bias in the study design or outcomes		
Empirical studies†	Citations by a systematic review/meta- analysis (Has the publication been	Systematic reviews and meta-analyses evaluate and compare existing literature to assess for		

(N=205)	cited by any type of data synthesis	patterns, strengths, and weaknesses of studies
(14-203)		
	publication, and if so, was it explicitly	regarding a particular field or topic
	excluded?)	
Analysis Sci	ripts	1
	Availability statement (Does the	
	publication state whether or not the	
	analysis scripts are available?)	Providing access to the analysis script helps
Empirical	Method of availability (Ex: Are the	improve credibility by providing the replicators
studies‡	analysis scripts available upon request	the opportunity to analyze raw data with the
(N=202)	or in a supplement?)	same analysis procedure
	Accessibility (Can you view,	
	download, or otherwise access the	
	analysis scripts?)	
Materials		
	Availability statement (Does the	
	publication state whether or not the	
	materials are available?)	
Empirical	Method of availability (Ex: Are the	Providing the materials list allows replicators to
studies¶	materials available upon request or in a	reproduce study using the same materials,
(N=189)	supplement?)	promoting
	Accessibility (Can you view,	
	download, or otherwise access the	
	materials?)	
	1	<u> </u>

Pre-registra	tion	
Empirical studies‡ (N=202)	Availability statement (Does the publication state whether or not it was pre-registered?) Method of availability (Where was the publication pre-registered?) Accessibility (Can you view or otherwise access the registration?) Components (What components of the publication were pre-registered?)	Pre-registering studies may help mitigate potential bias and increase the overall validity and reliability of a study
Protocols		4
Empirical studies‡ (N=202)	Availability statement (Does the publication state whether or not a protocol is available?) Components (What components are available in the protocol?)	Providing replicators access to protocols allows the for more accurate replication of the study, promoting credibility
Raw Data		
Empirical studies‡ (N=202)	Availability statement (Does the publication state whether or not the raw data are available?) Method of availability (Ex: Are the raw data available upon request or in a supplement?)	Providing replicators with access to raw data can help reduce potential bias and increase validity and reliability



- † 'Empirical studies' are publications that include empirical data such as: clinical trial, cohort, case series, case reports, case-control, secondary analysis, chart review, commentaries (with data analysis), laboratory, surveys, and cross-sectional designs.
- ‡ Empirical studies determined to be case reports or case series were excluded in regard to reproducibility related questions (materials, data, protocol, and registration were excluded) as recommended by Wallach et al.

¶ Empirical studies determined to be either case reports, case series, commentaries with analysis, metaanalysis or systematic review were excluded as they did not provide materials to fit the category.

Ta	Table 2: Characteristics of Included Publications		
Ch	aracteristics	Variables	
		N (%)	95% CI
	University	11 (3.79)	1.63-5.95%
	Hospital	1 (0.34)	0-1.01%
	Public	54 (18.62)	14.22-23.03%
Funding	Private/Industry	15 (5.17)	2.67-7.68%
N=290	Non-Profit	11 (3.79)	1.63-5.95%
	Mixed	81 (27.93)	28.89-30.53%
	No Statement Listed	96 (33.01)	27.78-38.43%
	No Funding Received	21 (7.24)	4.31-10.17%
	No Empirical Data	72 (24.83)	19.94-29.72%
	Meta-Analysis	12 (4.14)	1.88-6.39%
	Commentary with		0-1.01%
	Analysis	1 (0.34)	0 110170
	Cost-Effectiveness	1 (0.34)	0-1.01%
	Clinical Trial	26 (8.97)	5.73-12.20%
Type of Study	Case Study	9 (3.10)	1.14-5.07%
N=290	Case Series	7 (2.41)	0.68-4.15%
	Cohort	43 (14.83)	10.81-18.85%
	Chart Review	3 (1.03)	0-2.18%
	Case Control	9 (3.10)	1.14-5.07%
	Survey	5 (1.72)	0.25-3.20%
	Cross-Sectional	34 (11.72)	8.08-15.36%
	Secondary Analysis	2 (0.69)	0-1.63%

	Laboratory	65 (22.41)	17.69-27.13%
	Multiple Study Types	1 (0.34)	0-1.01%
	Median	3.555	-
5 Year Impact			
_	1st Quartile	2.421	-
Factor			
	3rd Quartile	4.745	-
N=278			
	Interquartile Range	2.421-4.745	-

Supplemental 1: Additional Characteristics of Reproducibility in Neurology **Studies** Characteristics Variables N (%) 95% CI Statement, one or more conflicts of **Conflict of** 40 (13.79) 9.89-17.70% interest Interest Statement. no conflict of interest 154 (53.10) **Statement** 47.46-58.75% (N=290)No conflict of interest statement 96 (33.10) 27.78-38.43% Statement, some data are available 19 (9.41) 6.10-12.71% Data 0 **Availability** Statement, data are not available 0 (N=202)No data availability statement 183 (90.59) 87.29-93.90% Statement, some materials are available 17 (8.99) 5.76-12.23% Material 0 0 Statement, materials are not available **Availability** (N=189)No materials availability statement 172 (91.01) 87.77-94.24% **Protocol** Full Protocol 2 (0.99) 0-2.11% **Availability** 200 (99.01) No Protocol 97.89-100% (N=202)Statement, some analysis scripts are 1 (0.50) 0-1.29% available **Analysis Scripts** (N=202)Statement, analysis scripts are not 0 0 available

	No analysis script availability statement	201 (99.50)	98.71-100%
Replication	Reports Replication Study	0	0
Studies	No Clear Statement	202 (100)	100%
(N=202)	No Clear Statement	202 (100)	100%
Onen Access	Yes - found via Open Access Button	117 (40.34)	34.79-45.90%
Open Access (N=290)	Yes - found article via other means	5 (1.72)	0.25-3.20%
	Could not access through paywall	178 (61.38)	55.87-66.89%
Cited in a	No Citations	163 (79.51)	74.94-84.08%
Systematic	A Single Citation	28 (13.66)	9.77-17.54%
Review/	One to Five Citations	13 (6.34)	3.58-9.10%
Meta-Analysis (a) (N=205)	More Than 5 Citations	1 (0.49)	0-1.28%

Abbreviations: CI, Confidence Interval. a - No studies were explicitly excluded from the systematic reviews or meta-analyses that cited the original article.

Supplemental 2: Additional Characteristics of Reproducibility in Neurology **Studies** Characteristics Variables N (%) 95% CI Statement, says was pre--registered 7 (3.47) 1.40-5.54% Pre-Statement, says was not pre-registered 0 0 Registration No, there is no pre-registration statement (N=202)195 (96.53) 94.46-98.60% Animals 51 (17.59) 13.28-21.89% Humans 150 (51.72) 46.07-57.38% **Test Subjects** (N=290)Both 0 0 Neither 89 (30.69) 25.47-35.91% United States 164 (56.55) 50.94-62.16% UK 69 (23.79) 18.97-28.61% Netherlands 24 (8.28) 5.16-11.39% **Country of** Germany 6 (2.07) 0.46-3.68% **Journal** Ireland 6 (2.07) 0.46-3.68% **Publication** (N=290)Switzerland 4 (1.38) 0.06-2.70% Taiwan 1 (0.34) 0-1.10% Other (a) 16 (5.52) 2.93-8.10% United States 89 (30.69) 25.47-35.91% **Country of** 20 (6.90) Corresponding China 4.03-9.76%

Author	UK	19 (6.55)	3.75-9.35%
(N=290)	Netherlands	5 (1.72)	0.25-3.20%
	Turkey	7 (2.41)	0.68-4.15%
	France	10 (3.45)	1.38-5.51%
	Canada	13 (4.48)	2.14-6.82%
	Italy	12 (4.14)	1.88-6.39%
	Brazil	7 (2.41)	0.68-4.15%
	Australia	17 (5.86)	3.20-8.52%
	Unclear	0	0
	Other	91 (31.38)	31.38-26.13%

Supplemental 3: Additional Characteristics of Reproducibility in **Neurology Studies** Characteristics Variables N Personal or institutional 0 Supplementary information hosted by the 13 journal Material Online third party 3 **Availability** Upon Request 1 Yes, material was accessible 11 No, material was not accessible Personal or institutional Supplementary journal information 11 Online third party 4 Upon Request 4 Other (b) 0 4 Yes, data could be accessed and downloaded Data No, data count not be accessed and downloaded 15 **Availability** Yes, data files were clearly documented 1 No, data files were not clearly documented Yes, data files contain all raw data 1 No, data files do not contain all raw data 3 Unclear if all raw data was available 0

	Yes, there was a pre-regisration	7
	Registered on ClinicalTrials.Gov	4
Pre-	Registered on Other	3
Registration	Hypothesis was pre-registered	2
	Methods were pre-registered	7
	Analysis plan was pre-registered	6