

RESEARCH ARTICLE

Do funding applications where peer reviewers disagree have higher citations? A cross-sectional study. [version 2; referees: 2 approved]

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Abstract

Background: Decisions about which applications to fund are generally based on the mean scores of a panel of peer reviewers. As well as the mean, a large disagreement between peer reviewers may also be worth considering, as it may indicate a high-risk application with a high return.

Methods: We examined the peer reviewers' scores for 227 funded applications submitted to the American Institute of Biological Sciences between 1999 and 2006. We examined the mean score and two measures of reviewer disagreement: the standard deviation and range. The outcome variable was the relative citation ratio, which is the number of citations from all publications associated with the application, standardised by field and publication year.

Results: There was a clear increase in relative citations for applications with a better mean. There was no association between relative citations and either of the two measures of disagreement.

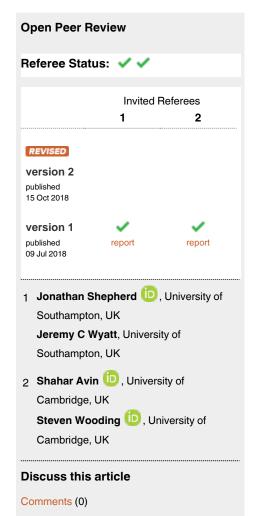
Conclusions: We found no evidence that reviewer disagreement was able to identify applications with a higher than average return. However, this is the first study to empirically examine this association, and it would be useful to examine whether reviewer disagreement is associated with research impact in other funding schemes and in larger sample sizes.

Kevwords

meta-research, research funding, peer review, citations, research impact



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REVISED Amendments from Version 1

We have added a new analysis of the variability in citations (using the inter-quartile range) compared against the application score statistics of the mean and standard deviation. We have added some additional analyses to those available on github, including a complete case analysis, exclusion of small panels, and examination of funding "buckets". Figure 4 has been replaced.

See referee reports

Introduction

Winning funding is an important stage of the research process and researchers spend large amounts of their time preparing applications¹. Applications are typically assessed using relatively small panels of 3 to 12 peer reviewers, sometimes including external reviewers with additional expertise, which is similar to the journal process of editors and reviewers. Given the importance of funding processes to researchers' careers and the progress of science, there is surprisingly little research on whether funding systems reliably identify the best research. A recent literature review found there are many unanswered questions in funding peer review, and concluded, "there is a need for open, transparent experimentation and evaluation of different ways to fund research"2. An earlier systematic review similarly concluded that studies to examine the accuracy and soundness of funding peer review are "urgently needed"3. Whilst a systematic review of innovations focused specifically on studies aiming to improve the effectiveness and efficiency in peer review funding found only eight studies and called for more studies of peer review4.

The majority of funding systems rank applications using the mean score from the review panel, and award funding from the highest to the lowest ranked applications, stopping when the budget is exhausted (exceptions are sometimes made for applications below the funding line because of national research priorities). An interesting recent idea is that an application's mean score may not be the only statistic worth considering, and that the standard deviation in peer reviewers' scores may also be a useful statistic for ranking applications⁵. A zero standard deviation means all panel members gave the same score. Larger standard deviations indicate more disagreement between panel members, and this disagreement may be useful for identifying high-risk research that may also have a higher return. A related alternative funding system is to fund all applications where reviewers agree on a high score, and then allocate the remaining budget at random where the reviewers disagreed but some reviewers gave the application a high score⁶.

Using the mean score for ranking may allow panel members to "sink" an application by awarding a low score that pulls the mean below the funding line. Including a measure of reviewer disagreement in funding could ameliorate such "sinking" and allow applications that have strong support from a few reviewers to be supported. This may also increase the diversity in what kinds of applications are funded. Some peer review systems already recognise this issue by giving each panel member a wildcard which allows them to "float" an application above the funding line regardless of other panel members' scores. At

least one funding scheme also includes patient and stakeholder reviewers to increase the diversity of viewpoints⁷.

A recent literature review found "suggestive" evidence that funding peer review can have an anti-innovation bias², whilst a survey of applicants and reviewers found that innovation and risk may not often be sufficiently addressed in review feedback⁸. There is evidence that riskier cross-disciplinary research has lower success rates⁹. Some researchers feel they need to write conservative applications that please all members of the panel to achieve a good mean score¹⁰. However, supporting risky research can have huge benefits for society when it pays off¹¹. In a survey of Australian researchers, 90% agreed with the statement: "I think the NHMRC [the main Australian funding body for health and medical research] should fund risky research that might fail but, if successful, would change the scientific field"¹².

Previous studies have investigated the association between an application's mean score (or ranking based on the mean) and subsequent citations, where citations are used as a measure of success. Many studies using large sample sizes found either no association or only a weak association between the mean score and the number of citations of subsequent publications^{13–17}. Other studies have shown a positive association between better mean peer review scores and increased citations^{18,19}, including a study that used the same data analyzed here²⁰. To our knowledge, no previous study has empirically estimated how the disagreement in peer reviewers' scores may also predict citations.

Methods

Application data

We examined 227 successful grant applications submitted to the American Institute of Biological Sciences between the years 1999 to 2006. These successful applications came from 2,063 total applications (overall 11% success rate). The applications covered a wide range of biomedical research areas, including vision, drug abuse, nutrition, blood-related cancer, kidney disease, autoimmune diseases, malaria, tuberculosis, osteoporosis, arthritis and autism. Applications were assessed by between 2 and 18 peer reviewers, with a median of 10 reviewers. Panels evaluated an average of 25 applications over two days. Ninety percent of applications were reviewed by on-site panels with an average size of 10 reviewers, and 10% of applications were reviewed via teleconferences of 3 reviewers. Further details on the funding process is available in a previous study of how the applications' mean scores predicted citations²⁰.

Our key predictor is the peer reviewers' scores. Individual peer reviewers, who were not conflicted, scored applications between 1.0 (best) to 5.0 (worst) in 0.1 increments. To determine funding, the score was averaged across all reviewers. In this study we also consider statistics that measure within-panel disagreement which are the standard deviation and the range (largest minus smallest score).

Citation counts

The primary outcome is the citation counts from publications associated with the successful application. The publication data

for the funded applications were taken from the mandatory final reports submitted by the applicants. On average, these reports were submitted 5 years after the application's peer review. Publications were produced from 1 to 8 (average 4.3) years after the review date. Only peer-reviewed publications were counted, confirmed through *PubMed* and *Web of Knowledge* searches. Publications listed in the final report as "submitted" or "in preparation", were included if they could be found as peer-reviewed published papers. Citations were counted in 2014 using *Web of Knowledge*.

This analysis used 20,313 citations from 805 peer reviewed publications. The total citation level per funded application was the cumulative citations of all publications. As citations are time-dependent they were standardized using the average citation level of all publications by scientific field and year, using data from a published calculation using the *Thomson Reuters Essential Science Indicators database*²¹. These published average rates were determined for 2000 to 2010 by scientific field, assessed in 2011 and displayed a linear relationship with time (e.g., $R^2 = 0.99$ for the field of molecular biology). We chose molecular biology because it was the highest cited field and in general was the field most applicable to the funded applications.

Because the *Reuters* curve was assessed in 2011, we extended the curve for 2014, back calculating using a linear fit which had a very high R² of 0.99. In this way, we could most accurately standardize the data for the relationship between publication date and citation level and could calculate the Total Relative Citation per application. A total relative citation of 1 meant the application achieved the average number of citations, whereas values above 1 meant a higher than average number of citations.

We note that a recent study that used both unadjusted citation counts and relative citations, found the two measures gave similar results when used as the key outcome variable²².

Statistical analyses

To graphically examine associations we used scatterplots of the total relative citations against the application score statistics. If a disagreement in scores indicates a high-risk high-return project, then there may be a greater variation in citations, with more unusually low and high citations for larger disagreements in scores. To examine this we plotted the estimated inter-quartile range in total relative citations by grouping applications using a scatterplot smoothing span²³. We used a span of 20 applications which was based on trial and error, and weighted the estimated inter-quartile ranges using a Gaussian kernel²³. We used the inter-quartile range instead of the standard deviation because of the strong skew in citations, and the standard deviation was strongly influenced by the application with the highest citations.

Regression model

The total relative citations were modelled using a multiple regression model. We ran two models with the three application variables:

- 1. Review year, mean score, score standard deviation
- 2. Review year, mean score, score range

Our aim was to examine two measures of panel disagreement: the standard deviation and range. We included mean score because it has already been shown to be an important predictor for these data²⁰ and we were interested in the additional value of a measure of disagreement. We adjusted for review year (1999 to 2006) because there was a difference in the application score statistics over time, and because year was associated with citation numbers, hence it was a potential confounder.

The citations were first base e log-transformed because of their positive skew. We added a small positive constant of 0.1 before using the log-transform because some citations were zero. The estimates were back-transformed and plotted to show the results on the original relative citations scale. Using equations the multiple regression model was:

$$\log_{e}(Y_{i} + 0.1) \sim N(\mu_{i}, \sigma^{2}), \qquad i = 1, ..., N,$$

$$\mu_{i} = \beta_{0} + \sum_{j=1}^{3} \beta_{j} f(X_{i,j}) + \gamma_{p(i),}$$

$$\gamma_{k} \sim N(0, \sigma_{v}^{2}), \qquad k = 1, ..., M,$$

where **Y** are the citations and γ are random intercepts to adjust for the potential within-panel correlation in citations (where p(i) is the panel number for application i and M is the total number of panels). The mean (μ) had a constant (β_0) and the three application predictors (\mathbf{X}) which were first transformed using a fractional polynomial function.

Associations between the score statistics and citation outcomes could be non-linear, for example a larger difference in citations for a change in mean score from 1.0 to 1.1 compared with a change from 2.0 to 2.1. To model this potential non-linearity we used fractional polynomials to examine a range of non-linear associations between the scores and citations²⁴. The fractional polynomial function is:

$$f(X_i) = \begin{cases} X_i^p, & P \neq 0, \\ \log_e(X_i), & P = 0. \end{cases}$$

We examined the eight transformations of: $P = \{-2, -1, -0.5, 0, 0.5, 1, 2, 3\}$ and chose the optimal P using the deviance. The optimal P was chosen for each of the three predictors, meaning we examined $8^3 = 512$ models in total. We only present results for the best model with the smallest deviance, but the results for the five best deviances are available: https://github.com/agbarnett/funding.disagree.

We checked the distribution of the model residuals for multimodality and outliers, and used Cook's distance to find influential observations.

Missing data

Sixteen (7%) observations were missing the score standard deviation and 32 (14%) observations were missing the score range because the individual peer reviewer scores were no longer available for some applications at the time of this retrospective analysis. These missing observations were imputed using linear regression with the application variables: review year,

mean score, score standard deviation, minimum score, maximum score and range. We used five multiple imputations.

All analyses were made using R version 3.4.4²⁵ with the imputations using the "MICE" package²⁶. The code and anonymized data are available here: http://doi.org/10.5281/zenodo.1452073²⁷.

We report our results using the STROBE guidelines for observational research²⁸.

Results

The histograms in Figure 1 show the distributions of total relative citations and the application score statistics: mean, standard deviation, minimum, maximum and range (maximum minus minimum). There was a strong positive skew in citations with one outlying relative citation of 104; the next largest citation was 34. To counter this positive skew we used a base e log transform in later analyses. There was also a positive skew in the score standard deviation and range, and we also log-transform these predictors.

The scatter-plots in Figure 2 show the associations between the application score statistics. There was a strong correlation between the standard deviation and maximum (0.80), but not between the standard deviation and minimum (0.05). This indicates the largest disagreement is where at least one panel member has given a poor score (remembering that the best possible score is 1.0 and the worst 5.0). Applications where there was one dissenting panel member with a good score were unlikely to be funded as their mean score would not be competitive, and hence are not in this sample. There was a strong positive

correlation between the two measures of panel disagreement, the standard deviation and range (0.93).

The scatter-plots in Figure 3 show the association between total relative citations and the score statistics. We used the log-transformed citations and the standard deviation and range to remove the skew and so show a clearer association. The variance in log-citations appears relatively stable over the score statistics, somewhat confirming the validity of log-transforming the citations²⁹. The points along the bottom of the y-axis are the 74 applications (33%) with no citations. Some association between mean score and citations is visible, with a generally downward pattern in citations for increasing score. There is no clear association between citations and either the standard deviation or range.

The inter-quartile ranges in total relative citations by the application scores' mean and standard deviation are in Figure 4. There was a general reduction in the inter-quartile range as the application score mean increased. The interquartile range also reduced somewhat as the application score standard deviation increased, although the reduction was not as clear as that for the mean

The results from the multiple linear regression models are in Table 1. Only the application's mean score had a statistically significant association with citation numbers.

The predictions from the multiple linear regression models are in Figure 5. There was a reduction in citations for applications with a worse mean score. The mean lines are flat for both

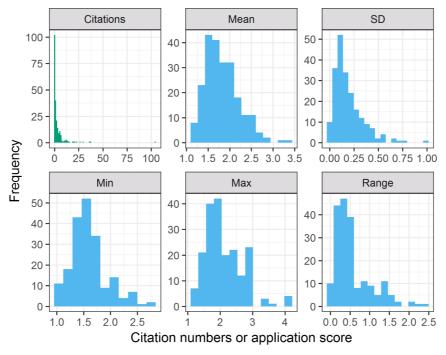


Figure 1. Histograms of total relative citations (green) and application score statistics (blue). The lower the mean score, the better the application did in peer review.

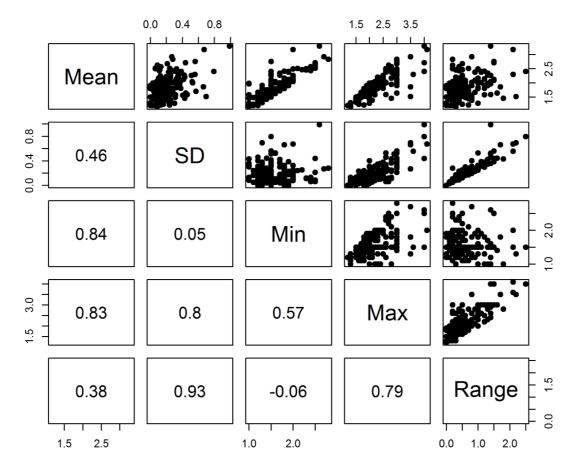


Figure 2. Scatter-plots and Pearson correlations of application score statistics. The numbers in the bottom-left diagonal of the plot matrix are the Pearson correlations.

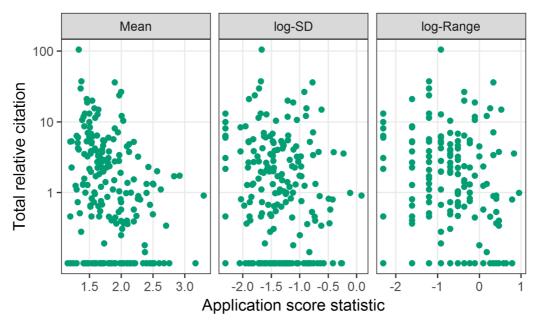


Figure 3. Scatter-plots of log-transformed total relative citations against application score statistics.

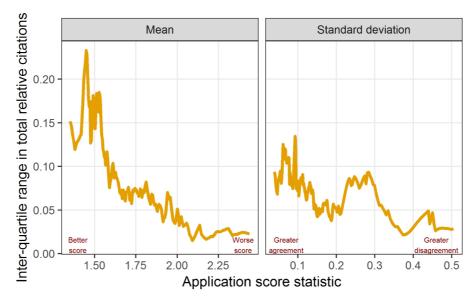


Figure 4. Scatter-plots of the standard deviation in total relative citations against application score statistics.

Table 1. Parameter estimates for the multiple regression models predicting citation numbers. FP = fractional polynomial.

Model 1	FP	Mean (95% CI)
Mean score	-0.5	5.9 (2.6 to 9.1)
Standard deviation	3	0.7 (-1.3 to 2.7)
Review year	-0.5	-1.2 (-3.0 to 0.6)
Model 2	FP	Mean (95% CI)
Model 2 Mean score	FP -0.5	Mean (95% CI) 5.4 (2.2 to 8.7)
Mean score	-0.5	` ,

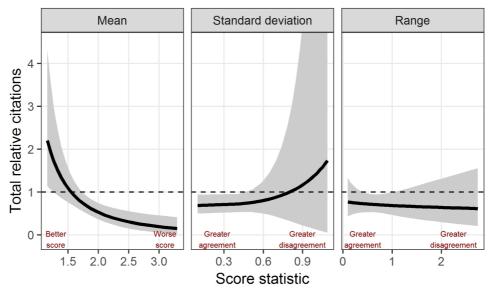


Figure 5. Predicted relative citations using the best fractional polynomial for the application score statistics. The solid lines are the means and the grey areas are 95% confidence intervals. The dotted horizontal line at 1 represents the average citations, so values above this are better than average.

the standard deviation and range, indicating no association between these score statistics and citation numbers. The 95% confidence intervals for the standard deviation are very wide for large standard deviations. The application with the largest standard deviation was influential according to Cook's statistic, and removing this application had little impact on the mean line but did reduce these wide intervals (see additional results at https://github.com/agbarnett/funding.disagree). The model residuals had an approximately symmetric and unimodal distribution with no outliers.

Discussion

We found a statistically significant association between an application's mean score and subsequent citations, with the result in the expected direction because applications with better scores had more citations (on average). The largest size of the increase is also practically significant as the highest scores have a mean of 2 (Figure 5), meaning double the average citations.

We found no association between the two measures of reviewer disagreement and citations counts. It appears that any disagreement between peer reviewers did not indicate an application with a potentially high return.

Disagreements between reviewers about an application can stem from different sources. Disagreements about the proposed methods may mean the study is not viable and would struggle to produce valid results and/or publish papers. Disagreements about the application's goals may reflect a difference of opinion about the potential impact, and it is these disagreements that are likely more subjective and hence where a higher return is possible if one reviewer is right. Disagreements between reviewers can also occur for more trivial reasons such as the dynamics of the panel and personal disagreements³⁰. A more sophisticated measure of panel disagreement to those used here may be more predictive of the benefits of the research, but such measures would need to be well-defined and prospectively recorded at the panel meeting. It may be possible to measure disagreement using an observer who watches the panel dynamics^{30,31}. Some reviews already breakdown scores into separate areas, such as track record and innovation, and reviewer disagreement could be examined using these separate scores.

Each reviewer brings their own experience and biases to the funding process and such intellectual differences influence application scores^{32,33}. Indeed, some recent research has indicated that there is more variability in scores across reviewers than across proposals³⁴ and previous studies indicate interrater reliability as very low³⁵. An application's average score is somewhat due to the "luck of the draw" of what reviewers were selected³⁶. Variations can also occur because of the way the application is summarised at the panel meeting^{30,37}.

If a larger disagreement leads to high-risk returns then we might expect an increase in the variance in citations for larger score standard deviations and ranges. This is because there might be more "failures" with zero citations, but also more big returns. Our multiple regression models only examined a change in mean citations and a different statistical model would be needed to examine a change in variance. However, the scatter-plots in Figure 3 show no sign of an increasing variance in citations for higher standard deviations or ranges, and the inter-quartile ranges in citations in Figure 4 show a slight decrease for greater disagreements.

Limitations

Some have argued that studies like ours are invalid because: 1) they only consider funded applications and do not include unfunded studies, 2) an application's score is not the only criteria used to award funding (e.g., applications with low scores awarded funding because of national priorities), and 3) because budgets are frequently cut, meaning the actual research may differ from the application³⁸. Studies that follow funded and unfunded fellowship applicants are possible, e.g., Bornmann et al (2008)³⁹, but this is very difficult when examining projects that need specific funding⁴⁰. We believe, despite the limitations of bibliometric measures, it is reasonable to expect a dose-response association between scores and citations within funded applications. Samples that include applications that were funded for reasons other than their mean score, such as national priorities, increase the variance in the key predictors of application scores statistics and hence increase statistical power. Cuts to the budget are important and can hinder the planned research. However, assuming the reviewers believed the study was still viable with the reduced budget, such studies still test the ability of a panel to predict what research will have the greatest return.

Citations are an imperfect measure of the impact of research because many citations have little worth and scientists often report that their most highly cited work is not their best⁴¹. Studies that examine more detailed outcomes such as translation into practice or cost-benefits would be incredibly useful, however these studies would themselves require funding as it would involve further data collection, analyses and interviews of the applicants.

Our results may not be generalisable to other funding schemes, especially as there are large differences between fields in their perceptions of what makes a good application³⁰. It would be useful to examine whether reviewer disagreement is associated with research impact in other funding schemes. It would also be useful to repeat the study in larger sample sizes, particularly because any conclusions could be influenced by a small proportion of applications that have a very high pay-off.

We only had summary statistics on the application scores and hence we could not examine the distribution of scores to look for interesting patterns such as bimodality in scores, indicating a strong split in the peer review panel.

Conclusions

We found no association between two measures of reviewer disagreement when assessing an application and the subsequent research impact of that application as measured by citation counts.

Data availability

The code and anonymized data are available here: https://doi.org/10.5281/zenodo.1452073²⁷

Author contributions

AGB had the idea for this analysis, performed the statistical analysis, and wrote the first draft of the manuscript. SG gave input to this study design, and conceived and designed the original experiments. SRG was involved in the data collection and commented on the draft manuscript.

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The notion that disagreements amongst reviewers might indicate a promising high-risk/high-reward project certainly has currency amongst those who consider how the system might be improved. In addition to the framework of Linton (2016)¹, the idea is also present in the model of Brezis (2007) and is implied by several works on the anti-innovation bias of grant peer review (e.g. Greenberg, 1998²; Gillies, 2008³). However, this intuitive notion has yet to be put to the test, until the current study, which is a welcome contribution.

The experimental design and statistical analysis are compelling for an initial study, though as the authors note further complications could be addressed in later work.

Given the prevailing policy and academic discussions the negative result is surprising and it rules out some suggested approaches to reform of the grant allocation system; however, the dataset limits the generality of the conclusions that can be drawn (through no fault of the authors) – the key limitation being that only 11% of applications can be analysed (the funded ones), a set that is heavily skewed towards the top scoring applications.

For example, as shown in Figure 4 there is plenty of potential for SD to be associated with increased citation, but not enough data to tell. The authors note that the application with the largest SD drove this large uncertainty – it could be that this application is an outlier, or it could be that it is typical of applications with large SDs but there are very few of them in the dataset because they tended to fall below the funding line.

Through its analysis, the study forces a refinement of the question of what sorts of disagreements (in extent and driven by what underlying logics) should be considered useful indicators. We suggest the study raises the following THREE questions for future work:

Firstly it would be valuable to understand the reasons and logics behind the disagreements that arise between reviewers, from the fact that "disagreements between reviewers about an application can stem from different sources". While the authors suggest addressing this with further information about scores, we can also ask if different score means might indicate different "regimes" for disagreement, e.g. disagreement along the lines of "not sure this will work" or "this goes against received wisdom" at relatively high score means, and disagreements along the lines of "not sure what this is about" or "hasn't



this been tried before?" at lower score means. Qualitative research asking reviewers about their scoring behaviour could help understand the reasoning that goes with different scoring.

Secondly, we could reframe the question to address the lack of information on unfunded applications by asking how can funding best be allocated if the amount of funding available is cut by 50%.

To test this, we reanalysed the data provided by the authors with the following question: assuming only half the funding was available, which of the following selection methods would perform better?

- 1. Rank proposals based on their mean scores until funding runs out.
- 2. Pool proposals based on their mean score into buckets, rounding to the nearest integer. Fund all proposals with a mean of 1 (1.0-1.49, n=45), then from the bucket with a mean of 2 (1.5-2.49, n=114) rank proposals based on their standard deviation and fund until funding runs out.

Under both methods, all proposals with a mean of 1 are funded, for a total TRC of 319.9. When we look at the proposals in the second bucket, when funding according to method 1, the total TRC for this bucket is 224.2, whereas for method 2 it is 195.5. However, the highest citation-receiving proposal in the second bucket under method 1 has a TRC of 19.9, whereas the highest citation-receiving proposal from the second bucket under method 2 has a TRC of 35.9 (the third highest TRC in the entire sample). This fits with Gillies' description of peer-review as going for less risky proposals, but at the cost of throwing out the occasional exceptional proposal. This is also born out by the distribution of TRCs, with the standard deviation of TRCs for the second bucket under method 1 being 4.2, and under method 2 being 6.2.

Thirdly, it would be valuable to understand what sorts of disagreement exist in the scoring of all the applications (both successful and unsuccessful). It could be argued that the type of disagreement likely to indicate high risk/high return proposals would be bi-modal – some good reviews maybe with scores of 1-1.5, some very poor reviews maybe with scores of 3-4. Given the small fraction of applications funded, very few applications of this nature would be funded (due to the understandable aggregation in the shared data we could not examine the exact number, but fewer than 10 applications have a score lower than 3). Do such applications with such bi-modal review scores exist? Or is the level of disagreement seen in the successful applications similar to that of the unsuccessful applications?

The above questions only suggest that there is further complexity here that needs to be explored, with more complex, higher power studies. We thank the authors for paving the path for such studies, and for making their underlying data and analysis available in a form that is easy to explore.

Two small notes:

- 1. It might be easier to read the paper if scores were consistently referred to as best/worst and better/worse using 'higher' is confusing when better scores are lower.
- 2. It would be more elegant to give 'e.g., Bornmann et al, 2008' rather than 'e.g., 36'.

References

- 1. Linton J: Improving the Peer review process: Capturing more information and enabling high-risk/high-return research. *Research Policy*. 2016; **45** (9): 1936-1938 Publisher Full Text
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- 3. Gillies D: How should research be organised?. College Publications. 2008.

Is the work clearly and accurately presented and does it cite the current literature? Yes



Is the study design appropriate and is the work technically sound? Yes

Are sufficient details of methods and analysis provided to allow replication by others? Yes

If applicable, is the statistical analysis and its interpretation appropriate? $\ensuremath{\text{Yes}}$

Are all the source data underlying the results available to ensure full reproducibility? Yes

Are the conclusions drawn adequately supported by the results? $\ensuremath{\text{Yes}}$

Competing Interests: No competing interests were disclosed.

We have read this submission. We believe that we have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Author Response 10 Oct 2018

Adrian Barnett, Queensland University of Technology, Australia

The notion that disagreements amongst reviewers might indicate a promising high-risk/high-reward project certainly has currency amongst those who consider how the system might be improved. In addition to the framework of Linton (2016)1, the idea is also present in the model of Brezis (2007) and is implied by several works on the anti-innovation bias of grant peer review (e.g. Greenberg, 1998 2; Gillies, 2008 3). However, this intuitive notion has yet to be put to the test, until the current study, which is a welcome contribution.

Thanks for the interesting references, we have now mentioned Brezis in our Introduction.

The experimental design and statistical analysis are compelling for an initial study, though as the authors note further complications could be addressed in later work.

Given the prevailing policy and academic discussions the negative result is surprising and it rules out some suggested approaches to reform of the grant allocation system; however, the dataset limits the generality of the conclusions that can be drawn (through no fault of the authors) – the key limitation being that only 11% of applications can be analysed (the funded ones), a set that is heavily skewed towards the top scoring applications.

We agree this is a key limitation, and it would be useful to repeat the study in a more generous scheme, where a greater proportion of applications were funded.

For example, as shown in Figure 4 there is plenty of potential for SD to be associated with increased citation, but not enough data to tell. The authors note that the application with the largest SD drove this large uncertainty – it could be that this application is an outlier, or it could be that it is typical of applications with large SDs but there are very few of them in the dataset because they tended to fall below the funding line.

This is entirely possible, and if there are a small number of extremely high pay-off projects then we would need a much larger sample to robustly test if there is a consistent pattern of more high pay-off projects associated with greater disagreement. We have now mentioned this in the limitations.

Through its analysis, the study forces a refinement of the question of what sorts of disagreements



(in extent and driven by what underlying logics) should be considered useful indicators. We suggest the study raises the following THREE questions for future work:

Firstly it would be valuable to understand the reasons and logics behind the disagreements that arise between reviewers, from the fact that "disagreements between reviewers about an application can stem from different sources". While the authors suggest addressing this with further information about scores, we can also ask if different score means might indicate different "regimes" for disagreement, e.g. disagreement along the lines of "not sure this will work" or "this goes against received wisdom" at relatively high score means, and disagreements along the lines of "not sure what this is about" or "hasn't this been tried before?" at lower score means. Qualitative research asking reviewers about their scoring behaviour could help understand the reasoning that goes with different scoring.

This is a good point. "Disagreement" is likely to be too general a word, and future studies could look in far more detail at the types of disagreement. We have previously used a qualitative researcher as an observer of funding panel dynamics, and we have now added this potential approach to our discussion.

Secondly, we could reframe the question to address the lack of information on unfunded applications by asking how can funding best be allocated if the amount of funding available is cut by 50%.

To test this, we reanalysed the data provided by the authors with the following question: assuming only half the funding was available, which of the following selection methods would perform better? Rank proposals based on their mean scores until funding runs out.

Pool proposals based on their mean score into buckets, rounding to the nearest integer. Fund all proposals with a mean of 1 (1.0-1.49, n=45), then from the bucket with a mean of 2 (1.5-2.49, n=114) rank proposals based on their standard deviation and fund until funding runs out. Under both methods, all proposals with a mean of 1 are funded, for a total TRC of 319.9. When we look at the proposals in the second bucket, when funding according to method 1, the total TRC for this bucket is 224.2, whereas for method 2 it is 195.5. However, the highest citation-receiving proposal in the second bucket under method 1 has a TRC of 19.9, whereas the highest citation-receiving proposal from the second bucket under method 2 has a TRC of 35.9 (the third highest TRC in the entire sample). This fits with Gillies' description of peer-review as going for less risky proposals, but at the cost of throwing out the occasional exceptional proposal. This is also born out by the distribution of TRCs, with the standard deviation of TRCs for the second bucket under method 1 being 4.2, and under method 2 being 6.2.

This is an interesting comment and thanks for the new analysis. As an aside, we note that only by sharing our data can we have such an in-depth discussion with our reviewers. We have added R code to run these suggested analyses to our results on github (

https://github.com/agbarnett/funding.disagree).

It is an interesting approach to use the mean for the "top" tier of applications and the standard deviation for the second tier. This may appeal to funders and panel members, because we believe there will always be a desire to fund those applications with the best mean. This two-tier approach may also be easier to explain than the Black–Scholes approach suggested by Linton, which combines the mean and variance in scores using an equation.

We are wary of using the standard deviation in the total relative citations because of the strong positive skew in its distribution, and the inter-quartile range in citations is actually larger for the approach using the mean at 4.3, compared with 2.9 when using the standard deviation.

Thirdly, it would be valuable to understand what sorts of disagreement exist in the scoring of all the applications (both successful and unsuccessful). It could be argued that the type of disagreement likely to indicate high risk/high return proposals would be bi-modal – some good reviews maybe with scores of 1-1.5, some very poor reviews maybe with scores of 3-4. Given the small fraction of



applications funded, very few applications of this nature would be funded (due to the understandable aggregation in the shared data we could not examine the exact number, but fewer than 10 applications have a score lower than 3). Do such applications with such bi-modal review scores exist? Or is the level of disagreement seen in the successful applications similar to that of the unsuccessful applications?

Unfortunately our data only contains the summary statistics on the applications' scores and so we can't examine these distributions. We have added this as a limitation.

The above questions only suggest that there is further complexity here that needs to be explored, with more complex, higher power studies. We thank the authors for paving the path for such studies, and for making their underlying data and analysis available in a form that is easy to explore.

Thanks, we agree that larger studies are needed.

Two small notes:

• It might be easier to read the paper if scores were consistently referred to as best/worst and better/worse – using 'higher' is confusing when better scores are lower.

Agreed and changed. We've also added text labels to Figure 5 and the new Figure 4.

• It would be more elegant to give 'e.g., Bornmann et al, 2008' rather than 'e.g., 36'. Changed as suggested.

Competing Interests: No competing interests were disclosed.

Referee Report 20 September 2018

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Jonathan Shepherd (1) 1, Jeremy C Wyatt 2

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We thank the authors for this very interesting study. We have some comments and suggestions which we think will enhance the manuscript.

- 1. The primary outcome measure is the citation counts from publications associated with the successful application. Publications were produced from 1 to 8 years after the peer review date (average 4.3 years). This does not appear to take account of varying time since the projects were funded (i.e. 7 year gap between projects that were funded from 1999 to 2006). Thus, older studies would have had more time for publications to be produced and cited. We therefore suggest a more meaningful outcome measure would be either the number of citations per year per study, or the total number of citations in, say, the 5 years following the final project report, or some other standardised project milestone. Adding the review year to the model does not seem to adequately control for this factor (though qualified statistical advice is needed to clarify this). We think this is the issue that likely affects the interpretation of the results the most in our critique.
- 2. The number of citations was standardised by academic field i.e. molecular biology. Was there any variation in study designs within this field that might also change the expected number of citations (e.g. systematic reviews may attract more citations than primary experimental studies in some fields)? It would be useful for context if there could be a table with some basic aggregate details of the funded studies, such as types of study design, molecular biological application, study sample characteristics, duration of



study etc. This would help to put the results into context.

- 3. The impact of a piece of research on which referees could not agree might be either lower or higher than those on which they could. So, it would be useful to plot the standard deviation of the citations against the standard deviation of the peer review score.
- 4. As well as using multiple imputation to correct for missing data, a sensitivity analysis in which cases with missing data are omitted would be useful.
- 5. Some measure of the model fit would be useful, eg. adjusted R squared
- 6. There was a wide variation in the number of reviewers per article from 2 to 18. Was this due to differences in the kind of research, amount of funding requested or some other perceived risk on behalf of the funder? Could this artificially influence the standard deviation of the score, confounding any association with citations?
- 7. Fractional polynomial model results are only presented for the best fitting model with the smallest deviance. This is acceptable in principle, but it would be useful for the authors to comment on whether there was any variation in the results according to the other fractional polynomial transformations (if available). This will provide confidence in the robustness of the findings.
- 8. Reference is made in the first paragraph to a "recent systematic review" by Guthrie et al (2018¹) (and also in the third paragraph). We note that this publication doesn't refer to itself as being a systematic review, and indeed, it is an update of a 2009 review which describes itself as a non-systematic review. We would suggest using the tern "non-systematic review", or just "literature review".
- 9. Thank you for citing our own recent systematic review on peer review of grants in health. You mention that the review included eight studies and called for further research in this area, which is correct. However, the review focused specifically on studies aiming to improve the effectiveness and efficiency of peer review. These were drawn from a wider set of 83 studies on peer review which we systematically mapped. In the map there were some studies which focused on assessing the impact of funded research, eg. In terms of bibliometrics. Thus, there is a body of evidence on this topic, though we didn't systematically review it in detail. We are happy to provide you with a list of these studies.
- 10. The sentence on page 3 beginning "A recent systematic review found "suggestive" evidence that funding peer review can have an anti-innovation bias² and that innovation and risk may not often be sufficiently addressed in review feedback⁷" needs re-wording as not only is the Guthrie et al paper a non-systematic review, but the second reference cited in that sentence by Gallo et al is a survey (i.e. not a review at all). The way the sentence is phrased implies that it is a systematic review.
- 11. Suggest amending the sentence on page 3 "Many studies using large sample sizes found either no association or only a weak association between the mean score and the VOLUME of citations of subsequent publications"

References

1. Guthrie S, Ghiga I, Wooding S: What do we know about grant peer review in the health sciences?. *F1000Res*. 2017; **6**: 1335 PubMed Abstract | Publisher Full Text

Is the work clearly and accurately presented and does it cite the current literature?



Yes

Is the study design appropriate and is the work technically sound? Partly

Are sufficient details of methods and analysis provided to allow replication by others? Yes

If applicable, is the statistical analysis and its interpretation appropriate? Partly

Are all the source data underlying the results available to ensure full reproducibility? Yes

Are the conclusions drawn adequately supported by the results? Yes

Competing Interests: No competing interests were disclosed.

We have read this submission. We believe that we have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Author Response 10 Oct 2018

Adrian Barnett, Queensland University of Technology, Australia

- 1. The primary outcome measure is the citation counts from publications associated with the successful application. Publications were produced from 1 to 8 years after the peer review date (average 4.3 years). This does not appear to take account of varying time since the projects were funded (i.e. 7 year gap between projects that were funded from 1999 to 2006). Thus, older studies would have had more time for publications to be produced and cited. We therefore suggest a more meaningful outcome measure would be either the number of citations per year per study, or the total number of citations in, say, the 5 years following the final project report, or some other standardized project milestone. Adding the review year to the model does not seem to adequately control for this factor (though qualified statistical advice is needed to clarify this). We think this is the issue that likely affects the interpretation of the results the most in our critique. Thanks for the comment. Actually, the citation values are normalized per publication year, thus older studies would not have an advantage as the resultant citations from publications are normalized by year. It is true that more time post peer review date would allow more publications, but all of these products were derived from the final reports, which were submitted on average five years after peer review. Thus, whatever is reported by the principal investigator in their final report is what was measured, which is what is presented by funding agencies as the output of the grant.
- 2. The number of citations was standardised by academic field i.e. molecular biology. Was there any variation in study designs within this field that might also change the expected number of citations (e.g. systematic reviews may attract more citations than primary experimental studies in some fields)? It would be useful for context if there could be a table with some basic aggregate details of the funded studies, such as types of study design, molecular biological application, study sample characteristics, duration of study etc. This would help to put the results into context. This is interesting, but would likely need to be a much larger, separate study to go back into each



application and characterize the study design, sample characteristics, etc., and is out of the current scope. That said, the applications included in this analysis were submitted to a 4-year, R01-style support mechanism. In every program year there was a significant proportion of both applied and basic research applications, with many applications encompassing varying degrees of both basic and applied research in their aims. There was a wide variety of research topic areas, including vision, drug abuse, nutrition, blood-related cancer, kidney disease, autoimmune diseases, malaria, tuberculosis, osteoporosis, arthritis, and autism research, among others. This is described in more detail in our original publication:

https://journals.plos.org/plosone/article?id=10.1371/journal.pone.0106474.

3. The impact of a piece of research on which referees could not agree might be either lower or higher than those on which they could. So, it would be useful to plot the standard deviation of the citations against the standard deviation of the peer review score.

We have now plotted the inter-quartile range in citations against the application score mean and standard deviation. We used the inter-quartile range instead of the standard deviation because of the strong skew in citations, and the standard deviation was strongly influenced by the application with the highest citations.

The new results show moderate support for a greater return for applications with a higher agreement between peer reviewers. See new Figure 4, with new explanatory paragraph in the "Statistical Methods" section, and an updated discussion.

4. As well as using multiple imputation to correct for missing data, a sensitivity analysis in which cases with missing data are omitted would be useful.

We have added a complete case analysis to the more detailed results available on github (https://github.com/agbarnett/funding.disagree). All the results were very similar to the imputed data.

- 5. Some measure of the model fit would be useful, eg. adjusted R squared We are not certain that the R-squared statistic would be useful here. The adjusted R-squared can be useful for comparing between alternative models, or where the goal is accurate prediction. Here we were just looking for any signal between application scores and citations, assuming that the detection of any signal could lead to improvements in the design of funding systems. We have investigated the residuals to look for poor model fit.
- 6. There was a wide variation in the number of reviewers per article from 2 to 18. Was this due to differences in the kind of research, amount of funding requested or some other perceived risk on behalf of the funder? Could this artificially influence the standard deviation of the score, confounding any association with citations?

The majority (90%) of applications were reviewed in panels (roughly 10 to 15 reviewers). Some were reviewed via a mail review mechanism (2 to 3 reviewers). The determination of review mechanism largely depended on the topic area. We don't believe this should be a large confounder of the results.

We have now included a sensitivity analysis using just the larger panels and excluding the mail review panels. The results were similar to the analysis using all panels. These new results are available on github (https://github.com/agbarnett/funding.disagree).

7. Fractional polynomial model results are only presented for the best fitting model with the smallest deviance. This is acceptable in principle, but it would be useful for the authors to comment on whether there was any variation in the results according to the other fractional polynomial



transformations (if available). This will provide confidence in the robustness of the findings. We have examined the predictions for the best five models and the average results were similar in terms of the mean. However, the confidence intervals for the second best model had a much narrow confidence interval for citation predictions using the application score standard deviation. In hindsight we prefer this model, as it has only a slightly poorer deviance, but a much more believable confidence interval. It would have been better to have a multi-criteria decision for the "best" model, that used both the deviance and average confidence interval width. This has been a useful lesson for future studies, but we continue to present the model with the wider CI in the main results because we feel we should stick to our original protocol.

These new results are available on github (https://github.com/agbarnett/funding.disagree).

8. Reference is made in the first paragraph to a "recent systematic review" by Guthrie et al (2018 1) (and also in the third paragraph). We note that this publication doesn't refer to itself as being a systematic review, and indeed, it is an update of a 2009 review which describes itself as a non-systematic review. We would suggest using the term "non-systematic review", or just "literature review".

Thank you for flagging this. We have now used "literature review".

9. Thank you for citing our own recent systematic review on peer review of grants in health. You mention that the review included eight studies and called for further research in this area, which is correct. However, the review focused specifically on studies aiming to improve the effectiveness and efficiency of peer review. These were drawn from a wider set of 83 studies on peer review which we systematically mapped. In the map there were some studies which focused on assessing the impact of funded research, eg. In terms of bibliometrics. Thus, there is a body of evidence on this topic, though we didn't systematically review it in detail. We are happy to provide you with a list of these studies.

We have changed our language to make the goal of your review clearer. We would gladly see the study list. Perhaps this is best uploaded to our github page for this project (if it can be made public): https://github.com/agbarnett/funding.disagree then click "upload file", you will need a github account (free) and request write access to our project.

10. The sentence on page 3 beginning "A recent systematic review found "suggestive" evidence that funding peer review can have an anti-innovation bias2 and that innovation and risk may not often be sufficiently addressed in review feedback7" needs re-wording as not only is the Guthrie et al paper a non-systematic review, but the second reference cited in that sentence by Gallo et al is a survey (i.e. not a review at all). The way the sentence is phrased implies that it is a systematic review.

Thank you for flagging this. We have now used "literature review" and added the fact that the second study is a survey.

11. Suggest amending the sentence on page 3 "Many studies using large sample sizes found either no association or only a weak association between the mean score and the VOLUME of citations of subsequent publications"

We have changed the text to read "number of citations".

Competing Interests: No competing interests were disclosed.



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