

# 10 big questions to dissect science papers

1. What is the central point the authors are trying to prove? Read carefully to determine whether the authors actually get there or fall short and/or over-interpret results.
2. Does it matter how often an experiment is repeated? Generally if you get a specific result once, it could be a mistake. If you get the same result twice, it could be a coincidence. If you get the same result three times, then maybe you have an interesting story.
3. Are there any controls? An experiment is usually only as good as its controls, which should address multiple aspects of the experiment including sensitivity of assays used, what a “positive” result should look like, and what a “negative” result should look like. This allows an experiment to be evaluated by comparing experimental findings to multiple standards.
4. Do they use the best methods available? This brings up several related questions. Check whether they used methods that are qualitative (i.e., is something there or not) versus quantitative (i.e., if it’s there, how abundant is it?). Consider whether they use rigorous and complementary approaches such as cellular lineage-tracing, clonal analysis, and/or functional assays. Do they overly rely on a simple phenotype (i.e., something sort of “looks” like “X,” so they call it “X”)?
5. Always ask yourself, “What experiment didn’t they do that could make the data more convincing?” For example, if a paper shows that a given cell expresses muscle-specific genes that look like muscle cells under the microscope, and that when transplanted into a mouse’s damaged muscle integrate and survive, what additional experiment would better convince you that they are in fact bona fide muscle?
6. Did they perform enrichment (i.e., it is mostly this) vs. purification (i.e., it is only this) vs. no idea what they used? If they knock-out/down a gene, is there also a knock-out model? Do they agree or disagree? One would hope that they

would produce similar effects. If so, what phenotype does it have? Is the result in an animal model the same as for the cognate human condition? If no, it may actually not be a very useful model of the human disease.

7. Did they increase confidence in the outcome by proving their results in multiple ways? Did they take the time to try to disprove their findings?
8. Is the paper “descriptive/hypothesis generating” or is it “hypothesis-driven” with multiple examples to back up the bottom line?
9. What do other articles say about the topic? If the paper is far afield from what others have published, are its big claims backed up by convincing data? How good is the bibliography? A good research paper should not cite too many review or opinion papers over peer-reviewed, primary research articles from high-impact journals.
10. Have the data been reproduced by other labs? Reproducibility is a key feature of good science and with a brand new finding, only time will tell if others are able to get it to work the same way.