

Someone who should have won a Nobel Prize

Martin Fenner, Gobbledygook

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The 2008 Nobel Prizes will be announced next week, starting with the Nobel Prize in Physiology or Medicine on Monday. There will be a live webcast on Monday at 9:30 AM GMT for those interested¹. As every year just before the announcement, speculation about this year's winners is in full swing. M. William Lensch, here on Nature Network correctly predicted last year's winners², and this year he is trying it again³. Thomson Reuters uses scientific methods for their predictions (including citation counts over 30 years and other awards), and they nominate⁴ these three discoveries:

- Toll-like receptors and innate immunity
- the role of microRNAs in gene regulation
- the development of meta-analysis and application to clinical medicine

There will be of course many more blog posts about the Nobel Prizes this week, both on Nature Network⁵ and elsewhere. But in this blog post I want to talk about someone who will not win a Nobel Prize, but who would very much have deserved to do so. I am talking about Judah Folkman who invented the field of angiogenesis research and worked on it for more than 35 years. But the prize can only be awarded to living people, and Judah Folkman passed away this January. A Nature obituary⁶ and recent PNAS article⁷ summarize his life and scientific achievements. Here I want to explain my personal reasons why I think he would have deserved a Nobel Prize.

Angiogenesis research is hypothesis-driven

Folkman first published his concept of tumor angiogenesis in 1971⁸. He postulated that the recruitment of dedicated blood vessels is essential to tumor growth. It took 25 years of experimental work by Folkman and others until the concept of angiogenesis became widely accepted. Interestingly, another hypothesis that changed our understanding of cancer biology was also published that year: Knudson's two-hit hypothesis of tumor suppressor genes⁹. Judah Folkman often said Science goes where you imagine it. A lot of today's research is not at all about proving an important hypothesis, and some people even mistakenly propose that concepts are no longer needed¹⁰.

Angiogenesis research is full of failures

The story of angiogenesis research is a very complicated one. Not only took it Judah Folkman a long time and a lot of stubbornness before his views became accepted, but there were many failures and setbacks along the way. Some scientific breakthroughs were an instant success, e.g. RNA interference awarded a Nobel Prize in 2006¹¹, but most research is just very complicated, including medical research. A September Science article about medical interventions found a median interval of 24 years between first description and earliest highly cited article¹².

Angiogenesis research has had an impact

The number of Pubmed citations is one good indicator for the research activity in a given field. Pubmed today lists 37,482 publications about angiogenesis, including 19985 publications from the last five years. This compares favorably to research about telomerase (8040 publications), and microRNA (3541 publications), two areas of research mentioned in the discussions about potential 2008 winners. Impact can be also measured in other ways, but it is clear that the concept of angiogenesis has not only profoundly changed cancer research, but is also important in many other diseases, summarized in a 2005 Nature article¹³.

Angiogenesis research is medical research

As the name already suggests, the Nobel Prize for Physiology or Medicine is actually two prizes in one. It is a Nobel Prize in Biology, awarded for major achievements in our understanding of all aspects of biology. But is also a Nobel Prize in Medicine, where major advances in our understanding of human disease are awarded. The 2005 Nobel Prize to Barry Marshall and Robin Warren for the role of *Helicobacter pylori* in gastric disease¹⁴ is a good recent example of the latter category.

The research by Folkman and others has changed the way we think about cancer, but also the way we treat cancer. Each year, more than one million patients with colon, lung or breast cancer are treated with the anti-VEGF antibody bevacizumab. Other drugs targeting angiogenesis are either already in clinical use (e.g. sunitinib for renal cancer) or in development. This makes angiogenesis research a prime example for translational research. I very much agree with the views expressed in a set of articles in Nature this June¹⁵, that more has to be done to connect our much better understanding of fundamental biological processes to the way we diagnose, prevent and treat diseases. And a Nobel Prize would be a strong signal.

fn1. Announcement of the Nobel Prize in Physiology or Medicine

fn2. Nobel Redux

fn3. Nobels 2008

fn4. 2008 Nobel Predictions

- fn5. Collective Blogging – Nobel Prize announcements
- fn6. Nature 2008 doi:10.1038/451781a
- fn7. PNAS 2008 doi:10.1073/pnas.0806582105
- fn8. N Engl J Med 1971 Tumor angiogenesis: therapeutic implications
- fn9. PNAS 1971 Mutation and Cancer: Statistical Study of Retinoblastoma
- fn10. The End of Theory: The Data Deluge Makes the Scientific Method Obsolete
- fn11. Nobel Prize in Physiology or Medicine 2006
- fn12. Science 2008 doi:10.1126/science.1160622
- fn13. Nature 2005 doi:10.1038/nature04478
- fn14. Nobel Prize in Physiology or Medicine 2005
- fn15. Nature 2008 doi:10.1038/453839a