

some clear positions. For example, it suggests roughly doubling average fiber intake, to 38 grams per day for men and 25 grams for women.

Certain recommendations—such as urging every adult to get an hour's daily exercise, twice the amount recommended in the past—seem to ignore the real-life lifestyles of North Americans. "I couldn't possibly do an hour of exercise a day," says Marion Nestle, chair of the department of nutrition and food studies at New York University. Nestle, who was not on the panel, complains that the report is too complex for "an already confused public." What causes obesity "isn't rocket science ... eating too much [does]."

Indeed, panel member Ronald Krauss, who studies diet and heart disease at Lawrence Berkeley National Laboratory in California, agrees that the report might not respond to the question people generally ask: "How much of this should I eat?" Unfortunately, science isn't able to deliver such detailed diet advice quite yet.

—JENNIFER COUZIN

## UNDERGRADUATE SCIENCE

### Biology Departments Urged to Bone Up

In 1998, Sheldon Wettack, a dean at Harvey Mudd College in Claremont, California, decided that undergraduates needed a better appreciation of the connections between biology and the physical sciences. He and a few faculty members devised a program, called the Interdisciplinary Laboratory, that parallels intro chemistry and physics classes and includes such exercises as how thermodynamics affects animal design. Wettack hoped that the new team-taught lab would strengthen the biology curriculum and maybe even attract majors from the more quantitative sciences into biology.

This kind of approach is exactly what's needed to train the next generation of biomedical researchers, says a new report by the National Research Council (NRC).<sup>\*</sup> "Biological research is already highly interdisciplinary, but undergradu-

ate education is not," says panel chair Lubert Stryer of Stanford University in Palo Alto, California. "And the gap is increasing."

The NRC panel, funded by the National Institutes of Health and the Howard Hughes Medical Institute (HHMI), found that undergraduate biology education also needs a more rigorous curriculum. Many of the recommended changes are longtime favorites of science education reformers (*Science*, 31 August 2001, p. 1607), including thought-provoking lab exercises and independent research projects. To improve quantitative skills, faculty members should include more concepts from math and physical sciences in biology classes. Ideally, the report says, the entire curriculum would be revamped to add more heft.

But these changes face many obstacles, including the expense of developing new course materials and the conservative influence of the Medical College Admission Test (MCAT), a national qualifying exam for would-be U.S. medical students. "It is time that the curriculum started driving the MCAT, not the other way around," says David Hillis of the University of Texas, Austin. When it comes to curricula, there is also a massive amount of inertia in higher education, says Peter Bruns, vice president for grants and special programs at HHMI. "People say the only institution more conservative is the [Catholic] Church."

Even when reform is on the agenda, it's hard for departments to agree on how to car-



**Now hear this.** New report emphasizes hands-on activities over lectures for undergraduates.

ry it out. "If you add something, you have to take something away," says Hillis, and no one wants his or her subject trimmed. Personal foibles can play a role, too: A professor who agrees that number-crunching skills would be useful might still be loath to admit ignorance. "Most faculty have trouble saying, 'I don't know much about this topic, but you should,'" says chemist Ronald Breslow of

Columbia University in New York City.

One way to achieve change is by sweetening the pot. Toward that end, next week HHMI will award \$1 million over 4 years to each of 20 senior faculty members who have proposed ways to improve undergraduate biology education at their institutions. The idea is to provide role models as well as the necessary resources. That approach makes good sense to Stryer, who says that energetic leadership is a key ingredient in making the panel's recommendations a reality.

—ERIK STOKSTAD

## INFORMATICS

### The Genome Chose Its Alphabet With Care

Of all the nucleotide bases available, why did nature pick the four we know as A, T, G, and C for the genomic alphabet? Researchers have long put it down to the composition of the primordial soup in which the first life arose. But Dónall Mac Dónaill of Trinity College Dublin says the answer is much more interesting. He believes that the choice of A, T, G, and C incorporates a tactic for minimizing the occurrence of errors in the pairing of bases, in the same way that error-coding systems are incorporated into ISBNs on books, credit card numbers, bank accounts, and airline tickets. "The answer may lie partly in the error-coding aspects of information transfer," he says.

There are 16 possible nucleotide bases that could pair up to make DNA, and researchers have created strands of synthetic DNA using all the combinations. Informatics might be the key to why nature ignored all but four of these possibilities, Mac Dónaill suspected, and he built on the structural work of biologist Eörs Szathmáry of the Collegium Budapest in Hungary to test his hunch.

In the error-coding theory first developed in 1950 by Bell Telephone Laboratories researcher Richard Hamming, a so-called parity bit is added to the end of digital numbers to make the digits add up to an even number. For example, when transmitting the number 100110, you would add an extra 1 onto the end (100110,1), and the number 100001 would have a zero added (100001,0). The most likely transmission error is a single digit changed from 1 to 0 or vice versa. Such a change would cause the sum of the digits to be odd, and the recipient of that number can assume that it was incorrectly transmitted.

Mac Dónaill asserts, in a forthcoming issue of *Chemical Communications*, that a similar process was at work in the choice of bases in the genetic alphabet. First he represented each nucleotide as a four-digit binary number. The first three digits represent the

<sup>\*</sup> *BIO2010: Undergraduate Education to Prepare Biomedical Research Scientists* (National Academy Press, 2002).

three bonding sites that each nucleotide presents to its partner. Each site is either a hydrogen donor or acceptor; a nucleotide offering donor-acceptor-acceptor sites would be represented as 100 and would bond only with an acceptor-donor-donor nucleotide, or 011. The fourth digit is 1 if the nucleotide is a single-ringed pyrimidine type and 0 if it is

a double-ringed purine type. Nucleotides readily bond with members of the other type.

Mac Dónaill noticed that the final digit acted as a parity bit: The four digits of A, T, G, and C all add up to an even number. Nature restricted its choice to nucleotides of even parity, says Mac Dónaill, because “alphabets composed of nucleotides of mixed parity would have catastrophic error rates.” For example,

nucleotide C (100,1) binds naturally to nucleotide G (011,0), but it might accidentally bind to the odd parity nucleotide X (010,0), because there is just one mismatch. Such a bond would be weak compared to C–G but not impossible. However, C is highly unlikely to bond to any other even-parity nucleotides, such as the idealized amino-adenine (101,0), because there are two mismatches. So, nature has avoided such mistakes by banishing all odd-parity nucleotides from the DNA alphabet.

Computational chemist Graham Richards of Oxford University thinks the finding is important: “It is a novel idea which should provoke others to explore aspects of informatics in the genetic code,” he says, adding: “Instinctively, one feels that the DNA code should have evolved systems to minimize errors. Mac Dónaill’s work shows how this could have been achieved.” Larry Liebovitch of Florida Atlantic University, Boca Raton, agrees. “Mac Dónaill’s clever analysis shows how well different nucleotides could serve as matches in DNA and how much different pairs differ from each other,” he says. “This analysis gives us a reason to believe that the A–T and G–C choice forms the best pairs that are the most different from each other, so that their ubiquitous use in living things represents an efficient

and successful choice rather than an accident of evolution.”

—DAVID BRADLEY

David Bradley is a writer in Cambridge, U.K.

## PATENT LAWS

### Report Urges Leeway For Developing World

An independent commission appointed by the British government is advocating weaker intellectual property (IP) laws in developing countries in hopes of fostering innovation. The report, released this week, also criticizes the World Trade Organization’s agreement on Trade-Related Aspects of Intellectual Property Rights (TRIPS), which in 1995 delineated minimum global standards that nations should achieve.

“We’ve recommended that patents aren’t necessarily a good idea for many developing countries,” says Charles Clift, a London economist and head of the secretariat that managed the six-member commission. Commissioners believe that developing countries can ill afford to pay fees on patented inventions, both domestic and foreign, that would help them expand their technological base. Devinder Sharma, an analyst at the Forum for Biotechnology and Food Security in New Delhi, India, says he’s pleased that “for the first time, a high-powered body is saying, ‘Go slow on patents.’”

The U.K. government launched the commission after a review of the problems associated with globalization. Its conclusions ([www.iprcommission.org](http://www.iprcommission.org)) reinforce arguments made for years by nongovernmental organizations. But some say a position that favors narrower patents is misguided. “A policy saying you should not grant patents on things that are really, truly inventive makes no sense for developing countries,” says Jeffrey Kushan, a patent attorney at Sidley Austin Brown & Wood LLP in Washington, D.C. “At the practical level, a lot of these issues don’t have any data.”

The commissioners acknowledge that a dearth of IP regulations in the least developed countries forced them to guess the impact of U.S. or European-type IP laws on a country’s technological development. After consulting with institutions in nine developed and developing countries, however, they concluded that many developing nations ought to avoid issuing patents that, for example, allow the kind of broad protection

available in the United States because it would stifle additional innovation. The commissioners also encouraged developing nations to adopt broad exemptions for educational and research use of patented materials. The group agreed, too, that a single IP system cannot serve the diversity of developing countries. Commission member Raghunath Anant Mashelkar, a polymer engineer and director-general of India’s Council of Scientific and Industrial Research, says the report’s message to authorities is “Don’t force-feed stringent IP [rights] laws to poor countries that do not have the inherent capacity to implement them.”

The commissioners were also concerned about the potential high cost of licensing a patent. For example, developing countries are generally not permitted to sell cheap generics of drugs still under patent protection (AIDS drugs being a notable exception). Stanford University law professor John Barton, who chaired the commission, suggests that such fees might prevent widespread use of an invention.

Kushan dismisses that argument, saying that “if a company wants to make money in a market, it’s going to adjust its license fees.” He also says that raising IP standards has historically promoted international competition and investment, citing reforms in Brazil in the late 1990s that preceded a \$2 billion infusion from U.S. drug companies. Lila Feisee, IP director at BIO, an industry group in Washington, D.C., agrees.

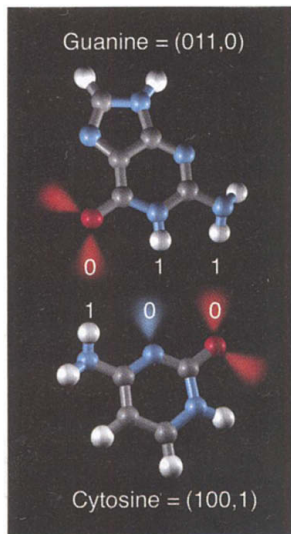


**Patent potpourri.** More lenient patent laws allow Indian biotech companies to produce generic AIDS drugs.

“Obviously, every country’s got its own special criteria,” she says. Still, “we push very hard to try and harmonize the patent laws of different countries.”

The report will be presented next week to officials from the World Trade Organization and the World Intellectual Property Organization. The commissioners hope that their ideas will be incorporated into TRIPS, which continues to undergo revisions, or other international agreements.

—JENNIFER COUZIN AND PALLAVA BAGLA



**Binary bases.** Representing nucleotides as binary numbers reveals how they were chosen to avoid errors.