The behind-the-scenes struggle to replicate ballyhooed cancer results



Interview with China's science minister

ning Margaret Hamburg said last week, is to build connections between emergency crews who first respond to a crisis and agencies like CDC that have expertise in exotic diseases. Most doctors today have never seen a smallpox or anthrax case, so the network will serve in part to educate physicians about recognizing and treating these diseases. It will also provide a secure communications link in a crisis.

HHS is planning to spend \$51 million on a stockpile of drugs and vaccines, says James LeDuc of CDC's National Center for Infectious Disease. The CDC's two top concerns are anthrax, a tough bacillus, and the variola virus that causes smallpox. Anthrax is treatable with antibiotics if detected quickly, but it's hard to spot an infection early, and it kills quickly. Anthrax usually is not communicated from person to person. Smallpox, on the other hand, is highly contagious, and would cut a devastating swath through unvaccinated urban populations.

CDC has about 15 million doses of smallpox vaccine in its 20-year-old reserve, but because the rubber seals are deteriorating, about a quarter are suspect. This vaccine should be replaced, preferably with a new type, LeDuc says, produced with modern techniques. (The existing vaccine is neither sterile nor pure.) But establishing the safety and efficacy of a new vaccine could be difficult, because there are no smallpox patients to test it on. CDC is talking to the National Institutes of Health (NIH) and USAMRIID about developing animal models for testing a new formula.

Researchers are also debating the wisdom of a plan to destroy the last stocks of small-pox virus, at CDC and at the Research Institute for Viral Preparations in Moscow. Henderson and some other researchers argue that the destruction, scheduled for June, would reduce the risk of a smallpox attack by keeping the virus out of terrorist hands. But some argue that stocks should be preserved to help develop new drugs and vaccines (*Science*, 19 November 1993, p. 1223 and 1225.)

For anthrax, there are no civilian vaccine stocks at all: Supplies have been purchased by the Defense Department for the troops, and the sole factory that makes the vaccine is shut for renovation. CDC officials agree that it will be necessary to develop a new anthrax vaccine soon. USAMRIID has candidates in development, but bringing them through a series of clinical trials will be costly. A new version may be ready by 2005. In the meantime,

CDC will store up antibiotics.

Another chunk of money, about \$25 million, will go to NIH for basic science supporting vaccine and drug research. The bulk of it will be channeled through NIH's National Institute of Allergy and Infectious Diseases (NIAID) to extramural grantees for genetic studies of pathogens (anthrax, smallpox, plague, and tularemia). Starting next year, according to NIAID's Catherine Loughlin, "We'd like to take advantage of the genomic information to identify targets" for drug and vaccine development.

One promising therapeutic development, according to NIAID staffer Bernard Moss, comes from an agency that isn't earmarked for a funding boost-USAMRIID. There, microbiologist John Huggins has been screening licensed antiviral drugs to find some that might help combat smallpox. Using a mouse model of smallpox he developed, Huggins found a good candidate: cidofovir, a drug used mostly by AIDS patients for cytomegalovirus eye infections. But it must be given intravenously, and it has strong side effects-problems that make it impractical for emergency use. Henderson, for one, sees no immediate application. But Huggins is collaborating with NIAID and CDC in a search for analog drugs, says Loughlin, although Huggins is "doing all the work."

While USAMRIID is pleased to have collaborators in its traditional line of research, long-time workers in the field wonder how long the enthusiasm will last. Mindful of the ephemeral quality of such policy initiatives, reporters asked Shalala last week at what point the government's antibioterror program would reach its objectives. Shalala shot back: "This is not a quick response. ... I will never say we have done enough." **–ELIOT MARSHALL** 

#### RADIOACTIVE WASTE

### Yucca Mountain Panel Says DOE Lacks Data

With just 2 years to go before deciding whether Yucca Mountain in southern Nevada should be a permanent home for spent fuel from the country's nuclear power plants, the U.S. Department of Energy (DOE) has run into another snag. In a report\* submitted 2 weeks ago, a panel of experts says major questions about the controversial site are still

unanswered and casts doubt on DOE's ability to make a final decision in 2001.

Congress chose Yucca Mountain as the sole site to be studied as a high-level radioactive waste repository in 1987, and DOE has spent \$6 billion toward reaching that goal. While waste piles up in 72 temporary facilities, political and legal battles have pushed back its original start-up date of 1998 to the current target of 2010. In December, the department announced that the latest study, an assessment of the remote mountain's ability to entomb the waste safely for thousands of years, had identified "no show stoppers." Although safety questions remain, DOE officials said then, they were confident that the repository "would protect public health and the environment for thousands of years."



A long haul. Panel wants more testing at Yucca Mountain and in the lab.

But on 11 February, a blue-ribbon panel of six experts hired to peer review the agency's study raised doubts about that conclusion in what the panel calls a "highly critical" report. "There's a lot to be done" before DOE can make such a prediction, says panel chair Chris Whipple, a risk assessment engineer at ICF Kaiser Engineers Inc., in Oakland, California. "Can they do it on their current schedule? That seems unlikely."

DOE is taking the panel's report in stride. "I think they overstated [the uncertainties] a bit, [but] it's what we paid for," says Abe Van Luik, senior technical adviser for performance assessment in the Yucca Mountain Project. "We're taking them seriously." DOE is still aiming for a decision in 2001, he says.

The report faults the department's current model for predicting the repository's behavior, which takes into account everything affecting the movement of radioactive elements out of

<sup>\* &</sup>quot;Final Report, Total System Performance Assessment Peer Review Panel." For a copy, see www.ymp.gov

the fuel rods and into the distant environment over millennia. The panel agrees that DOE has done a good job assessing such possible disruptions as earthquakes, volcanic eruptions, and nuclear reactions suddenly taking off, but notes that other assumptions "may be unduly optimistic." For example, the cladding that encases the enriched uranium rods and provides the first line of defense may not hold up as well as assumed. More lab work on the cladding's behavior under repository conditions is needed, says the report.

The behavior of the radioactive material once it leaks out, as it eventually must, is also unclear, says the report. More exploratory holes should be drilled into aquifers far from Yucca Mountain, where the radioactivity will ultimately spread, it suggests. The panel is especially concerned about the assumptions behind the repository's "hot" design, in which heat from the waste is supposed to keep temperatures well above boiling and thus initially keep out moisture that could corrode the rods. "We don't think anybody can model that convincingly," says Whipple. Such stubborn problems might be handled by making some conservative, simplifying assumptions, says Whipple, an approach DOE has yet to accept.

Van Luik says he's "a tad surprised at the amount of material they think we need to do." Some of the suggested work is already under way, he notes, and project staff are still debating the merits of a hot design. "This is not our final design, nor our final understanding of the site," he explains. But he's concerned by the fact that "the panel recommends that we do additional work that would extend us beyond our current schedule."

Kevin Crowley, staff director of the National Research Council's Board of Radioactive Waste Management in Washington, D.C., says DOE would be wise to take the panel's advice because its current schedule is unrealistic. The panel's emphasis on gathering more data and dealing with the intractable complexities, he adds, could be key to resolving the technical issues. "The DOE has some real challenges ahead," he warns.

-RICHARD A. KERR

#### MATERIALS

# Stress Profiling Gets The Best Out of Glass

Try to bend a piece of window glass, and you'll get a vivid demonstration of glass's brittle behavior. When stressed, it shatters without warning into thousands of shards. Now an international team of researchers has developed a clever way to make glass a little more pliant and predictable. On page 1295, materials scientist David Green of Pennsylvania State University in University Park and his team describe a chemical

toughening process that resulted in glass that both resists fracture better and delivers a warning before it finally fails, in the form of small cracks on its surface.

"Usually, when a piece of glass starts to break, that's the end of the story. In this glass you can arrest the cracks and you get some warning before the final failure," says Green. "The fact that multiple cracking can be observed in glass is indeed remarkable," agrees William Tredway, advanced ceramics group manager at United Technologies Research Center in East Hartford, Connecticut.

The traditional method for making glass more resistant to fracture is called tempering. Manufacturers use either heat or chemicals to increase the "residual stress"—the compres-



Flexiglass. Cracks form on a piece of treated glass as it is flexed, but it does not break.

sive forces between atoms-at its surface. Before an external stress forms a crack, it must overcome not only the normal strength of the material, but also this extra residual stress. Tempered glass is more resistant to fracture, but when a crack does form at the surface it quickly moves deeper where the stress is lower, and the material fails catastrophically. In 1991, Green

and Rajan Tandon, now at Caterpillar Inc. Technical Center, a construction machinery manufacturer in Peoria, Illinois, did theoretical studies that pointed to a better way to strengthen glass. The studies showed that a compressive stress "profile," with relatively weak stress at the surface increasing to a maximum at a depth of 20 to 30 micrometers, would stop cracks because they would face increasing compressive stress as they moved deeper into the material. "The idea went against the current dogma of what you are supposed to do," says Green. "Usually people try to get the maximum compression at the surface."

To create the required stress profile, the researchers developed a two-stage chemical tempering process. The main skeleton structure of glass is composed of silicon and oxygen atoms, interspersed with sodium atoms. The researchers immersed a glass sample in a bath of molten potassium nitrate at high temperature, allowing some of the potassium ions in the bath to swap places with sodium ions in the glass—a process called "ion exchange." Potassium atoms have a radius that is 25% larger than sodium, says team member Vincenzo Sglavo of the Uni-

## ScienceScope

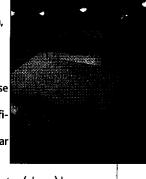
Food Fight Fed up with the ongoing media feeding frenzy surrounding genetically modified (GM) food, 19 of Britain's most eminent scientists, all Fellows of the Royal Society, have called for the use of peer review rather than public opinion to judge scientific results. Writing in the 23 February issue of London's Daily Telegraph and Guardian newspapers, they say that "it is a dangerous mistake ... to assume that all statements claiming to be scientific can be taken at face value."

The letter follows a public statement from 21 scientists in defense of protein chemist Arpad Pusztai (*Science*, 19 February, p. 1094). Last summer, Pusztai was suspended from his post at the Rowett Research Institute near Aberdeen after declaring in a TV documentary that his unpublished research indicated that potatoes genetically altered to resist pests stunted growth and suppressed immunity in rats.

One signatory of the new letter, botanist Ghillean Prance, director of the Royal Botanical Gardens at Kew, says that the Fellows are concerned about several recent incidents in which unpublished results have been promoted in the media. "So much bad science is going into the press," he says. The society is convening an expert panel to review Pusztai's results and broader issues related to GM foods.

**Meltdown** Nuclear research reactors continue to disappear from U.S. university

campuses as the field loses ground to other disciplines. Last month, Iowa State University in Ames finalized plans to dismantle by next year its 10-kilo-watt minireactor, whose \$200,000-a-year budget became an unjustifiable expense after the school closed its nuclear engineering program a few years ago.



The 40-year-old reactor (above) becomes the 43rd campus-based atomic plant to close since 1975, leaving about 30 operating academic reactors. Up to a third of the remaining plants could close within the next decade, experts say, due to operating cost concerns and relicensing expenses. Several advisory panels have called on the Department of Energy to stem the decline, which they say threatens a wide range of engineering, materials, and physics research.

Contributors: Daniel Clery and David Malakoff