

patient who had died from vCJD. For the recent study, the team collected a large number of samples of lymphoid tissues—including tonsils, spleens, and lymph nodes—from a variety of sources. These included tonsil biopsies from 20 patients suspected of having some sort of prion disease—nine of whom were later shown to be suffering from vCJD. The rest were from tonsillectomies or stored autopsy materials taken from vCJD victims, normal controls, and sufferers of other neurological diseases, including “classic” forms of CJD not linked to BSE.

In search of PrP, Collinge’s team then subjected samples to two types of laboratory test. The first, immunohistochemistry—in which a target protein is “stained” with antibodies that specifically recognize it—can tell if PrP is present but not whether it is abnormal. The team found that PrP was detectable in lymphoid tissue only among vCJD sufferers; they found no PrP in any of the other samples. The second test, known as Western blotting, detects proteins both by their molecular weight and their reactions with antibodies. Collinge has claimed that this test can distinguish different prion strains because they have different patterns of sugar residues on their surface and hence different molecular weights. The new results may support that claim: They confirmed that only vCJD sufferers had PrP in their lymphoid tissue and found that all the vCJD patients shared the same prion strain.

The study “looks very convincing,” says Oxford University epidemiologist Roy Anderson. And molecular biologist Chris Bostock, director of Britain’s Institute for Animal Health, says that the new test “looks like a promising tool, along with others, to confirm diagnosis of vCJD.” But researchers caution that use of the test for wide-scale screening raises serious ethical questions. For example, with no cure for vCJD in sight, should people be told they are harboring the disease? “The situation is analogous to the early stages of the AIDS epidemic,” says Anderson. Health officials are therefore considering an anonymous screening program, for research purposes only.

Yet some researchers say that screening tonsil tissues could give rise to misleading data on the extent of the epidemic. Moira Bruce of the Institute for Animal Health’s Neuropathogenesis Unit in Edinburgh points out that tonsils and appendixes are normally removed because they are inflamed and flooded with immune cells such as lymphocytes. “We know that expression of PrP on lymphocytes is elevated as part of the immune response,” Bruce says, so such a screening program could lead to “false positives” and overestimate

the infected population.

On the other hand, some researchers believe abnormal PrP may be undetectable in the disease’s early stages, and because nobody knows at what stage the protein moves from infected beef in the gut to lymphoid tissues, screening might underestimate the epidemic. Nevertheless, screening will be required if health officials are to know what they are up against. Says Collinge: “It would be irresponsible not to make use of this test. We might find evidence of a major problem, and we need to know sooner or later.”

—MICHAEL BALTER

DEVELOPMENT

Brain Stem Cells Show Their Potential

Brains memorize organic chemistry equations, control typing fingers, and integrate the sensory input needed to navigate snarled traffic—all feats of profound sophistication and versatility. Now the brain is proving itself even more of a renaissance organ, for some of its cells can perform the tasks of a completely different tissue.

On page 534, Angelo Vescovi, a neurobiologist at the National Neurological Institute Carlo Besta in Milan, Italy, and his colleagues report that neural stem cells, which give rise to the three main types of brain cells, can also become blood cells when transplanted into mice whose own blood-forming tissue, the bone marrow, has been mostly destroyed. It wasn’t until the early 1990s that scientists

found ways to isolate neural stem cells and grow them in the lab. “Now the brain’s making blood,” Vescovi says.

“What’s interesting is the idea that cells can shake their fates,” says Ron McKay, a neurobiologist at the National Institute of Neurological Disorders and Stroke in Bethesda, Maryland. The result provides a strong push to find other stem cell types with similar capabilities. And it opens the possibility of using neural stem cell transplants to treat human blood cell disorders such as aplastic anemia and severe combined immunodeficiency—an appealing idea, as bone marrow stem cells don’t replenish themselves well in lab cultures.

Several observations had hinted at brain cells’ versatility. During development, muscle cell types—which arise from a layer of embryonic cells distinct from that which

generates the brain—appear in the central nervous system, says Vescovi. Furthermore, scientists sometimes see muscle tissue in a particular type of brain tumor. Because no one knows where the muscle cells come from, “we theorized that maybe there’s a brain cell that possesses a much wider potential for differentiation than previously thought,” says Christopher Bjornson, a developmental biologist currently at the University of Washington, Seattle, and a co-author of the *Science* paper.

To find out, the team isolated neural stem cells from adult and embryonic mice and grew them singly in lab cultures. After irradiating mice to kill most of their bone marrow cells and create a vacancy that new cells might occupy, the researchers injected the neural stem cells into the animals. Because the donor mouse cells carried distinctive genetic markers, the researchers could trace their fate in the injected animals.

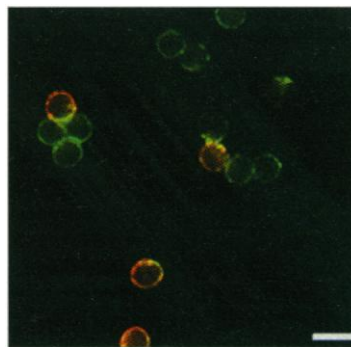
Five months later, the investigators found that the blood of the recipients contained cells that not only displayed the donor cell marker protein but also produced proteins that only mature blood cells make. They also showed that the animals’ bone marrow carried immature blood precursor cells that were descended from the neural cells.

No one knows exactly what caused the neural cells to turn into blood cells. But Vescovi and his colleagues suspect that the neural cells might be responding to the same signals that normally stimulate the few remaining blood stem cells to reproduce

and mature after irradiation wipes out most of the bone marrow. “The result suggests that there’s something quite powerful in the mature adult blood system that can instruct cells from a different origin what to do,” says Arturo Alvarez-Buylla, a neurobiologist at The Rockefeller University in Manhattan.

Whatever that is, its effects appear to be long-lasting. The Vescovi team could detect the neural-derived blood cells a year after the injection. That means that the transplants may persist long enough to be clinically useful, the researchers say. If the Vescovi team’s work can be replicated in humans, agrees Irving Weissman, a stem cell biologist at Stanford University School of Medicine, the neural cells “could become a source for blood stem cells.”

—EVELYN STRAUSS



Double duty. Yellow color identifies white blood cells produced by neural stem cells. (Bar equals 10 micrometers.)