

high risk of colon cancer, so that they could be watched carefully and treated early.

Identification of the new APC mutation was serendipitous, the result of a social visitor to Johns Hopkins, who mentioned that he had had several colorectal polyps and a slight family history of colon cancer. Vogelstein, whose lab had already uncovered a fistful of genes involved in the disease, offered to test the 39-year-old male for mutations.

Vogelstein's group did find a change in the APC gene. But at first glance it appeared to be innocuous—a simple switch from a thymine (T) to an adenine (A) at position 1307 that didn't look like it would disrupt the gene's ability to function. Such gene changes, called polymorphisms, are common.

What raised the researchers' suspicions was a strange phenomenon that occurred when they tested the patient's APC gene in a routine assay that allows it to make its protein product. They found that the protein began to pick up extra mutations in and around the region that contains the T-to-A switch. That apparently

happened, Kinzler says, because the mutation creates a stretch of eight consecutive adenines, which are often misread by polymerase enzymes that transcribe genes into messenger RNAs (mRNAs)—the first step toward making proteins. "The DNA strands can get a little one-to-two base-pair bubble," says Kinzler. "That allows the polymerase to put in an extra base without realizing there is a mistake." Such "frameshift" mutations can totally garble the rest of the message, creating shortened forms of the protein or rendering it useless.

If just mRNA synthesis were affected, the situation might not be harmful, because it wouldn't lead to permanent loss of all functional APC protein. But the same kind of error can also occur in the DNA itself during replication. The Johns Hopkins team found that this may in fact be happening in the colon cells of some patients who have an increased risk of colon cancer but don't carry the original APC mutations. When the investigators looked at the APC gene in tumor and blood samples of these patients, they found that all the tumors

that carried the second type of inactivating mutation also had the T-to-A change. Blood cells from the same patients only had the T-to-A switch, however. These results suggest that the patients inherited the base change and developed the other mutations later, but only in the colon cells that became cancerous.

The work so far has shown that the T-to-A mutation is present in about 6% of Ashkenazi Jews and in 28% of Ashkenazim with familial colon cancer. The Hopkins team and others are now expanding those studies and looking at how common the gene is in the general population. Another big question is whether other tumor-suppressor genes are prone to similar problems. "Everyone has seen polymorphisms in cancer genes," says Vogelstein. "And all of us have assumed they are just harmless variants. This study suggests that those kinds of mutations may not really be harmless, but rather a kind of wolf in sheep's clothing."

—Trisha Gura

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PHYSICS

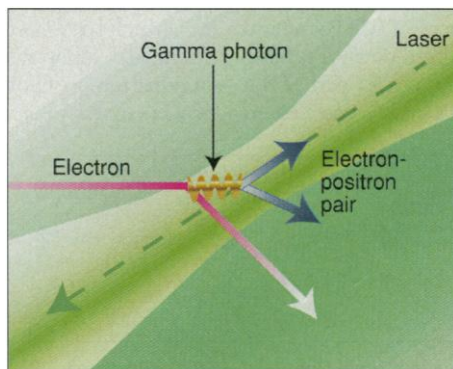
Conjuring Matter From Light

Turning matter into light, heat, and other forms of energy is nothing new, as nuclear bombs spectacularly demonstrate. Now a team of physicists at the Stanford Linear Accelerator Center (SLAC) has demonstrated the inverse process—what University of Rochester physicist Adrian Melissinos, a spokesperson for the group, calls "the first creation of matter out of light." In the 1 September *Physical Review Letters*, the researchers describe how they collided large crowds of photons together so violently that the interactions spawned particles of matter and antimatter: electrons and positrons (antielectrons).

Physicists have long known that this kind of conjuring act is possible, but they have never observed it directly. The experiment is also a proof of principle for a technology, based on intense laser beams boosted to enormous energies with the help of SLAC's electron beam, for exploring a theory known as quantum electrodynamics. QED describes electromagnetic fields, such as those of light, and their interactions with matter, and its predictions are notoriously accurate. But physicists are eager to study it at so-called "critical" electromagnetic fields—fields so strong that their energy can

be converted directly into the creation of electrons and positrons.

To create a field as close as possible to critical, the 20-physicist collaboration started with a short-pulse glass laser that packs a half-trillion watts of power into a beam measuring just 6 micrometers across at its narrowest point, resulting in extraordinary intensities. To increase the energy of the photons, the team collided the pulses with SLAC's 30-



Flash dance. An electron beam intersects a laser pulse, boosting photons to gamma energies and triggering an interaction that spawns particles.

micrometer-wide pulsed beam of high-energy electrons—a feat that required precise alignment and synchronization. When laser photons collided head-on with the electrons, they got a huge energy boost, much like ping-pong balls hitting a speeding Mack truck, changing them from visible light to very high energy gamma

rays. Because of the laser's intensity, these backscattered gamma photons sometimes encountered several incoming laser photons simultaneously; a collision with four of them concentrated enough energy in one place to produce electron-positron pairs.

Melissinos views the result as the first direct demonstration of "sparking the vacuum," a long-predicted phenomenon. In it, the en-

ergy of a very strong electromagnetic field promotes some of the fleeting, "virtual" particles that inhabit the vacuum, according to QED, to become pairs of real particles.

Electron-positron pairs are often spawned in accelerator experiments that collide other particles at high energies, and photons produced in the collision are what actually generate the pairs. But at least one of the photons involved is virtual—produced only for a brief moment in the strong electric field near a charged particle. The SLAC experiment marks the first time matter has been created entirely from ordinary photons.

Princeton University physicist Kirk McDonald, another spokesperson for the collaboration, which also includes the University of Tennessee and SLAC, thinks the high-field experiments could shed light on phenomena at the surface of a neutron star, where magnetic fields are very strong, and in other exotic astrophysical settings. On a more practical level, the conversion of light into matter could also give particle physicists a new source of positrons that are exceptionally uniform in energy and momentum.

The result is also the first step toward using powerful lasers and electron beams to test high-field QED predictions, such as what McDonald calls "vacuum optics"—the behavior of light in a strong-field environment. "We're exploring new regimes and trying to map out the basic phenomena," he says. Physicist Tom Erber of the Illinois Institute of Technology looks forward to the results: "Hopefully, this will open the door to future experiments which will approach [more probing] tests of QED."

—David Ehrenstein