UCDAVIS BIOLOGICAL SCIENCES

News from the Division of Biological Sciences

Vol. 9 No.1

BIOTECH SCHOOL'S IN FOR SUMMER

ho do you call when a new research technique debuts on the scientific stage and you want to learn it fast?



protein analysis class (above) in conjunction with the UC Davis Molecular Structure Facility," says Biotechnology Program Director Martina McGloughlin. "The students in the class are exposed to techniques that make them aware we're heading into an era during which we'll be exploring areas beyond proteomics."

During the summer, the UC Davis Biotechnology Program offers a series of intensive courses that focus on a particular biotechnology research technique. "We teach the

Hundreds of scientists, with backgrounds in academia and industry, have turned to the UC Davis Biotechnology Program.

Since 1988, the Biotechnology Program has offered two- to five-day intensive courses, open to the public, that combine lectures with laboratories designed to give participants hands-on experience with a particular research technique.

Martina McGloughlin, director of the Biotechnology Program, is aware of only a few other places in the United States where people can take classes similar to the ones her program offers.

The courses change to keep pace with and reflect developments in biotechnology.

Says McGloughlin, "In 1989 the Biotechnology Program offered its first intensive summer short course. It was a course in DNA manipulation, and we offered it because many people had gone

> through school before these new techniques were taught. We had mostly faculty in the first classes—biochemists and physiologists who were trying to catch up on the new technology."

> McGloughlin realized the initial DNA manipulation course targeted a finite audience since the techniques would now be part of a student's education. Therefore, in 1990, the program offered a DNA sequencing class. "When we offered the class, people didn't have automatic DNA sequencers," says McGloughlin.

> Once automatic sequencers supplanted the do-it-yourself method the sequencing class became obsolete.

McGloughlin mentions another example

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THE BIG PICTURE

hink big is the message **Pete Smietana**, Ph.D., Biophysics, 1979, hopes students took away from the bioinformatics courses he taught for UC Davis' Biotechnology Program.

"The amount of information currently being generated in the life sciences is mind boggling," says Smietana. "For example, in the company I'm currently with, LumiCyte, Inc., we don't run one microarray chip, think about the results for awhile, then run another. We run one microchip every day—that's about 10,000 data points a day, every day. This quantity of information needs to be in electronic form. Most students coming out of universities are not prepared for that."

While pursuing his doctoral studies in biophysics,

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UC Davis Biological Sciences

undergraduate, and parents

of current division students.

We welcome news from

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RESEARCH NEV

WATCH DNA UNZIP IN MICROMOVIE

BY CARL T. HALL

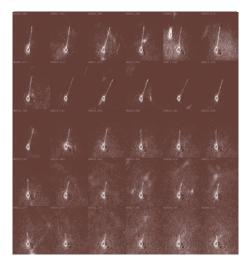
Reprinted with permission from the *San Francisco Chronicle*

sing some extraordinary camera tricks, scientists at the University of California at Davis have produced Lilliputian action shots of molecular "motors" unwinding strands of DNA.

Close to four years in the making, the grainy black-and-white movie stars an enzyme called helicase, chugging along brightly lit tracks of fluorescent-dyed microbial DNA.

The footage lasts only about a minute, but it's already attracting attention from other researchers who are trying to peer into the excruciatingly tiny realm of molecular motion.

"I have seen the movie," said Ron Vale, a biochemist at UC San Francisco. "It's striking, and it's completely clear.... It's



(above) A series of "stills" taken from video tape of the enzyme helicase (large round spot) unwinding DNA. The film took nearly four years to complete.

really amazing that you can now see single protein molecules in motion doing their work."

The images underscore recent dramatic advances in the field of nanotechnology, a discipline that scientists hope will allow

precise control over the very fabric of matter. The techniques used in the UC Davis experiments might ultimately be developed to repair DNA in patients with genetic illnesses.



Clip zips to No. 1. Professor of microbiology Stephen Kowalczykowski (above); Ronald Baskin, professor of molecular and cellular biology, and post-doctoral researcher Piero Bianco produced a film that shows the enzyme helicase unwinding strands of DNA. In late January, when the film was placed on the *San Francisco Chronicle*'s Web site, www.sfgate.com, it was clicked on more times than any other picture or video file.

A report on the experiments appeared in the January 18 issue of the journal *Nature*. Authors include microbiologist Stephen Kowalczykowski; Ronald Baskin, professor of molecular and cellular biology, and post-doctoral researcher Piero Bianco, all of UC Davis, collaborating with physicist Laurence Brewer and colleagues at Lawrence Livermore National Laboratory.

The experiments described in *Nature* build on earlier work—also led by UC Davis researchers—that showed how the helicase enzyme, called RecBCD—moves along in the fashion of an inchworm, jumping with each step 23 base pairs, the chemical units that make up the message-encoding parts of the DNA double helix.

Powered by ATP, the same energy source that fuels our muscles, the enzyme roars along at an astonishing speed—the equivalent of about 1,500 miles an hour, if DNA were scaled up to the width of a highway.

In its natural setting, this movement is a key part of natural DNA-repair processes. Defects in this system are linked to

various inheritable diseases, sensitivity to sunlight and certain forms of cancer.

It has been difficult to figure out how the enzymes go about their routine business in a molecular world too small to be

glimpsed directly. Now, Kowalczykowski and his colleagues are hoping to harness the process for medical purposes, including gene therapy and ultraprecise delivery of DNA-repairing drug payloads.

"Our very simple goal was to see in real time a molecular motor running along a strand of DNA, something that has never been visualized before," Kowalczykowski said.

It took a combination of sophisticated tools and custom engineering to make the DNA stretch out sufficiently

and hold still long enough to be photographed while the enzyme molecules were attached.

Lighting was provided by special fluorescent dyes that make DNA glow when it is in its typical double-stranded form, but not when the molecule has been unzipped into two single strands.

Glowing polystyrene beads were attached to one end of each piece of DNA to help anchor them down. The researchers also wielded laser beams as pairs of high-precision "optical tweezers" to help keep things under control.

A single helicase molecule, attached to a DNA molecule, was then loaded into one channel of a Y-shaped micromachined "flow cell." The action began when the ATP was added as fuel through the other arm of the "Y."

At that point, the video camera began recording the scene through an optical microscope, capturing the glowing DNA as the enzyme molecules marched along toward the polystyrene anchors.

The enzyme molecules are too small to be seen one at a time, even under the

microscope. And technical problems made it impossible to use more advanced nanotech imaging methods that make pictures by recording subtle atomic forces.

So the researchers had to settle for the indirect strategy of the special dye. As the enzyme unzipped the DNA into its two separate strands, the light appears to blink out in the enzyme's wake. The pictures show the DNA strands seemingly growing shorter and shorter.

In the end, only the polystyrene bead is still visible, even though the unlit DNA still dangles in place.

Kowalczykowski said additional experiments are planned to better understand how the enzymes work at the level of individual molecules—activity that biochemists traditionally have studied by mixing test tubes and beakers containing molecules by the millions.

Looking at a single molecule as it changes form is "a very powerful approach," Kowalczykowski said. "If you want to know how a car works, you can take an aerial view of Interstate 80 and see thousands of cars at once, which can tell you something," he said. "But if you really want to understand how a car works, eventually you have to take a close look at a single car."



FIRST FLOWERING PLANT **GENOME SEQUENCED**

he first complete genome sequence for a flowering plant, Arabidopsis thaliana, was published on Dec. 14 in the journal *Nature*. Anne Britt, professor in the Section of Plant Biology, contributed to the report.

Arabidopsis is a small, fast-growing plant widely used in plant biology research. The completed genome sequence may open up ways to study human diseases using plants.

The researchers found unexpected similarities to, and differences from, other organisms that have been sequenced, such as the yeast Saccharomyces cerevisiae, the fruitfly Drosophila, and the soil roundworm Caenorhabditis elegans.

Teams of scientists examined the DNA sequence for genes with particular functions. Britt, in collaboration with Jonathan Eisen of The Institute for Genome Research, looked for genes that repair damaged DNA.

"Twenty-seven genes were identified in *Arabidopsis* that were closely related to human disease genes," said Britt. Of these, a third were DNA repair genes, including genes linked to some types of



The first complete genome sequence for a flowering plant, Arabidopsis thaliana, was published on Dec. 14 in the journal Nature. Anne Britt (above), professor in the Section of Plant Biology, contributed to the report.

breast and colon cancer, and to the hereditary disease xeroderma pigmentosa, which makes children extremely susceptible to skin cancer.

"Arabidopsis has a similar distribution of repair genes to humans. Arabidopsis might turn out to be a very good model for the study of DNA repair in mammals," said Britt.

UC Davis has established a national presence in the study of DNA repair and chromosome biology, according to Stephen Kowalczykowski, director of the Center for Genetics and Development.



MICROBIOLOGISTS PLUMB THE DEPTHS

remote-controlled submarine is being used to study mysterious bacteria living half a mile below the surface of Monterey Bay. UC Davis microbial ecologist Doug Nelson, professor in the Section of Microbiology, leads the research.

The team uses a remote operated vehicle (ROV) owned and operated by the Monterey Bay Aquarium Research Institute (MBARI). The ROV Ventana is equipped with high-definition cameras and laser measuring equipment, and its arms can be fitted with a variety of collecting and sampling equipment.

> Because of their strange and hostile habitat, deep-sea bacteria have evolved survival strategies found nowhere else on Earth. Far from the sun, they live on hydrogen sulfide seeping from cracks in the seafloor and use nitrate in seawater instead of oxygen.

One example is Thiomargarita namibiensis, giant bacteria that grows to almost a millimeter in size—100 times the size of any other known bacterium. Thiomargarita was discovered in 1999 in deep water off the coast of Namibia in southern

Africa, by a team led by German scientist Heide Schulz.

This year, Schulz will join the UC Davis researchers studying bacteria around sulfide seeps in Monterey Canyon. Nelson's lab is particularly interested in Beggiatoa, bacteria that grow in long filaments on the seafloor mud.

Although these bacteria are deep in the ocean, they can be affected by fertilizer runoff, aquaculture and dumping of waste at sea, said Nelson.

"If you insult the ocean in a significant way, it will result in a die-off," said Nelson. That could lead to a release of hydrogen sulfide from the seafloor, which could affect fish and other marine animals and plants.

Website: http://www.mbari.org/dmo/ ventana/ventana.html.



Solid as an Oak

Retired professor's gift to benefit herbarium and arboretum

By KATHLEEN HOLDER

Botany Professor Emeritus John Tucker, who played a leading role in developing UC Davis' herbarium and arboretum, is helping to ensure the future health of both facilities with a \$500,000 gift.

The money will be divided equally between the herbarium and arboretum, enabling each, among other things, to maintain world-class collections of oaks—trees that have been the subject of Tucker's intellectual passion for most of his life.

The arboretum's share will establish the John M. Tucker Oak Collection Endowed Fund to maintain its oak trees, particularly those in the Peter J. Shields Oak Grove. The grove contains more than 80 kinds of oaks, many of them grown from acorns collected for Tucker from throughout the United States and northern Mexico as part of his research.

The portion for the herbarium is the first major gift toward construction of a \$3 million state-of-the-art facility to house its more than 200,000 dried plant specimens. Many of those specimens, including an extensive oak collection, were acquired during the 39 years Tucker directed the herbarium.

The new herbarium will be located in the future Sciences Laboratory Building, which is slated to open in mid-2004 east of the Life Sciences Addition.

The herbarium's \$2 million costs are part of an \$8.5 million "Opportunities for Distinction" fundraising campaign by the Division of Biological Sciences. The College of Agricultural and Environmental Sciences, which jointly administers the herbarium with the biological sciences division, is also raising private funds for the facility.

Tucker, 85, said he had been considering giving such a gift for a number of years.

"I've been interested in these two facilities my entire career, since the very beginning," Tucker said. "Both of them, at least in the early years, have had tough-going financially. Things have changed in recent years very much for the better. I just wanted to help things along."

Tucker joined the UC Davis faculty and began overseeing the Botany Herbarium in 1947 while he was completing his research for his Ph.D. from UC Berkeley. He received his degree three years later.

When he started at the herbarium, the specimens numbered fewer than 10,000. During his tenure, he started an exchange program and vastly expanded collections of weeds and California flora, as well as oaks. By the time he retired in 1986, the facility was renamed in his honor.

(In addition to the J. M. Tucker Herbarium Collection, the current UC Davis Herbarium also includes the Beecher Crampton Herbarium Collection—known for its California native grasses—which was started in 1913 and moved from another campus location in 1988).

Tucker also became involved in the arboretum's direction early in his career, joining its administrative committee in 1953, filling in as acting director in 1965-66 and serving as director from 1972 to 1984.

Ellen Dean, the herbarium's director and curator, said Tucker's gift is a demonstration of his "vision for and a true belief in the importance of the herbarium."

"As director of the herbarium in 1960, John designed our current herbarium space," Dean said. "We will always remember that he stepped forward first, when we needed to move beyond the space that he designed and into new facilities."

Arboretum Director Kathleen Socolofsky said Tucker's gift will "provide the critical resources necessary to ensure the long-term health of the



oaks and to fund educational and interpretive programs relating to the Shields Oak Grove.

"Dr. Tucker's gift is especially meaningful because of his role in developing our outstanding oak collection and because of his strong leadership as arboretum director in the 1970s and 1980s."

An authority on oak hybrids, Tucker said he developed a fascination for the stately trees while participating in scouting activities as a teenager in Santa Barbara in the 1930s.

With little money during the Depression for other activities, his scoutmaster took him and other members of his Boy Scout troop on weekend hikes and camping trips where they learned to identify the local trees and shrubs.

About the same time, the local natural history museum had an exhibition on oaks native to Santa Barbara County. "I was amazed to see how many different kinds there were. So I went back repeatedly."

Tucker said he pestered the curator into telling him where the different varieties grew. "It wasn't too long before I found myself hitchhiking to places where I could see these oaks."

On his longest trip, when he was about 16, he hitchhiked nearly 120 miles to the foothills of the San Gabriel Mountains in northern Los Angeles County to see the southernmost grove of Oregon oak. A forester who drove him the last leg of the trip in his pickup truck became a lifelong friend.

Tucker was the third of eight children born to a farming family in the tiny western Oregon community of Amity. His family moved to Ithaca, N.Y., when he was a toddler, then settled in Santa Barbara when he was four.

After graduating from high school in 1934, he enrolled at what was then Santa Barbara State College (now UC Santa Barbara). He transferred as a junior to UC Berkeley where he graduated with honors in 1940.

He entered graduate school at UC Berkeley, where he studied under the likes of G. Ledyard Stebbins, a pioneering plant evolutionist who would join UC Davis not long after Tucker.

World War II interrupted Tucker's doctoral studies. With poor eyesight keeping him from enlisting, he went to work as a machine welder for a Naval shipyard in Richmond. Later, he worked for a Shell Oil research plant in Emeryville as an overnight lab monitor, checking on chemists' experiments, until his major professor persuaded him to return to graduate school. He did even though it meant sacrificing his \$200 monthly salary to earn \$95 a month as a teaching assistant.

He and his wife, June, a fellow UC Berkeley graduate and the only child of a bank vice president, moved to Davis in 1947 while June was expecting the first of their three children.

Tucker joined the faculty of what was then the Botany Department and is now the Section of Plant Biology.

His research focused on taxonomy of oaks and intermingling of oak species. "Oaks have a long reputation of being a difficult group to classify taxonomically. They're horrendously variable, so it's hard to draw a line from where species A starts off and species B begins."



Oak trees in the arboretum (above) will benefit from John Tucker's gift, part of which establishes the John M. Tucker Oak Collection Endowed Fund.

Tucker shares an office near the herbarium and still conducts research. However, he said he no longer approaches it with the same single-minded intensity as he did before his wife died of cancer in 1987.

Other interests also occupy his time, he said. A selfdescribed newspaper addict, he reads three newspapers daily as well as a number of other publications. He also has traveled to China, Central America, Scandinavia and Russia since retiring.

"There's no question I'm slowing down. The desire is still there but my research has tapered off."

However, directors of the herbarium and arboretum said Tucker's gift will ensure a long-lasting legacy for his many contributions to a better understanding of oaks and other plants.











...BIOTECH SCHOOL

(continued from front page)

of technological evolution rendering a class obsolete. "In 1990, we offered a class in alternatives to radioactive labelling, which were newly developed," she says. "The health, environmental, disposal and monetary costs of using radioactive labels had become apparent, and I felt it was important to introduce these techniques." Today the techniques are standard, and the course isn't offered separately, its content having been incorporated into the biotechnology program's other classes.

Enter polymerase chain reaction (PCR), a technique used to make numerous copies

Martina McGloughlin (above), director of UC Davis' Biotechnology Program, is aware of only a few other places in the United States that offer classes similar to the Biotechnology Program's courses.

of a specific segment of DNA quickly and accurately.

"By the early 1990s, PCR was really taking off," says McGloughlin. "I contacted someone I knew at Perkin-Elmer, who was willing to provide equipment and teach a class."

McGloughlin continues, "We still offer an advanced PCR class and it's one of the most highly subscribed to. We constantly update it because the technology is constantly evolving." She laughs as she recalls that a few years ago several employees from Chiron, a biotechnology firm, attended the class. ""The Chiron

employees were essentially coming from where PCR originated because Chiron took over Cetus, the company that invented PCR. So it was funny that they came here to learn the technique.

Apparently people in the company were simply too busy to teach the technique."

She adds, "We offered *in situ* PCR for two years, but the technique was so system-specific and temperamental that it didn't lend itself to a generic lab course. A course on carbohydrate research didn't garner enought interest to warrant the time and energy involved in offering it. So in addition to courses that became obsolete, courses that just didn't work out have been a part of the program's growth."

McGloughlin then took the initiative to discover what type of biotechnology instruction was needed.

"I thought we should offer a protein analysis class," says McGloughlin, "although the first time we offered it not many people enrolled. Since then people have moved from genomics to proteomics [quantifying all the proteins expressed at any given time in a cell] and the enrollment has increased. The class has also evolved from basic protein analysis—isolation, purification, and character-

ization—to true proteomics using 2-D gel analysis and advanced mass spectometry. We teach the class in conjunction with the UC Davis Molecular Structure Facility, which helps familiarize people with the powerful technology that's available on this campus."

A burgeoning new discipline motivated McGloughlin to expand course offerings in 1995-96. "It became clear to me that we needed to focus on bioinformatics, which addresses how to electronically store, organize, and access the massive datasets being generated in the life sciences," says McGloughlin. "For so long

biologists went into biology because they were math averse—we can't afford to be that way any longer."

The initial bioinformatics course was a survey course and McGloughlin discovered it wasn't focused enough.

"For the bioinformatics course we had a huge spectrum. At one end were biologists who wanted to know something about bioinformatics but didn't want to know what happens inside the black box; they didn't want to know how to program computers. At the other end were the computational people who wanted to know more about biology, and then there was a whole bunch of people in the middle who wanted to know a bit of both." In 2001, McGloughlin therefore split up the one bioinformatics class into two sections, each of which covers a different aspect of the subject.

Overall the fast-paced courses receive excellent evaluations. "The content of the courses usually challenges the students," says McGloughlin, "and often they learn more than they had expected to." She laughs again as she says, "A woman in the protein analysis class said her head hurt afterward, but she gave us a great evaluation."

Depending on pre-enrollment, McGloughlin plans to offer the following courses in summer 2001:

- ► In Situ Diagnostics and Analysis
- ► Confocal Microscopy
- ► Advanced Polymerase Chain Reaction Techniques
- ► Bioinformatics I: Instrumentation and Biological Samples
- ➤ Bioinformatics II: Databases, Visualization, Data Mining and Integration, Algorithms and Image Processing
- ► Protein/Proteome Analysis

Information about enrollment, costs, and course content is available at http://www.biotech.ucdavis.edu/courses/new courses.htm.

FOCUS ON FACULTY

CARL SCHMID: A STORY WITH REPETITIVE ELEMENTS

he Christmas he was 7 years old, Carl Schmid was walking with his father when he spied a shiny chemistry set displayed in a store window. Schmid asked if he could have the set; his father replied, "No, you have to become a chemist to have that." The younger Schmid immediately thought, "Then I'll become a chemist."

His resolve never wavered.

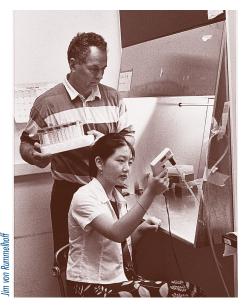
"Chemistry was my childhood hobby," Schmid says. "I set up a home lab and cut grass to buy chemistry sets. My friends and I made explosives and concocted our own experiments. It's something I really wanted to do."

As an undergraduate, Schmid attended Drexel Institute, Philadelphia, Pa., where students attend school for six months and work for six months. His job not only made college affordable, but also exposed him to the research that would become a primary interest. "I was placed in a laboratory and became interested in nucleic acids, an interest that led me to pursue my doctoral studies under John Hearst, who was studying DNA structure using physical methods, at UC Berkeley."

After receiving his doctorate in biophysical chemistry in 1971, Schmid took a post-doctoral position at California Institute of Technology in Pasadena, where he became interested in studying eukaryotic genome structure at a time when sequencing and cloning techniques had not yet been developed. He continued his post-doctoral studies at UC Davis "with the idea," says Schmid, "that I'd study the structure and organization of the human genome using physical methods developed at Cal Tech."

Schmid became a faculty member in UC Davis' Chemistry Department in 1973. His research on DNA structure led. in 1990, to a joint appointment in the chemistry department and Department of Genetics (now the Section of Molecular and Cellular Biology).

Since the late 1960s scientists had been aware that the eukaryotic genome



Carl Schmid (standing) and his colleagues discovered, in 1979, a family of repetitive DNA sequences. Known as "Alu repeats," these repetitive sequences were characterized as "junk DNA." However, the recently published human genome maps revealed that Alu repeats constitute 10 percent of the genome and are located near genes. These findings have caused scientists to reexamine Schmid's controversial theory that the Alu repeats aren't junk DNA but are activated in response to cellular stress.

contained repetitive DNA sequences. The prevailing theory was that many distinct families composed the repetitive DNA, each distinct family having a few thousand members interspersed throughout the genome.

Careful analyses carried out by Schmid and his laboratory revealed, in 1979, a situation very different from the prevailing theory: a major fraction of the repetitive sequences belonged to a single family—the Alu family, which appeared to constitute about five percent of the human genome.

Walter Eckhart, a molecular biologist with The Salk Institute, noted in 1981, "His [Schmid's] recent work on interspersed sequences in the human genome has been a major contribution to a

particularly interesting and important area of modern genetics. It has won him international recognition and has helped to advance the work of many others in the field "

Most scientists believed the ubiquitous Alu repeats were purposeless—along with other types of DNA sequences that don't code for genes, they fell under the rubric "junk DNA"—but in 1998 Schmid proposed the controversial theory that Alu repeats are activated when cells are damaged in some way, for example by heat, toxic compounds or lack of essential nutrients, and help repair damage by controlling genes that make proteins.

"We found that these repeating elements are expressed, normally at very low levels, but when a tissue is stressed, their expression increases," says Schmid.

The recent publication of the human genome maps stimulated a reexamination of Schmid's hypothesis.

"There's tremendous excitement now that the genome has been sequenced," says Schmid, "and it's been observed that Alu repeats are extremely abundant throughout the genome, and are often found near genes that code for proteins."

In fact, the maps show that Alu repeats make up 10 percent of the genometwice as much as was previously supposed.

In regard to the widespread presence and strategic placement of Alu repeats, Francis Collins, director of the National Human Genome Research Institute, was quoted in a New York Times article as saying, "...the natural conclusion is, this part of our junk DNA isn't junk."

Schmid comments, "I can't say at this point that my theory is right or wrong, but the new genome maps certainly have intensified interest in my laboratory's research."

Prescott Deininger, Zimmerman Chair for Cancer Research at Tulane University's School of Public Health and Tropical

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...Schmid

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Medicine, New Orleans, La., also studies Alu repeats and was a graduate research assistant in Schmid's laboratory from 1974 to 1978.

Says Deininger of Schmid's research, "The recent focus on the sequence of the human genome has really highlighted the incredible role that Alu has played in shaping the human genome.

"Carl has been one of the major contributors to the characterization of this type of element for over 25 years. In recent years his has been the major laboratory in the world focusing on the quest for possible functions of those elements. He's a rigorous and imaginative scientist."

According to Schmid the new genome maps will lead to more studies of how Alu repeats influence genes.

"Until we resolve the question of whether these sequences are junk, or have a defined function, it's impossible to understand the evolution of genome structure," he says.





ALUMNI

1996

Kevin A. Morano, Ph.D. Microbiology, 1996, is currently an assistant professor in the Department of Microbiology and Molecular Genetics at the University of



Kevin Morano

Texas Medical School in Houston.

He conducted his doctoral research in Professor Daniel Klionsky's laboratory (Klionsky is presently with the University of Michigan) studying targeting and

assembly of vacuolar proteins. He continued his education with postdoctoral



Three UC Davis graduates in New Caledonia having breakfast at their motel, Le Grand Cerf, which means "the big deer." From left to right: Mike Irwin, Jason Bradford, Barry Donovan.

studies in the Department of Biochemistry at the University of Michigan Medical School (1996-2000), studying yeast stressresponse mechanisms.

Morano is currently studying the functions of a class of proteins called "molecular chaperones," which help proteins fold in the cell. Many important cellular regulatory proteins are very unstable and require a protein chaperone called Hsp90 to protect them. Says Morano, "My laboratory is interested in how the cell in turn regulates the function of the chaperones themselves. We're also using the latest in DNA microarray technology

We welcome news from alumni for the "People" section; please e-mail the editor at dccleveland@ucdavis.edu. Or visit the alumni Web site at www.dbs.ucdavis.edu/ alumni/postcards/ and send us a postcard.

to identify new stress genes in the model eukaryotic cell Saccharomyces cerevisiae, or baker's yeast. Many of these genes have close human homologs, and by studying the genes in yeast, we may learn about how they are recruited for 'stressful' pathological states in our own cells, like cancer and aging."

1992

Jason Bradford, B.S., Biological Sciences, 1992, received his Ph.D. in evolutionary and population biology from Washington University, St. Louis, Mo. Says Bradford, "My current position is research associate at the Missouri Botanical Garden in St. Louis. I'm also a course instructor in ecology at Washington University.

"I was traveling in New Caledonia, in December 2000, on a grant from the National Geographic Society to study the systematics and pollination biology of a family of flowering plants, Cunoniaceae, that I specialize in. Along with me was Barry Donovan from New Zealand, who knows more about pollination biology than I do, and is an expert on the bees of New Caledonia. Barry received his doctorate from UC Davis in 1969 in entomology.

"While staying in a small motel in the northern town of Koumac, Barry and I met Mike Irwin. It turns out that Mike

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...PEOPLE

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was there to study a group of flies he is doing a worldwide monograph on, and he also graduated from UC Davis—with a B.S. in entomology in 1963. He also has a Ph.D. in entomology from UC Riverside and is currently a professor at University of Illinois.

"My wife Kristin Bradford (née **Sparks)** also graduated from UC Davis in 1992 with a B.S. in physiology. She was the division's first student commencement speaker. She now has a medical degree from University of Vermont and is finishing her residencies here in St. Louis (family practice and preventive medicine). We plan to move back to Davis in July 2001 with our twin boys, Curtis and Davis, who will be 2 1/2 years old by then."

FACULTY

The Botanical Society of America awarded its 2000 Merit Award, its highest honor, to **Leslie Gottlieb**, a professor in the Section of Evolution and Ecology. An announcement in Plant Science Bulletin, the society's newsletter, described Gottlieb as "one of the most influential plant evolutionary biologists over the past several decades." The award committee chair, University of Wisconsin genetics professor John Doebley, cited three of Gottlieb's publications as classics, including a 1984 article in American Naturalist that "has been called one of the most important papers in plant evolutionary biology during the past half century." However, Doebley added that Gottlieb's "greatest contribution may have come through his influence on the careers and research of a substantial number of plant evolutionary biologists, including many of the people most active in this field today."

IN MEMORIAM

UC Davis exercise biologist Paul **Molé**, whose research contributed to an understanding of how muscles function during exercise, died on January 20 of complications following a heart attack. He was 62.

Molé became ill on Jan. 8 while teaching a class. Thanks to prompt action by his teaching assistants, he was taken to Sutter Memorial Hospital and later transferred to the UC Davis Medical Center in Sacramento. Although doctors were able to remove a blood clot from a coronary artery, his condition deteriorated, and he died early Saturday morning.

Molé's main research interest was in skeletal muscle metabolism, and how it responds to exercise, said UC Davis Professor Emeritus Ed Bernauer, a friend and colleague for more than 40 years.

"He reanalyzed much of the important documented research, and formulated a mathematical model that challenged the accepted view," said Bernauer. While the conventional view held that carbohydrate, not fat, was the primary fuel for muscles, Molé believed that fat played a greater role earlier in exercise, according to Bernauer. He then went on to test his ideas using magnetic resonance imaging (MRI) techniques.

"He'll be sorely missed," said Tom Jue, a biochemist in the School of Medicine who worked closely with Molé. Using MRI technology developed in Jue's laboratory, they could look inside an athlete's limb to see how the muscles used oxygen and fuel during exercise. As a physiologist with expertise in muscles, Molé's contribution was very important in helping to develop the technology, said Jue. Eventually, the same technology might be used to study heart disease, he added.

Molé was vice chair of the Department of Exercise Science and held an adjunct appointment at the Department of Physical Medicine and Rehabilitation. He was a faculty member of the graduate groups in exercise biology, nutrition and physiology.

Molé played a pivotal role in the recent restructuring of the exercise biology program, and was actively involved in

recruiting new faculty, said department chair Chuck Fuller.

"We're going to miss his insight as we build for the future," said Fuller.

"He was an intense and committed educator, who felt it was the obligation of the professor to bring students to the highest level of understanding," said Bernauer. "Paul was always challenging his students, always pushing them to think," said Fuller.

Brought up in Jamestown, N.Y., Molé entered the University of Illinois, Urbana-Champaign, on a football scholarship, graduating in 1960. He stayed at Illinois to complete a M.S. in physical education and a Ph.D. in physiology. He joined UC Davis in 1977.

Molé was a Fellow of the American College of Sports Medicine, and a member of the American Physiology Society and the New York Academy of Science. He served as president of the Southwest chapter of the American College of Sports Medicine from 1999 to 2000.

Molé loved the outdoors, whether walking in the woods, fishing or gardening, said his wife of 42 years, Patty. He was a keen amateur photographer. His parents were bakers, and he liked to bake and cook, making fresh bread every weekend, she said.

He is survived by Patty; their three children, Pam, of Oklahoma, Greg, of Pittsburgh in the Bay Area, and Michael, of San Jose; and five grandchildren. In his freshman year on a football scholarship, Molé told his coach that he was marrying Patty, his high-school sweetheart. The coach told him he could either get married or play football, but not both. Molé responded by giving up football and taking up fencing, joining a team that went on to win the Midwestern Big Ten championship, an achievement that typified his determination and persistence, said Bernauer.



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...BIG PICTURE

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Smietana supported himself by programming computers. He thus possesses both biological and computational perspectives. Smietana has been associated with the biotechnology industry for more than 18 years; while with Ciphergen Biosystems and GeneLogic he created novel methods for integrating protein and DNA expression data with public and private clinical and genomic databases. For the past three years, he has taught the summer bioinformatics short courses.

Underscoring the need to think big, Smietana talks about the direction in which the life sciences industry is headed. "I want students to be aware of the scaling up of technologies in industry," he says. "As an example, LumiCyte is going to have 20, perhaps 30 or 40 instruments, with multiple sites throughout the world, collecting information on proteomic biochips. Again, the amount of information is staggering—we're talking about terabytes of information."

According to Smietana, what this translates into is a need for people who know how to design complicated, integrated experiments.

"We're moving into a time when we'll need to integrate our knowledge of genes with our knowledge of proteins," says Smietana. "Genes don't mean very much without the context of proteins and vice versa. So the scope of your experiment has to change. You now have to consider bringing in more information than you ever have before. That means you're going to have to learn something about the tools that are out there. A student may study one protein for her doctoral research, but we now have the ability to study sets of proteins."

When asked how undergraduates can prepare themselves for careers in bioinformatics, Smietana replied with a brief anecdote. "I was recently at a meeting with representatives from Genentech, DoubleTwist, Applied Biosystems, and Iconics," he says. "Asked for description of an ideal candidate for a position in bioinformatics they responded, 'A full professor in biological sciences with 20 years experience in the computer field.' We know that person doesn't exist, so how can you parlay that down? The best strategy for an undergraduate is to take a diverse program."

And remember: think big.



