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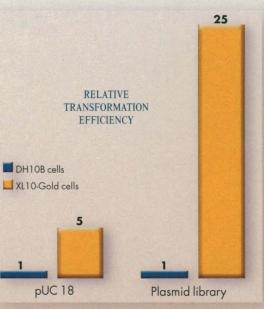
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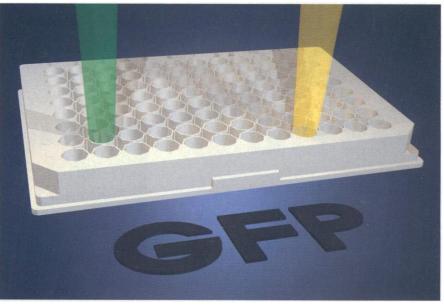
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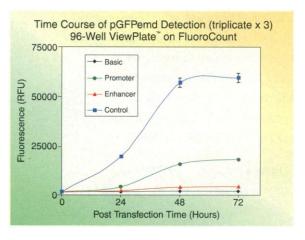
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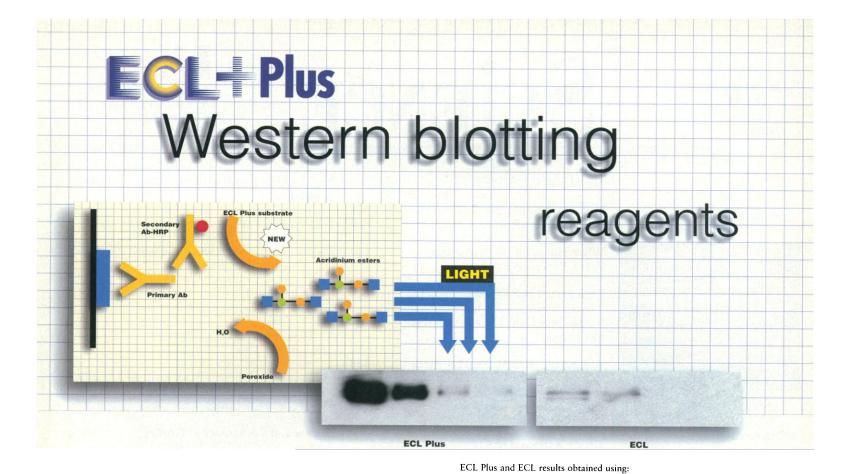
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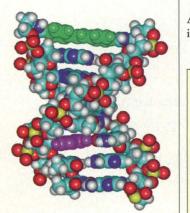
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ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE



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COVER

Plot of the structure in comet Hyakutake. The nucleus is the bright peak, and the dust coma (green area at top) is visible toward the sun. Trailing the nucleus is an arc of cyanogen (CN) emission (green arc) and two dust condensations (below the green arc). Simulations imply that the CN arc resulted from interaction between the gas coma and a secondary source, possibly the inner dust condensation. Field of view, 8100 kilometers. See page 676. [WIYN Observatory image, by Walter Harris, University of Wisconsin, Madison]

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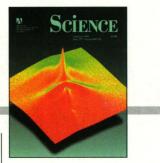
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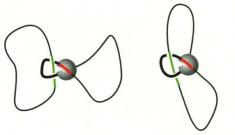
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Untying knots





681 Delayed delivery



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 "The I.M.A.G.E. Consortium: An Integrated Molecular Analysis of Genomes and their Expression", Lennon, G.G., Auffray, C., Polymeropoulos, M., and Soares, M. B. (1995) Genomics.

THIS WEEK IN SCIENCE

edited by BROOKS HANSON

Electrons through vibrations

Electron transfer (ET) plays an important role in many chemical and biochemical reactions, but because ET is usually very fast, it is difficult to study. It o et al. (p. 660) synthesized a series of model compounds that allow tuning the rate at which ET occurs in an intramolecular transfer reaction over a wide range of times. The system allows them to correlate vibrational absorption line shapes with the ET dynamics.

Chemical plants

The secondary metabolites of plants, which contribute to their interesting flavors, have other functions, including warding off insects and microbial pathogens. Frey et al. (p. 696) found that in maize a series of five enzymes is sufficient to synthesize one of these defense compounds, DIBOA, from a common precursor. The first of the enzymes resembles a subunit of tryptophan synthase but represents a branchpoint in the pathway that leads away from tryptophan synthesis. The other enzymes are cytochrome P-450dependent monooxygenases. Together these five genes confer on transgenic yeast the ability to synthesize DIBOA from indole-3-glycerol phosphate.

Electrons through hairpins

Electron transfer (ET) through DNA has been controversial; some results suggest that DNA is an efficient "molecular wire" and others suggest that it acts as a more protein-like insulating state. Lewis *et al.* (p. 673) measured photoinduced ET rates in DNA hairpins in which a stilbene dicarboxamide group bridges two connecting DNA arms. No ET was seen in a six AT base pair hairpin, but introducing a

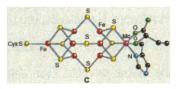
DNA in knots

At thermodynamic equilibrium, a distribution of isoforms of closed circular DNA (superhelical coils, knots, and catenanes) is formed from linear DNA. DNA topoisomerases can catalyze strand passage between two DNA segments, thereby affecting DNA topology. Isoforms formed with type I DNA topoisomerase approach that which is observed at equilibrium. However, Rybenkov *et al.* (p. 690) show that with topoisomerase II the distribution of isoforms is below the equilibrium values. Type II topoisomerases use the energy of adenosine triphosphate hydrolysis to reduce the number of knots and catenanes and simplify the topology. These findings have implications in DNA replication and chromosome segregation, as highlighted in a Perspective by Pulleyblank (p. 648).

single GC pair led to distancedependent fluorescence quenching. Although not a molecular wire, ET is more efficient than in proteins.

Clusters in proteins

Many proteins contain ironsulfur clusters, which facilitate electron transfer reactions and can act as catalytic centers and as sensors for iron or oxygen.



Beinert *et al.* (p. 653) review our understanding of the various structures and properties of these prosthetic groups, including results on studies of model compounds and on the incorporation of the clusters into proteins.

Inducing labor

Observations that aspirin-like drugs could delay childbirth led to speculation that prostaglandins might be involved in the regulation of labor. However, oxytocin seemed more effective. Sugimoto *et al.* (p. 681) have now sorted out these observations. They used a knockout mouse lacking a prostaglandin receptor to show that prostaglandin $F_{2\alpha}$ is required to inactivate the corpora lutea. This inactivation in turn leads to a decline of serum progesterone levels and induction of oxytocin receptor, followed shortly by labor and delivery.

Boundary effects

The thin boundary between Earth's core and mantle may play a key role in convection in the mantle as well as in the origin and structure of Earth's magnetic field (see the Perspective by Kellogg, p. 646). Two reports provide information on the character of this boundary. Earle and Shearer (p. 667) measured scattering of seismic waves propagating through the layer. The scattering may be explained by topography on the boundary of about 300 meters over wavelengths of about 8 kilometers. Revenaugh and Meyer (p. 670) examined seismic waves reflected from just above the boundary. They were able to map several regions along the boundary where the seismic waves were slowed considerably. These data suggest that much of the boundary contains a thin, <15-km-thick boundary of melt.

Snowball fights

Comet Hyakutake recently passed within about 0.1 AU of Earth. Harris *et al.* (p. 676; cover) used narrow band filters

on the WIYN telescope at Kitt Peak to image the structure of the inner coma for a 7-hour period during Hyakutake's closest approach. They found that there was an extended region of icy particles that is separated from the gas and forms arcs and extended clumps well beyond the comet nucleus. They conclude that there is another local source of icy grains behind (away from the sun) the nucleus of the comet that produced some of the volatiles detected from Hyakutake.

Baby talk

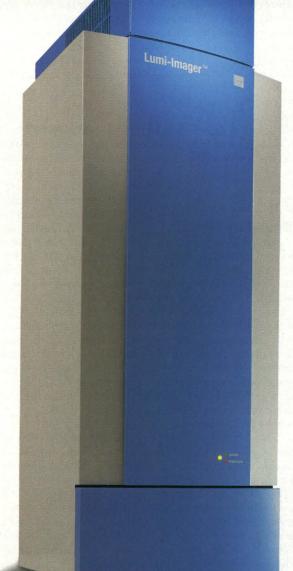
Why do mothers speak differently when talking to their infants than when talking to their husbands? Kuhl et al. (p. 684) report the results of a cross-cultural study using 30 mothers fluent in English, Russian, or Swedish. The data reveal that vowels are exaggerated when speech is directed toward infants. Because infants appear to be able to discriminate normally pronounced vowels, the authors suggest that mothers are emphasizing the parameters that infants must learn in order to speak themselves. See also the news story by Barinaga (p. 641).

Right place and time

The El Nino-Southern Oscillation (ENSO) phenomenon dominates tropical Pacific climate and influences global climate, but the exact mechanism of the oscillation is not yet fully understood. Picaut et al. (p. 663) used recent data on the eastward or westward advection of surface waters and the resulting convergence of water in the central Pacific to modify the classical delayed action oscillator model. When these features are included in a simple model of ENSO, observations and simulation match well, indicating that these features are central to the ENSO mechanism.

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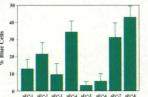


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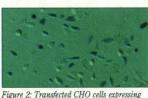
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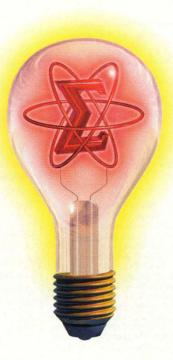
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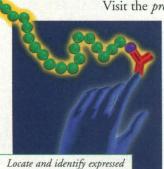


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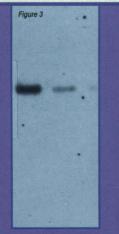
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Figure 2. Mouse B-actin was detected using a fluorescein-labeled ssRNA probe and ready-to-use CDP-Star. This film demonstrates results obtained using Renaissance RNA Fluorescein Labeling Kit with Antifluorescein-AP (NEL633) in conjunction with ready-to-use CDP-Stat (NEL601). Blots were exposed to Reflection film for 5 min-

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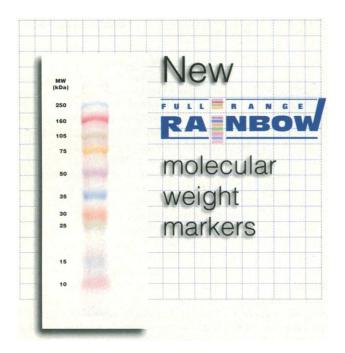


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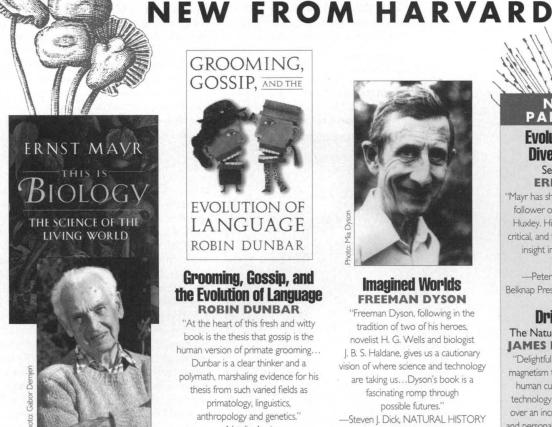
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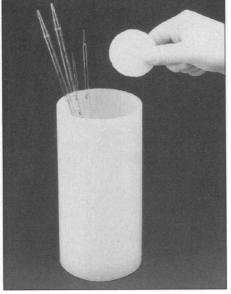
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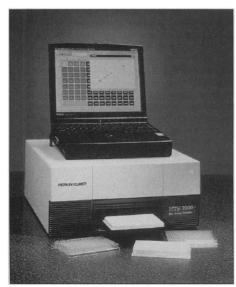
HazeBuster and MicroTome are software packages that remove out-of-focus haze from images acquired with microscopes, including confocal microscopes. Prior to the introduction of these stand-alone versions, researchers had to purchase additional, expensive image processing software to work in conjunction with these applications. Because they are modular, the new versions allow researchers to use image processing software from other developers. For example, Micro-Tome and HazeBuster can be used alone or in conjunction with Optimus, Image-Pro Plus, or IP Lab Spectrum. HazeBuster is designed for researchers who need to keep costs down, but must have low-noise, highly detailed images normally obtained from expensive confocal equipment. MicroTome is an upscale digital deconvolution program that provides accelerated performance and many additional options, including user-selectable algorithms, automatic, true-color imaging, cell counting, measurements, and more. Both programs work with a broad range of fluorescence, transmitted, reflected, and other image modes and have applications in medical imaging, industrial inspections, nondestructive testing, and other microscopy procedures. Vaytek. For information call 515-472-2227 or circle 145 on the reader service card.

PCR Product Quantitation

The Sunrise Amplification Detection System permits the simultaneous amplification and detection of nucleic acids within the polymerase chain reaction (PCR) tube. The method is based on the incorporation of energy transfer-labeled hairpin primers into the amplified DNA product. The Sunrise Uni-Primer I, for amplification of any target, is designed so a fluorescent signal is generated only when the primer is unfolded by incorporation into the amplification product. It has virtually no fluorescent signal, eliminating the need to purify the PCR product prior to detection. Oncor. For information call 301-963-3500 or circle 146 on the reader service card.

Bio Assay Reader

The HTS 7000 Bio Assay Reader is a versatile system for rapid, high-throughput screening of a large array of small-volume samples routinely performed in many molecular biology, cell biology, and biochemistry labo-



ratories. Applications for the new system include DNA and protein quantitation; drug screening and therapeutic drug monitoring; immunoassays; cell function studies; and DNA hybridization, gene expression, recombinant enzyme, and reporter gene assays. The HTS 7000 includes software designed for processing up to thousands of samples per day. Top or bottom reading of luminescence or through-well absorbance measurements are achieved automatically through simple software selection. Users with few samples can take advantage of the convenience of pipetting in strip wells and perform traditional cuvette-based analyses more rapidly. Perkin-Elmer. For information call 800-762-4000 or circle 147 on the reader service card.

Literature

Chemicon 1997 Immunological Reagents catalogs more than 500 new reagents for use in neuroscience, signal transduction, adhesion molecule, extracellular matrix, cytokine, cytoskeletal, and infectious disease research. Chemicon International. For information call 800-437-7500 or circle 148 on the reader service card.

The Cytosensor Microphysiometer at Work Reference Guide lists more than 150 scientific references describing the use of this instrument for a variety of applications. These applications include receptor activation and inactivation, signal transduction elucidation, agonist and antagonist profiling, experimental in vitro toxicology, microbial metabolism study, and study of growth factor, neurotrophic factor, and cytokine responses. This instrument can detect responses triggered by a wide variety of receptor classes coupled to different second messengers; these receptor classes include G protein-linked receptors, ligand-gated ion channels, and tyrosine kinase receptors. Molecular Devices. For information call 408-747-3542 or circle 149 on the reader service card.

Dako 1997 Catalog features a wide variety of monoclonal and polyclonal antibodies targeted against apoptotic markers, cell cycle proteins, multi-drug resistance markers, and cancer markers as well as a comprehensive listing of conjugated and unconjugated secondary antibodies. The catalog also describes immunostaining systems for immunohistochemistry and in situ hybridization and products for flow cytometry. Dako. For information call 800-424-0021 or circle 150 on the reader service card.

Newly offered instrumentation, apparatus, and laboratory materials of interest to researchers in all disciplines in academic, industrial, and government organizations are featured in this space. Emphasis is given to purpose, chief characteristics, and availability of products and materials. Endorsement of any products or materials mentioned in Tech.Sight by *Science* or AAAS is not implied. Additional information may be obtained from the manufacturers or suppliers named by circling the appropriate number on the Readers' Service Card and placing it in a mailbox. Postage is free.



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At SmithKline Beecham the application of combinatorial chemistry and highthroughput screening is resulting in an extraordinary increase in the numbers of compounds and corresponding data being generated for drug discovery. We are expanding our transnational Cheminformatics group to work closely with medicinal chemists and screening scientists in the UK and US in the collection, transfer, manipulation and exploitation of these data.

This expansion opens opportunities for careers which can be based at either our US or UK state-ofthe-art facilities in the following areas:

- Medicinal Cheminformatics developing a Medicinal Chemistry information system to meet the demands of combinatorial and automated chemistry; developing new methods to analyse molecular diversity.
- Screening Informatics developing methods for dynamic design of efficient screens; developing sophisticated statistical analysis and data fitting; data interfaces to new screening technologies.
- Knowledge Discovery in Databases developing and applying computational tools to find relationships within and among large chemical and biological databases.
- Intelligent Data Integration designing and developing systems for model-based integration and presentation of chemical and biological data.

Specific opportunities exist for the following positions:

Group Leader - Screening Informatics

Requirements include a PhD in biological sciences or chemistry with at least 5 years' directly related experience in information analysis, high-throughput screening or laboratory automation, and a record of successful performance in pharmaceutical discovery or biomedical research. **Refer to Job Code H7-0344**.

Group Leader - Knowledge Discovery

Requirements include a PhD in physical, biological or computer sciences or statistics, with at least 5 years' experience in pattern recognition, machine learning or chemometrics and a proven record of performance in chemical or biological database analysis. **Refer to Job Code H7-0244**.

Group Leader - Data Integration

Requirements include a PhD in chemical or biological sciences, plus a broad understanding of computer technology and at least 5 years' relevant experience with a record of success in computational applications to chemistry and biology. **Refer to Job Code H7-0273**.

PhD Scientist -Medicinal Cheminformatics

Requirements include a PhD or equivalent experience in chemistry or a related discipline plus significant experience providing computer support within a scientific environment or extensive experience in the use of chemistry software packages and databases. The position entails ensuring efficient selection and use of computer tools, development and implementation of a training strategy for Cheminformatics and Medicinal Chemistry applications, and working with multidisciplinary groups to identify and evaluate new applications. This position is US-based. **Refer to Job Code H7-0318**.

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