

# SAME-DAY DETECTION OF MYCOPLASMA INFECTION

# The First PCR\*-based Technique for Rapid Detection

- Produces a Diagnostic Fingerprint
- Eliminates Preculturing with Antibiotics
- Detects Infection in <30 Cell Equivalents
- Sensitive and Fast

Mycoplasma infection of cell lines represents a major problem in tissue culture. Besides affecting cellular growth and morphology, mycoplasma infections can have drastic effects on cell metabolism involving DNA, RNA, and protein. To assure the integrity of cell cultures, various

serological, cell culture, or molecular means typically are employed to look for mycoplasma infection. The problem with these methods is that they can be time-consuming and costly, and may yield results which are subject to interpretation. Now Stratagene introduces a convenient, rapid, and sensitive means of testing for the presence of mycoplasma in eucaryotic cells.

Stratagene's Mycoplasma PCR Primer Set detects all forms of mycoplasma associated with infection of cell lines, including *Mycoplasma arginini*, *Mycoplasma hyorbinis*, *Mycoplasma orale*,

Mycoplasma fermentans, and Acholeplasma laidlawii. The easy-to-interpret results consist of a fingerprint indicative of infection. Uninfected cells do not yield any PCR product. This non-radioactive PCR-based test offers sensitivity of 30 cell equivalents or less. And with no need to preculture cells, it delivers results the same day you begin testing.

Catalog # 302007

\*The Polymerase chain reaction (PCR) process is covered by U.S. patents owned by Hoffman-La Roche. Use of the PCR process requires a license.

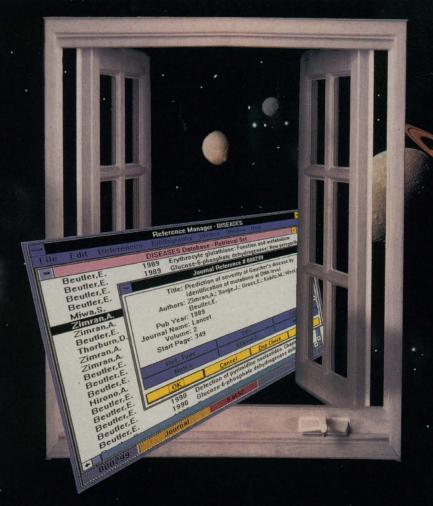


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ptimizing each part of the pre- and post-PCR\* process — primers, sample preparation, and product detection — maximizes the speed and cost efficiency of the process as a whole. Now laboratories can com-

plete their research faster, more economically and with reliable results, by improving the critical steps of the PCR process using products from Beckman.

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interface with a variety of labware offers the flexibility of using standard PCR plates and tube formats. Plus, contamination is effectively controlled with disposable aerosolbarrier pipette tips.

# DNA Synthesis Sample Preparation PCR PCR PHY-1 AND HLAD DALES EXCHENCES WITH THAZOLE PROMATE WHESTAL A STRAW TS\_1 HV-1 AND HLAD DALES AL STRAW PROMATE WHESTAL A STRAW PROMATE WHESTAL A STRAW TS\_1 HV-1 AND HLAD DALES AL STRAW PROMATE WHESTAL A STRAW PROMATE W

Start with high quality primers, automate sample preparation, and improve the detection process.

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- \* PCR is covered by U.S. patents owned by Hoffman-La Roche Inc.

  <sup>†</sup> Mark Holodniy, Mark A. Winters and Thomas C. Merigan. BioTechniques 1992,
- <sup>1</sup> Mark Holodniy, Mark A. Winters and Thomas C. Merigan. BioTechniques 1992 Vol. 12, a: 1, 36-39.
- <sup>2</sup> Herbert E. Schwartz and Kathi J. Ulfelder. Anal. Chem. 1992, 1737-1740.

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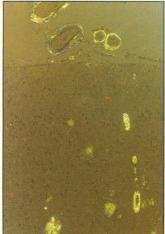
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# Science

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Changing of the guard at NIH



Apolipoprotein E type 4 and Alzheimer's disease

# NEWS & COMMENT Varmus Tapped to Head NIH 820 How Varmus Got the Job DOE Pulls Plug On SSC Contractor 822 Academic Earmarks: Report Takes Aim 823 at 50 New Projects Scientific Misconduct: Army Clears 824 Redfield-But Fails to Resolve Controversy RESEARCH NEWS New Piece in Alzheimer's Puzzle 828 Protein Signals Frog Embryo to Start 830 Getting a Head

How Collapsing Stars Might Hide Their

Artificial Atom Unveils Quantum Effects

Things Start Getting Sticky for a Cell

Tracks in Black Holes

Surface Enzyme

## SPECIAL SECTION COMPUTING IN SCIENCE **NEWS REPORTS** Beyond Databases and E-Mail • AI Helps Researchers Find Meaning in Molecules • Fitting Planet Earth Into a User-Friendly Database • Frustrated With Fortran? Bored by Basic? Try OOP! **PERSPECTIVES** The Ever-Present Packaging Challenge 851 L. M. Thorndyke and J. P. Riganati Workstation Clusters Rise and Shine 852 B. Buzbee The Collaboratory Opportunity 854 W. A. Wulf **ARTICLES** Parallel Scientific Computation 856 W. D. Hillis and B. M. Boghosian Microprocessors: From Desktops to 864 Supercomputers F. Baskett and J. L. Hennessy Genetic Algorithms: Principles of Natural 872 Selection Applied to Computation S. Forrest

# 828 & 921

#### DEPARTMENTS THIS WEEK IN SCIENCE 809 SCIENCESCOPE 819 **EDITORIAL RANDOM SAMPLES** 811 Computing in Science Making a Chemical Warfare Treaty Work • J. I. Brauman Oregon Natural History Gets Its Own Museum • DOE Doles Out Millions for High Tech Tools, etc. Disposing of Weapons-Grade Plutonium: E. Lyman, **BOOK REVIEWS** Parascript, reviewed by S. A. Nadler • The Merging F. Berkhout, H. Feiveson • Fisheries Management: W. Aron, D. Fluharty, D. McCaughran, J. F. Roos of the Senses, A. J. King . A Continent Revealed, P. Choukroune • Vignettes • Books Received Determining Carcinogenicity: H. Vainio, B. Armstrong, L. Tomatis • Choose Your Poison: **PRODUCTS & MATERIALS** D. Rosenberger

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COVER

895

Structure of a hypothetical protein with four helices (purple cylinders) modeled by the computer program Sculpt, a tool for guessing the structure of existing proteins or for designing new ones. In this special issue, other aspects of computing in science are discussed in

News stories, Perspectives, and Articles beginning on page 841. [Sculpt was developed by Mark Surles, University of California San Diego Supercomputer Center; Jane and Dave Richardson, Duke University; and Fred Brooks, University of North Carolina, Chapel Hill]



# RESEARCH ARTICLE

Metal Ion–Dependent Modulation of the Dynamics of a Designed Protein

T. M. Handel, S. A. Williams, W. F. DeGrado

#### REPORTS |

Reversible Rotation of Antimony Dimers on the Silicon (001) Surface with a Scanning Tunneling Microscope

Y. W. Mo

Multiple Ion Association in Supercritical
Aqueous Solutions of Single Electrolytes
E. H. Oelkers and H. C. Helgeson

Protein Catalysis of the Retinal Subpicosecond Photoisomerization in the Primary Process of Bacteriorhodopsin Photosynthesis

L. Song, M. A. El-Sayed, J. K. Lanyi

Micromachining a Miniaturized Capillary Electrophoresis-Based Chemical Analysis System on a Chip

D. J. Harrison, K. Fluri, K. Seiler, Z. Fan, C. S. Effenhauser, A. Manz

High-Density Nanosecond Charge
Trapping in Thin Films of the
Photoconductor ZnODEP
C.-y. Liu, H.-I. Pan, M. A. Fox, A. J. Bard

Electron Diffraction and Imaging of
Uncompressed Monolayers of Amphiphilic
Molecules on Vitreous and Hexagonal Ice

J. Majewski, L. Margulis, D. Jacquemain, F. Leveiller, C. Böhm, T. Arad, Y. Talmon, M. Lahav, L. Leiserowitz

Early and Late Alkali Igneous Pulses and a 902 High-<sup>3</sup>He Plume Origin for the Deccan Flood Basalts

A. R. Basu, P. R. Renne, D. K. DasGupta, F. Teichmann, R. J. Poreda

Influence of Productivity on the Stability of Real and Model Ecosystems
J. C. Moore, P. C. de Ruiter, H. W. Hunt

Determinants of Binding-Site Specificity
Among Yeast C<sub>6</sub> Zinc Cluster Proteins
R. J. Reece and M. Ptashne

Development of Mature CD8<sup>+</sup> 911
Thymocytes: Selection Rather Than
Instruction?

J. P. M. van Meerwijk and R. N. Germain

C. G. Hall, J. Sancho, C. Terhorst

Reconstitution of T Cell Receptor

ζ-Mediated Calcium Mobilization in
Nonlymphoid Cells

T Cell Development in Mice That Lack the ζ Chain of the T Cell Antigen Receptor Complex

P. E. Love, E. W. Shores, M. D. Johnson, M. L. Tremblay, E. J. Lee, A. Grinberg, S. P. Huang, A. Singer, H. Westphal

Gene Dose of Apolipoprotein E Type 4 2 921 Allele and the Risk of Alzheimer's Disease in Late Onset Families

E. H. Corder, A. M. Saunders, W. J. Strittmatter, D. E. Schmechel, P. C. Gaskell, G. W. Small, A. D. Roses, J. L. Haines, M. A. Pericak-Vance

# **TECHNICAL COMMENTS**

Composition Limits of Fe<sub>x</sub>O and the Earth's Lower Mantle

R. Jeanloz and R. M. Hazen; C. McCammon

Simplifications of a Self-Replication Model

Indicates accompanying feature

igitoda iioop



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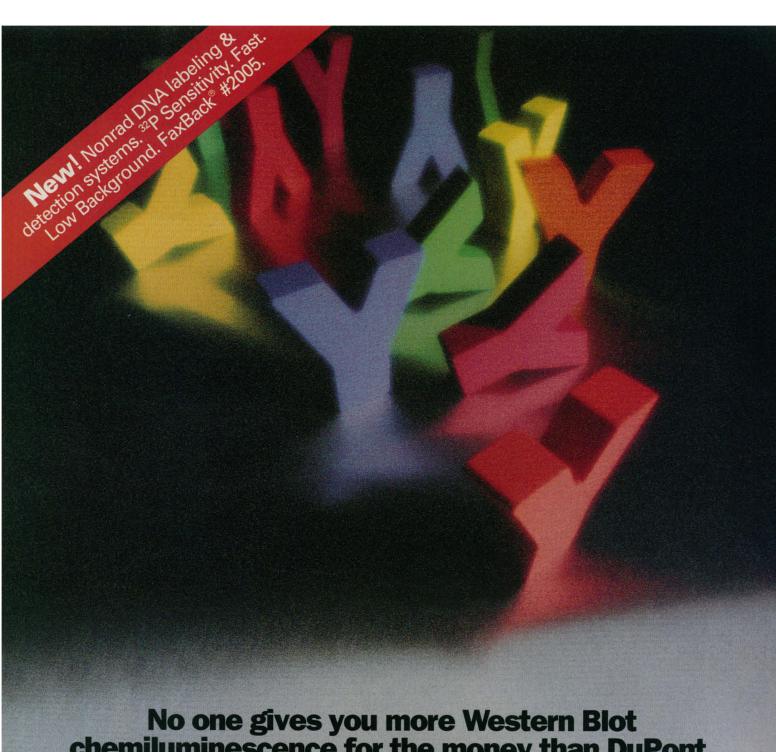
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edited by PHIL SZUROMI

## Going native

Inclusion of metal binding sites into a simplified, designed protein has led to some insights into how specific interactions can affect protein stability. The designed protein  $\alpha 4$  is a fourhelix bundle that forms a molten globule structure that is stabilized by hydrophobic interactions. Handel et al. (p. 879) found that introducing two zinc binding sites into the protein destabilized the molten globule state. Addition of zinc ions stabilized the ground-state structure so that it more nearly resembled a native protein.

## **Dumbbell exercises**

Many types of irreversible surface modifications have been made with the scanning tunneling microscope (STM) as a form of writing. Mo (p. 886) describes the rotation of antimony dimers on the silicon (001) surface, a reversible surface modification







that could have memory applications. Scanning with the STM tip exerts an atomic-scale torque that rotates the dimer between two orthogonal positions, a rotated state and a more stable final state.

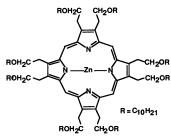
## **Charging ahead**

Spectroscopic studies of bacteriorhodopsin mutants by Song *et al.* (p. 891) indicate that charged residues are critical for maintaining a fast rate of retinal photoisomerization. Replacing to the charged residues in the retinal cavity (Arg<sup>82</sup>, Asp<sup>85</sup>, and Asp<sup>212</sup>) reduced the photoisomerization rate. Isomerization about bonds

## Charge trapping and data storage

High-density, high-speed information storage can be achieved with a device that uses a photoconductive material to record data. Liu *et al.* (p. 897) sandwiched a zinc porphyrin compound between sheets of the transparent conductor indium-tin-oxide. Shining light on

this material creates electronhole pairs; a bias voltage applied between the conducting sheets causes them to migrate. In this way a potential difference can be frozen into the material that can serve as a long-term, nonvolatile memory. The voltages can be retrieved on a subnanosecond time



scale by shining light on the material under short-circuit conditions. No voltage is applied during readout, so the traps are depleted slowly and can be accessed hundreds of times. Single elements as small as 0.2 micrometer can be obtained readily, which translates into a storage density of 3 gigabytes per square centimeter.

other than the double bond between carbon atoms 13 and 14 is probably favored in the mutants, thus slowing the preferred isomerization pathway.

# Miniature instruments under glass

A chemical analysis system has been fabricated on a small glass chip. Harrison et al. (p. 895) performed amino acid analysis with a capillary electrophoresis instrument that was micromachined on a centimeter-sized glass wafer. Moving parts are not needed, as electroosmotic flow was used to pump fluids through capillaries with cross sections on the order of tens of micrometers. Such small dimensions allowed separations to be performed in tens of seconds. Liquid flow at junctions could be directed by selectively applying voltages, so sample manipulations such as dilution could be performed. The amino acids were fluorescently labeled and detected by laser excitation. This miniaturized system combines the essential elements of an analytical system, including sample pretreatment, injection, separation, and detection.

# **Dating the flood**

One model for the origin of flood basalts, which represent the most voluminous volcanism on Earth, is that they are related to the ascent of a plume of hot material in the mantle, perhaps from the lower mantle. To evaluate this model and the evolution of melting as the presumed plume nears the surface, Basu et al. (p. 902) studied igneous rocks that mark the inception and demise of the Deccan flood basalts, which cover a large part of west-central India and were predominately erupted about 65 million years ago, near the time of the Cretaceous-Tertiary boundary. The 40Ar-39Ar dates for the early intrusions show that Deccan magmatism actually began about 3.5 million years earlier. The earliest and latest magmas were particularly alkalic, and the early magmas have high <sup>3</sup>He/<sup>4</sup>He ratios, similar to certain ocean island

basalts, whereas the latest magmas show clear evidence of crustal contamination.

# T cells and $\zeta$ chains

Signal transduction by the T cell receptor (TCR) is accomplished by invariant cytoplasmic subunits such as the  $\zeta$  chain. Two reports examine the role of ζ chain in T cell signaling and development. Hall et al. (p. 915), working in COS-1 cells, show that coexpression of the  $\zeta$  chain with the protein tyrosine kinase p59fynT is necessary and sufficient to phosphorylate the  $\zeta$  chain and phospholipase C (PLC)yl and to release intracellular calcium stores. This result suggests that p59fynT may directly link the  $\zeta$  chain to the PLC- $\gamma$ 1 activation pathway. Love et al. (p. 918) used gene targeting to generate mice that lacked the  $\zeta$ chain. The  $\zeta$  chain is needed to transport the assembled TCR to the cell surface. A substantial decrease in the number of single positive CD4<sup>+</sup> and CD8<sup>+</sup> cells was seen, but unexpectedly some single positive cells were still detected in the periphery. Normal T cell expression was restored with a  $\zeta$  transgene.

#### Hammering iron

Knowledge of the oxidation state of the lower mantle and whether the mantle is in equilibrium with metallic iron that forms Earth's core bears directly on models of the origin or the core and subsequent evolution of the mantle. Recent experimental data on the pressure dependence of the Fe<sup>3+</sup> content of Fe<sub>x</sub>O in the mantle and their implications for these questions are considered in a technical comment by Jeanloz and Hazen (p. 923) and reply by McCammon (p. 924).

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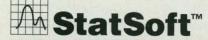
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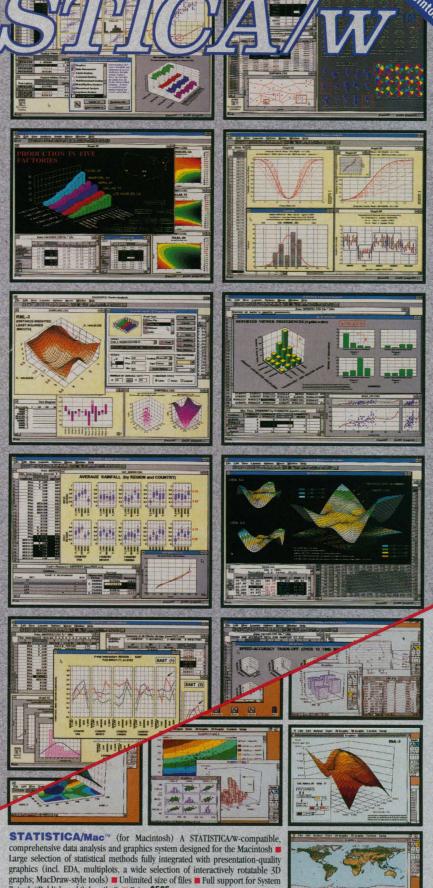
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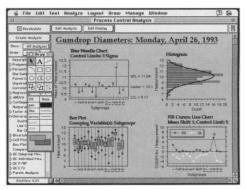
This summer, we're introducing QC Tools. QC Tools adds a "no compromise" set of QC statistics and graphs to StatView. With QC Tools, you can do Xbar, R, S, I, MR, CUSUM (with FIR), Capability, p/np, c/u, Pareto analysis, as well as ANOVA, box plots, normality plots, and all the other analyses that come standard with StatView.

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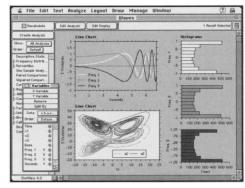
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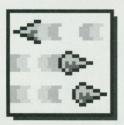
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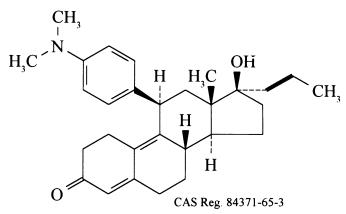
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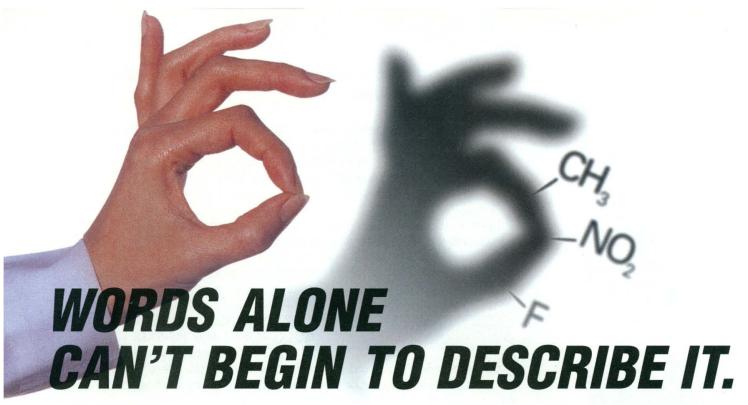
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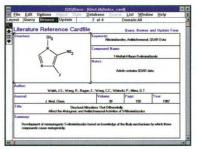
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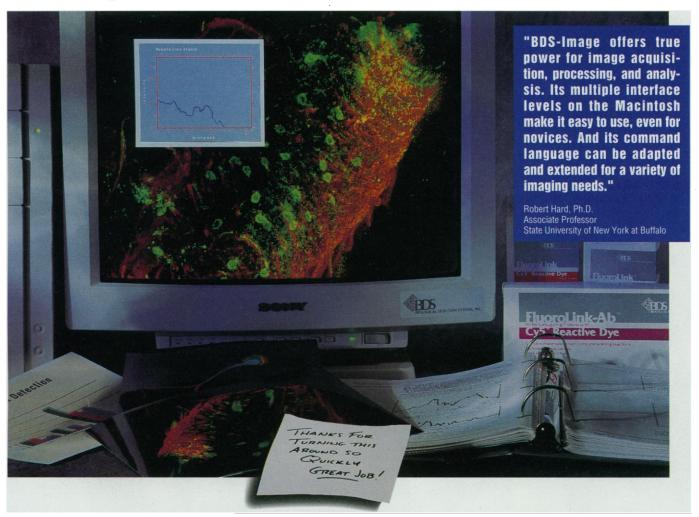
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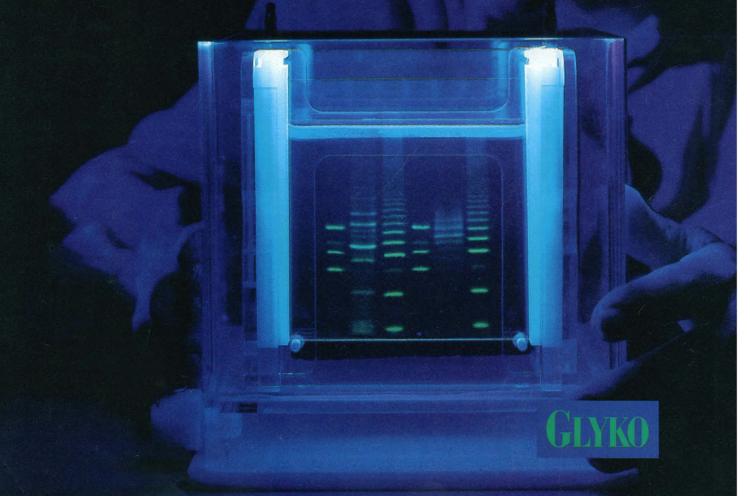
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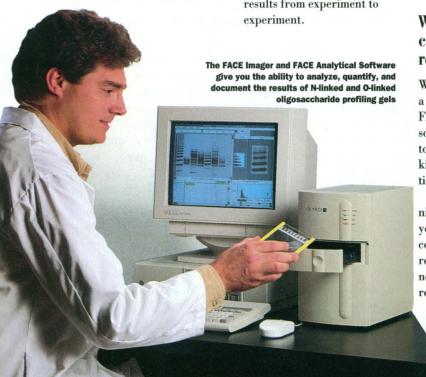
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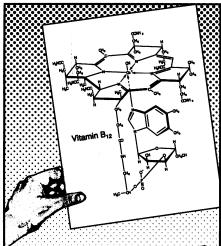
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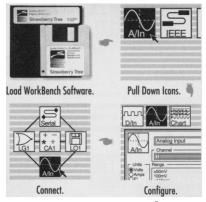
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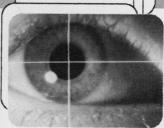
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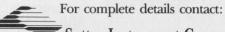
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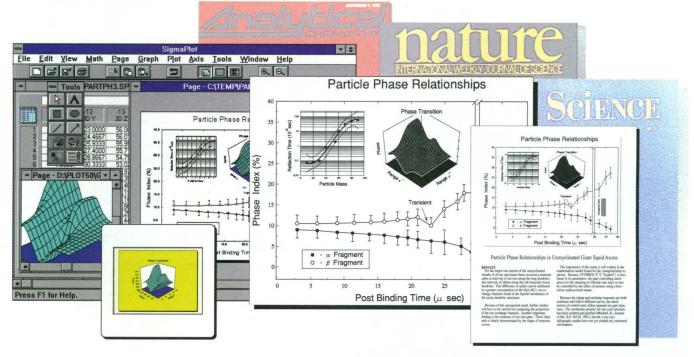
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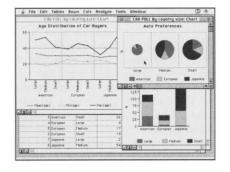


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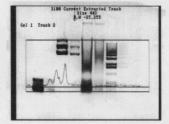
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