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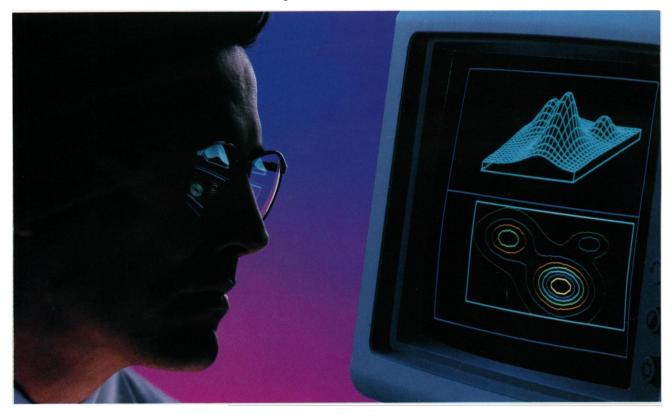
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COVER In Greek mythology, the chimera was a creature with characteristics of several animals. The construction of chimeric molecules from diverse pieces of DNA is now a cornerstone of recombinant DNA technology. See pages 1223 to 1263. [Cover illustration by Sharon Wolfgang, Washington, D.C.]

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Stephen Kress's innovative techniques successfully repopulated near-extinct colonies of birds on Maine islands and will soon be

employed in attempted recolonizations in the south of Japan.

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ticular species, and will complete a book on the subject.



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## Religious teaching and environmental education.



Nancy Lee Nash will glean teachings about nature from Buddhist writings and incorporate them into an educational syllabus to be used

in Thailand colleges.

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Johan Gjefsen Reinhard will explore the peaks and lakes of the Peruvian Andes to determine the role of those ancient religious sites in the

formation of pre-Colombian society.

*The Spirit of Enterprise, The 1987 Rolex Awards,* a book about the five laureates' projects, as well as 238 other outstanding proposals, is available in bookstores and from the publisher, Van Nostrand Reinhold.

Additional information about the awards is available from The Secretariat, The Rolex Awards for Enterprise, P.O. Box 178, 1211 Geneva 26, Switzerland.

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## **Disease diagnosis**

s a result of developments in recombinant DNA technology, growing numbers of diseasesgenetic, infectious, and neoplastic-are able to be diagnosed (page 1223). A review of how gene probes, restriction fragment length polymorphisms, and other recombinant tools are aiding clinical diagnoses is presented by Caskey in this special issue on developments in recombinant DNA technology (see also page 1157 and pages 1223 to 1268). Some genetic diseases can now be diagnosed in the fetus and the inheritance of late-onset diseases can be tracked in families; for some of these, thorny ethical and legal considerations remain to be addressed, such as the responsibility of insurance companies to asymptomatic individuals who test positive for a late-onset disease and to normal individuals who are critical to a linkage analysis. Cancer diagnosis is improving and, as additional translocations in oncogenes are associated with specific cancers, is likely to increasingly rely on recombinant techniques. So is the diagnosis of infectious diseases, particularly viral diseases for which rapid and reliable methods were not formerly available. Accurate disease diagnosis is a crucial first step in treatment and prevention.

## **Enzyme tinkering**

→ ITE-DIRECTED mutagenesis is a recombinant technique that has been used for altering single amino acids in enzymes in a controlled fashion; similar precision cannot be achieved through chemical modification procedures (page 1252). Knowles summarizes results from some experiments in which single amino acids in or near an enzyme's active site have been replaced by another naturally occurring amino acid (at some future time, unusual substituents or unusual analogs of amino acids may be integrated into enzymes). Changes in single amino acids have caused subtle and dramatic structural alterations; these, in turn, can affect the enzyme's stability to heat and

proteolysis, the specificity of the enzyme for its substrate, the energetics of and steps in catalysis, and the types of intermediates generated in the catalytic reaction. As understanding of enzyme structure and function and their interdependence improves, new strategies may arise for restoring catalytic activity in defe ctive enzymes (in diseases, for example) and for developing designer enzymes that efficiently catalyze useful biochemical reactions.

## Rock magnetic properties

AGNETITE is an important ore of iron; it is one of the most L common and powerful magnetic minerals (page 1287). An understanding of its properties may be key to explaining physical events in the earth's history because magnetite retains information about earlier directions of the earth's magnetic field. Crysta's of magnetite ranging in size up to 1 millimeter were grown by Heider et al. by heating seed crystals in mineralizing solutions at temperatures between 400° and 800°C. Physical properties of these laboratorygrown crystals and of crushed magnetite that had been used previously as a model for natural magnetite were evaluated. The laboratory-grown crystals resembled similarly sized natural magnetite particles found in the earth far better than did the crushed magnetite. These crystals will be useful for elucidating properties of the natural magnetite, including the possible determination of how rocks are magnetized. Magnetite is used in permanent magnets and has other valuable applications in the preparation of magnetic recording devices.

## Bridging communication gaps

Can take place through groups of channels that form junctions, called gap junctions; this form of intercellular communication is mediated by electric and chemical signals (page

1290). A 28,000-dalton gap junction protein from rat liver is one constituent that participates in the formation of functional channels for intercellular communication. Dahl et al. inserted complementary DNA for this protein into an oocyte expression system. Channels of two types formed between pairs of oocytes, and they were readily distinguishable because they had different electric properties and responded differently to ions. One type was endogenous to pairs of oocytes; the other type, which formed as a result of the injected complementary DNA, had properties like those of channels that form between cells in liver, the parent tissue. This oocyte system should be useful for studying the mechanism of channel formation and other facets of the intercellular communication process.

## Viral probe for sialic acid derivative

EMAGGLUTININS on the surface of influenza C viruses have a special affinity for the sialic acid derivative 9-O-acetyl-N-acetylneuraminic acid that resides on many cell surfaces; Muchmore and Varki capitalized on this relation to study properties of the hemagglutinin molecules and to develop a specific probe for this sialic acid derivative (page 1293). The hemagglutinin molecules have an esterase activity that typically destroys the 9-Oacetyl substituent of the sialic acid derivative. Therefore, esterase activity was specifically inhibited. The hemagglutination properties of the protein and the fusion properties of the virus were unaltered, and the viruses bound specifically and irreversibly to mammalian cells that had the derivatives of interest. Only the infectivity of the virus was decreased. Sialic acid derivatives are widely distributed amino sugars found in both bacterial and animal cells; they are thought to participate in a variety of processes involving surface-to-surface interactions, such as cellular homing. The development of this probe will help in understanding their distribution and functions.

THIS WEEK IN SCIENCE 1155

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# Science

5 JUNE 1987 VOLUME 236 NUMBER 4805

## Frontiers in Recombinant DNA

cartoon in the New Yorker shows an aristocratic dowager in a Chippendale chair looking up from her newspaper and saying to her patrician-looking husband, "Dear, are they tampering with our genes?" Such vague worry that the young Turks of DNA may not only inadvertently create pestilence, war, and famine but may even invade the regions of good taste is prevalent in the general public. To the cognoscenti these worries are remote but not absurd; yet they rush on. If there is even the remotest chance of some misuse, should one not slow down and proceed with great caution?

The answer is that, in fact, fuller understanding of the DNA methods has revealed the areas in which caution must be applied as well as the power of this research tool. The sweep of subjects in which recombinant DNA has provided breakthroughs is incredible. In this issue a few of many examples are selected to show how it has been applied to the understanding of learning, development, diagnosis of disease, stimulation of colony growth, generation of new plant foods, design of new catalysts, and the switching on of genes and viruses. Although the applications described here relate to these fundamental biological processes, the potential for practical applications is clear.

Maniatis et al. describe a research area that has moved with lightning speed, but still leaves many mysteries. The role of promoters and enhancers in turning on and turning off gene expression and the apparent crazy quilt pattern of upstream and downstream elements could never have been unraveled by classical genetics. Gehring illuminates a phenomenon that was initially revealed by classical genetics: the homeotic genes, mutations in which cause legs to grow out of the heads of flies. By recombinant DNA studies the homeo gene has led to new understanding of morphological development. Cocking and Davey describe the breach of a new barrier-the cereals-previously resistant to the techniques that allowed genetic engineering of other types of plants. Clark and Kamen discuss the application of recombinant DNA techniques to the elusive elements that stimulate cell growth, the colonystimulating factors. Caskey outlines recombinant DNA procedures that can lead to the identification of molecular defects accounting for heritable diseases and acquired neoplasia in Homo sapiens, and Black et al. describe the application to one of the most difficult experimental challenges, the understanding of the molecular events in learning and memory. Knowles tells of the application of genetic engineering to the vital catalysts that control all dynamic biological processes-the enzymes.

The intellectual excitement of rapid progress in some of the most vital problems in biology is, by itself, an enormous stimulus; but in each case practical applications are almost at hand. The prospect of an age of designer catalysts, of quick medical diagnosis of genetic defects, of new cereals for starving millions, and of brain therapy in neurological diseases is another spur to progress in these already rapidly advancing areas.

Biologists asked for a moratorium to assess risks in the initial days of recombinant DNA. How is it, now that the tools appear to be more powerful than was expected then, that there can be confidence that these experiments no longer threaten humankind? The answer has two parts. First, much is now known about the events that lead to uncontrolled growth of cells and about the safe application of recombinant DNA. Second, experiments have revealed that genetic engineering has been occurring in nature for eons without catastrophic consequences. Recombination in vivo results in events such as attachment of the hind end of one protein to the front end of another. Nature's genetic engineering through selection is much slower than modern laboratory manipulation, but it has been going on for billions of years. That fact can not only induce some humility in molecular biologists but also ease some fears of onlookers.

Ecological and moral dilemmas created by these new technologies are appreciable and will require new ideas. Tampering with the mind is generally considered to be bad, but should genetic engineering to alleviate Alzheimer's disease be outlawed? To feed starving populations is desirable, but if new crops help add a billion people to a crowded globe, is that necessarily good? The powerful new methods are here. Applying them may well require the use of brain enhancers.—DANIEL E. KOSHLAND, JR.

## **BIOSYSTEMS UPDATE**

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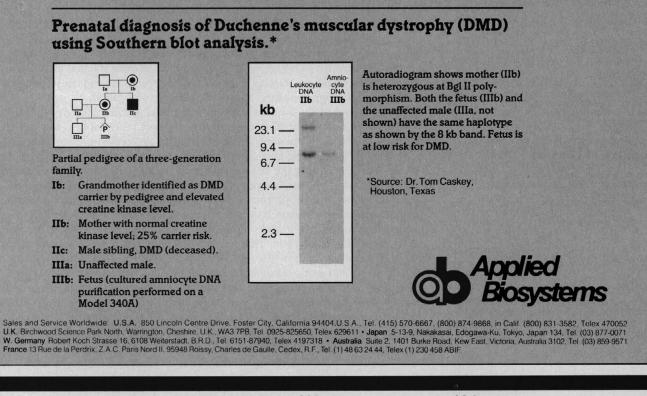
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by 46 countries (Argentina and Chile not among them), including all of the major maritime nations. It imposes strict rules on tanker operators to prevent oil pollution. These rules not only prohibit the type of dumping that Boersma highlights but also incorporate vessel design and equipment requirements. These include segregated ballast tanks, crude oil washing, and other features that minimize the amount of oily waste that is generated aboard a tanker. The rules also require the installation of oil-inwater monitors that measure and record the oil content of ballast that is discharged by a tanker. In fact, these monitors are designed to stop the discharge of ballast automatically should oil concentration in the water exceed strict limits.

Thus, there are international regulations that prohibit tankers from "dumping" oil into the sea, and, contrary to Boersma's assertion, most tankers follow these rules.

In dealing with the "oily penguin problem," it appears first necessary to consider possible sources of contamination, for example, spills occurring during drilling and producing operations in the Tierra del Fuego area, indiscriminate discharges of oily engine room bilge wastes from any type of vessel operating in the area, and discharge of oily ballast water from tankers that may operate in the area and not conform to international operating standards.

Second, it will be necessary to heighten the level of awareness of government, industry, and the public with respect to the nature of the problem. Oil-producing and transportation operations are routinely conducted in other environmentally sensitive areas without adverse effects on those areas. Such environmentally acceptable operations should be sought and are achievable in Tierra del Fuego.

> T. S. WYMAN Government and Public Affairs Division, Chevron Shipping Company, 555 Market Street, San Francisco, CA 94105-2870

#### Man's Role in Space

Peter M. Banks and David C. Black (Perspective, 17 Apr., p. 244) write optimistically and philosophically about what might be gained through manned research facilities in space. James A. Van Allen has already written factually (1) about what has been gained through unmanned spacecraft and what further surely could be gained. One might add that that work (in which I participated some three decades ago) (2) led to new and largely

unforeseen technological applications of unmanned spacecraft as communication satellites. Since then earth observation satellites of various sorts and navigation satellites have assumed great importance.

Our present humiliating position is that exaggerated emphasis on man-in-space has left us, temporarily, we hope, without any domestic means for launching either scientific or technological spacecraft.

Man-in-space is an old dream which I exploited in science fiction (3) at a time when no one had thought of the new capabilities we have seen in unmanned spacecraft. The old dream of man-in-space, magnificently realized in Apollo, has been extended in the Shuttle, with an emphasis that has had disastrous consequences for our exploration and exploitation of space.

Surely, the exploration of man's role in space is worthy of continued and intense investigation. The survivability of man-inspace, to which Banks and Black give no emphasis, calls for continued and intense investigation, but not at the cost of scientific and technological benefits which we know we could attain through unmanned spacecraft-if we had the resources to construct and launch them.

> JOHN R. PIERCE Center for Computer Research in Music and Acoustics and Department of Music, Stanford University, Stanford, CA 94305

#### REFERENCES

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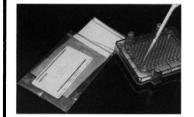
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 J. R. Pierce, The Beginnings of Satellite Communications (San Francisco Press, San Francisco, CA, 1968).
 \_\_\_\_\_\_, "Relics of the earth," Sci. Wonderstories, March 1930, p. 894.

#### Neutrino Shield?

M. Mitchell Waldrop (Research News, 1 May, p. 523) quotes a theorist as saying that the neutrino flux alone from supernova 1987A would have killed an "unprotected human" out to nearly a billion kilometers. I am curious about what sort of protection the theorist had in mind. Is this one of the theorists behind the Strategic Defense Initiative?

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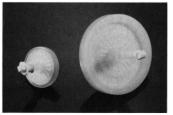
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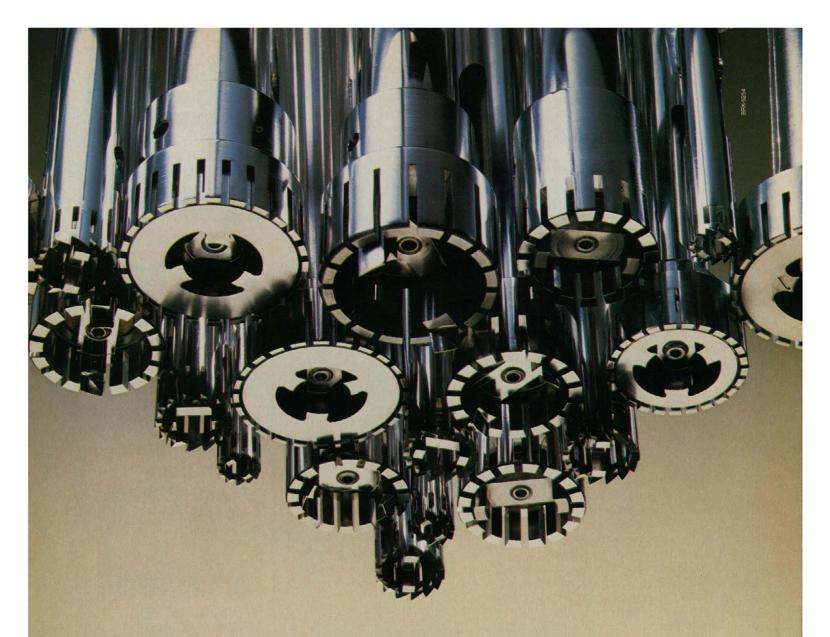
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Erratum: In Constance Holden's article "Adjusting to an aging population" (News & Comment, 15 May, p. 772), Secretary of Labor William E. Brock III is incor-rectly described as "Commerce Secretary."



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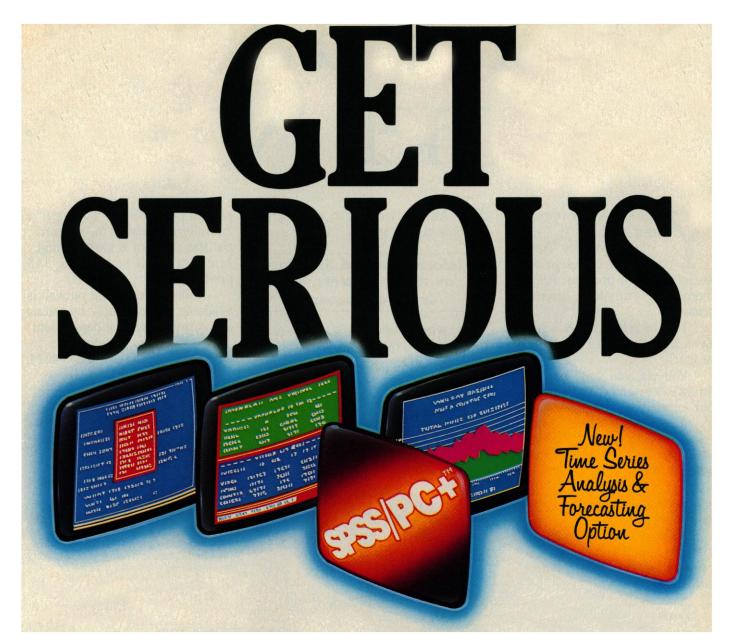
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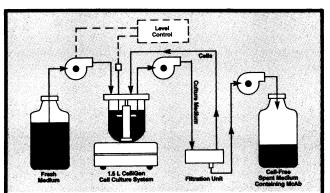
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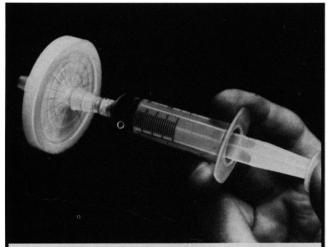
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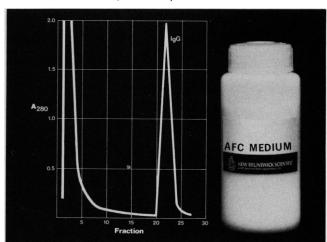
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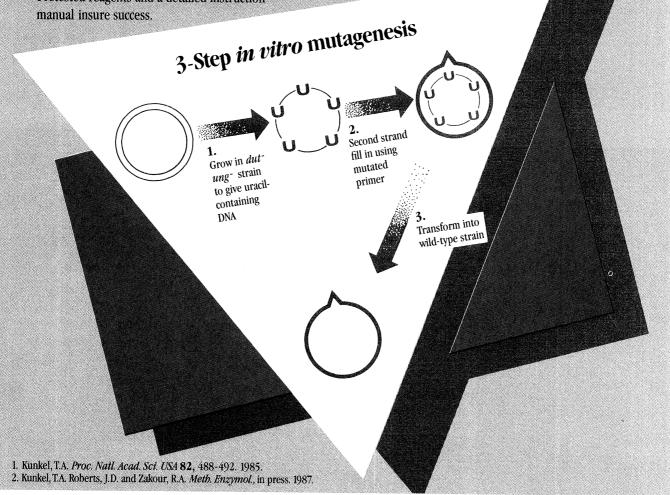
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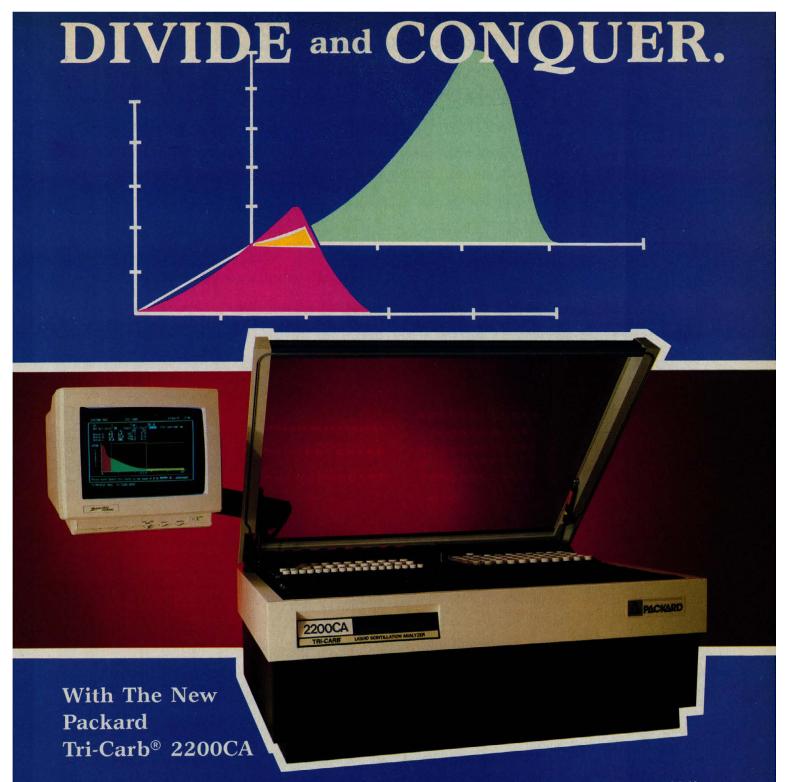
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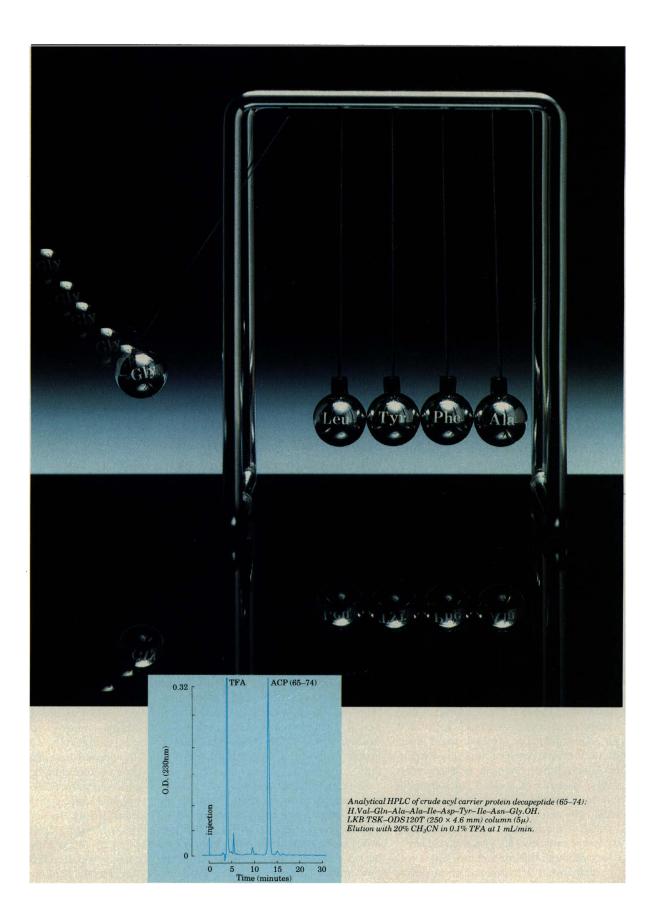
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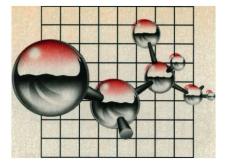
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#### August 9-13, 1987 San Diego, California



First Symposium of the Protein Society

#### The Protein Society

A scholarly society of scientists interested in the structure and function of proteins. The purpose of the society is fundamental; to foster discussion and interaction among scientists interested in all aspects of protein structure and function. The society already has over 700 members covering a broad spectrum of the different disciplines bearing on protein structure and function.

Interaction among many types of investigators utilizing the myriad of approaches and techniques to probe protein molecules is essential. The Protein Society ensures these interactions will occur by providing regular opportunities for protein scientists to meet and exchange ideas and findings.

#### **Full Membership**

Individuals holding a doctoral degree or equivalent experience with a demonstrated interest in the structure and function of proteins. \$50, 2 yrs. Associate Membership

Individuals with an interest in proteins not qualifying for full membership. \$25, 2 yrs.

#### **Corporate Membership**

Commercial/Industrial companies interested in fostering a synergistic involvement of industry in protein chemistry. \$1500, 2 yrs.

#### The Second Symposium-

San Diego, August 9-13, 1987

The first biannual symposium of the Protein Society.

#### Location

The second San Diego symposium is the first opportunity for the membership of the Protein Society to meet and interact. The program of this first meeting is designed to attract a variety of individuals representing all the differing aspects of protein research.

The program committee under the chairmanship of Professor Jack Richards of CalTech is made up of expertise in all disciplines of protein structure and function enforcing the purpose of the Protein Society.

The symposium program is made up of plenary sessions focusing on major impact topics of general interest.

Dr. Eisenberg of UCLA and Dr. Huber of the Max Planck Institute will open the meeting. The meeting will close with plenary lectures from Dr. Würtrich of ETH, Zurich and Dr. Hood of CalTech.

To facilitate interaction among protein chemists from all aspects of the science, the remainder of the program is given over to minisymposia and poster presentations on the following topics:

#### Session Topics and Invited Speakers for San Diego '87

- 1. 3D Structure: Theory Folding and Dynamics. (Baldwin, Goddard, Rose)
- 2. MicroAnalytical Strategies: Structural Analysis and Micro Purification.
  - (Simpson, Kent, Beiman)
- Protein Kinases: Biological 3. Control of Protein Function. (Walsh, Ullrich)
- Protein/Nucleic Acid 4. Interactions. (Altman, McPherson)
- 5 Studies with Synthetic Peptides and Proteins.

(Kaiser, Degrado, Lerner, **Richards**) Photoreceptors as Signal

- 6. Transducers. (Khorana, Zucker)
- 7. Expression, Processing and Secretion.
- (Varshavsky, Beckwith, Wickner) 8 Mutagenic Studies of Protein
- Function: Design and Selection. (Schimmel, Shortle, Winters) 9. New Developments in
- Instrumentation, Chemicals and Accessories for the Protein Chemistry Lab

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#### Stein and Moore Award and Symposium

Part of the program will be used to honor the recipient of the first Stein and Moore award in the form of a half day symposium. The subject and the speakers taking part will be selected by the recipient of the award. Announcement of the awardee and the subject to be presented will be made in the spring of 1987. The award, to be presented biannually by the Protein Society will recognize significant achievement in protein research. In addition to the symposium, the awardee will receive \$5000 and a commemorative plaque presented at a banquet which will feature Dan Koshland as an after dinner speaker.

#### Sample Workshop

All registrants will be invited to participate in a workshop designed to demonstrate the variety of modern techniques available for the study of protein structure and function.

#### **Scientific Exhibition**

Leading manufacturers of scientific instrumentation and techniques relevant to protein research will be represented.

#### Symposium Registration

\$275-Full Members \$150—Associate Members \$330—Non Members \$180—Students

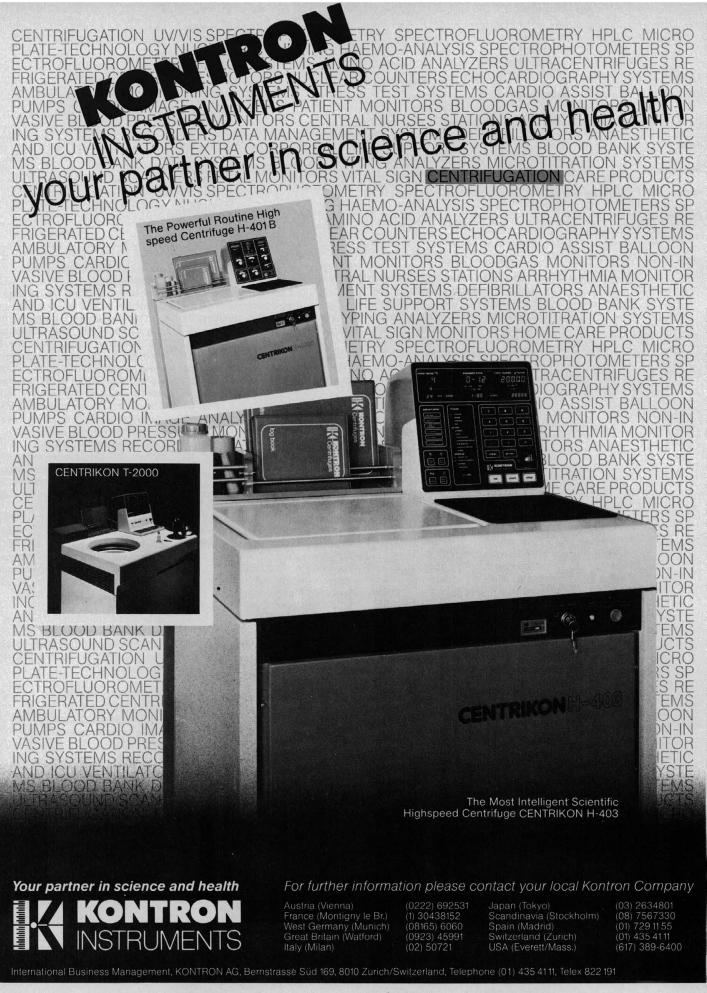
#### Symposium Announcement and Call for Papers went out **January '87**

If you are interested in becoming a member of the Protein Society or attending the San Diego Symposium of the Protein Society, please circle the appropriate number on the reader information card, or contact the Protein Society directly:

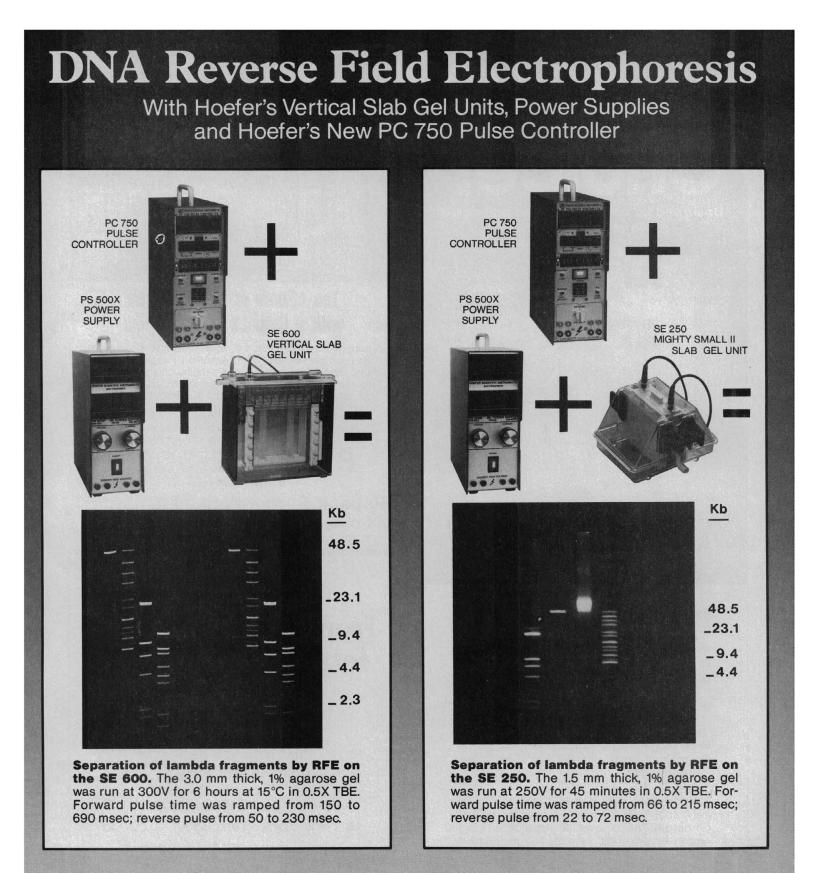
The Protein Society Shirley E. Schlessinger Symposium Manager 400 E. Randolph, Suite 105 Chicago, Illinois 60601 (312) 527-2011

First Symposium

American Protein Chemists September, 1985 San Diego, California Second Symposium The Protein Society August 9-13, 1987

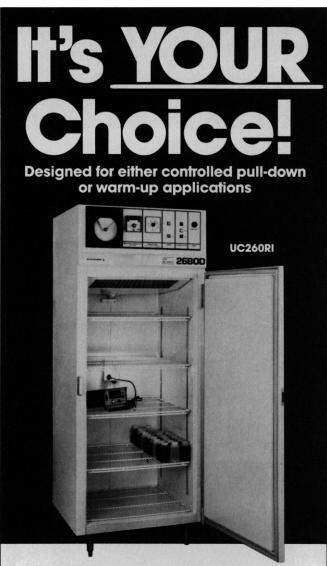


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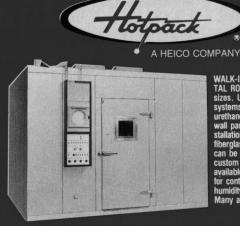


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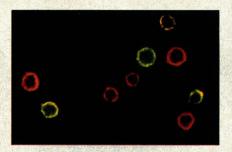


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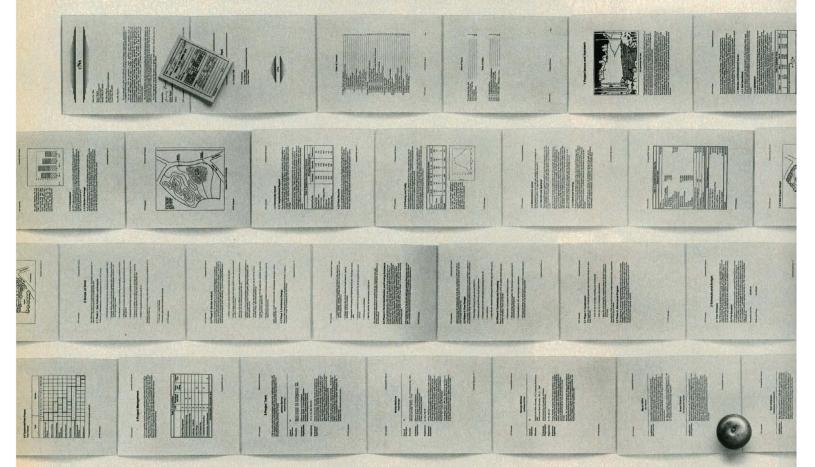
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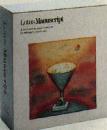
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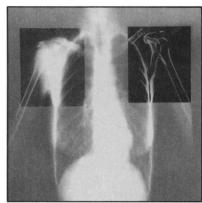
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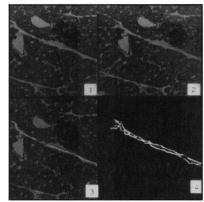


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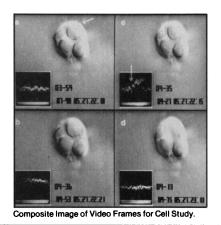
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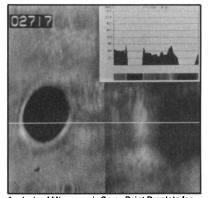
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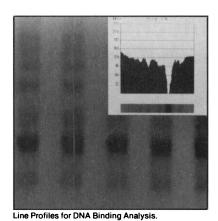
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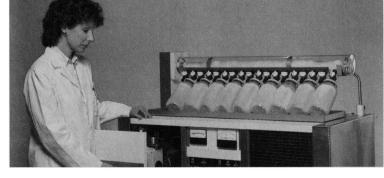
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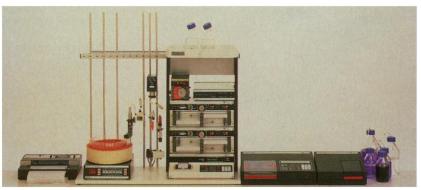


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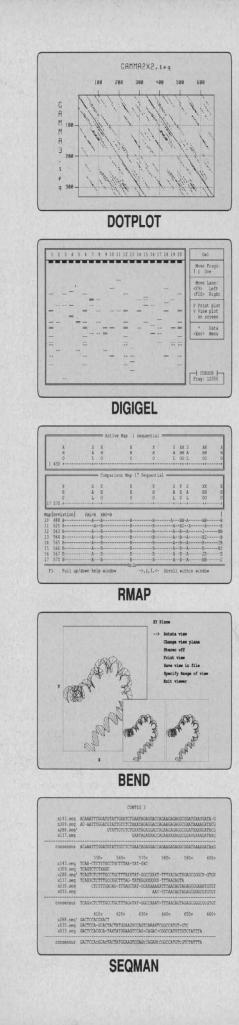
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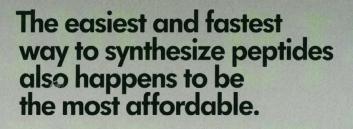
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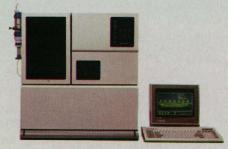
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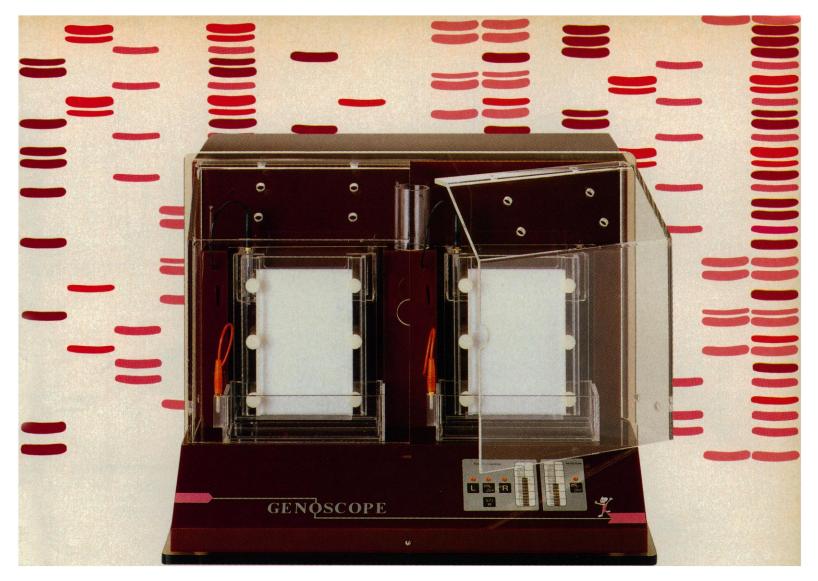
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<sup>1</sup> Aebersold, R.H., Teplow, D.B., Hood, L.E., and Kent, S.B.H. (1986) *J. Biol. Chem.* **261**, 4229-4238.





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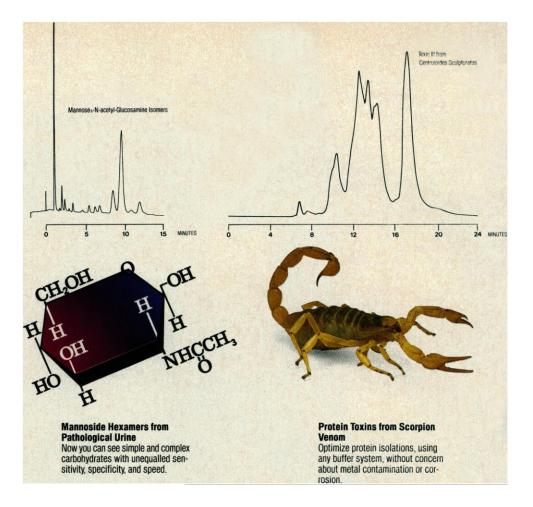
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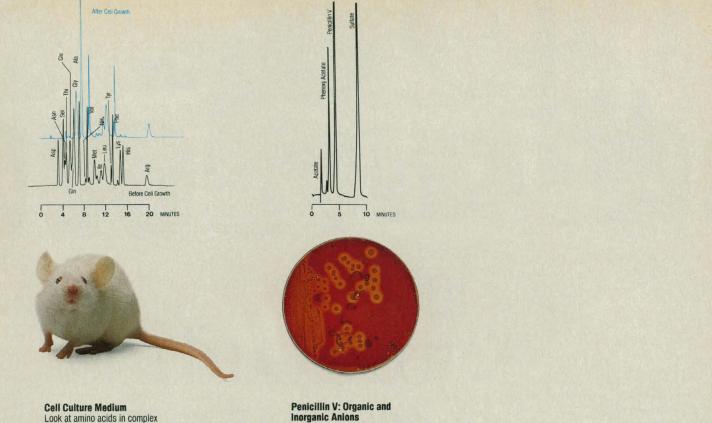


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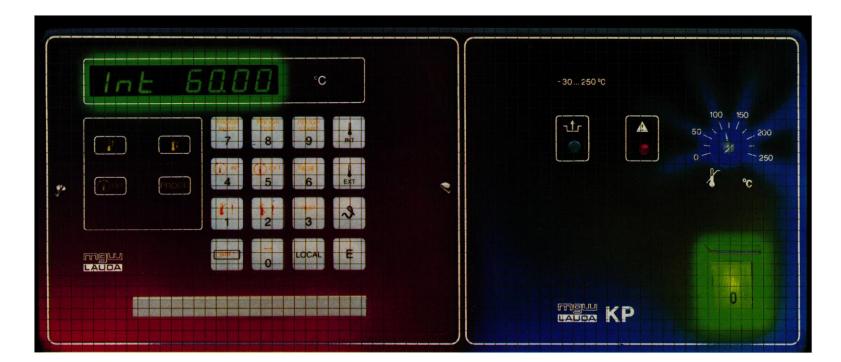
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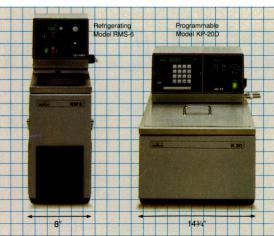


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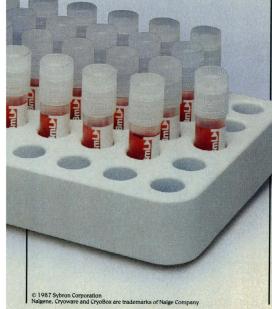
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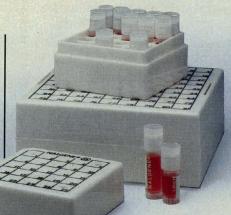
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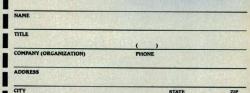
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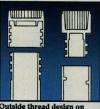
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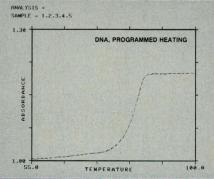
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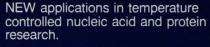
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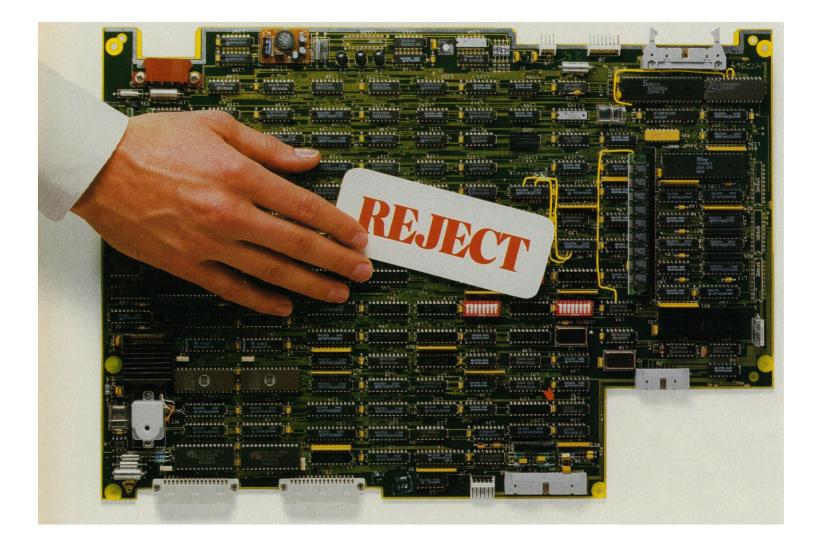
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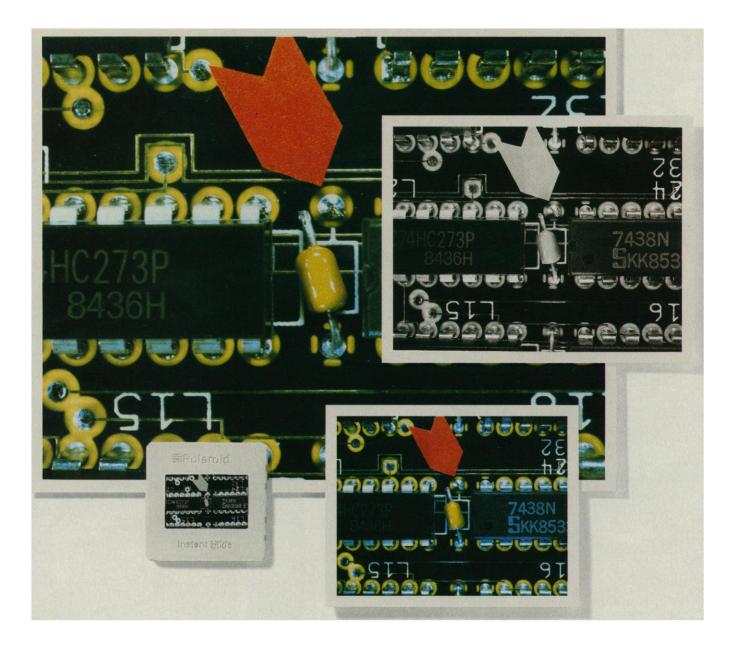


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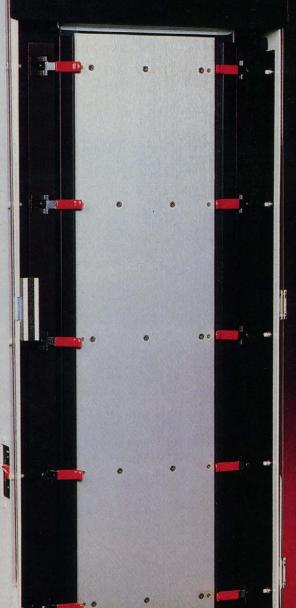
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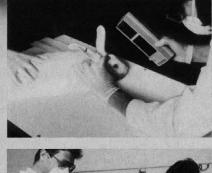
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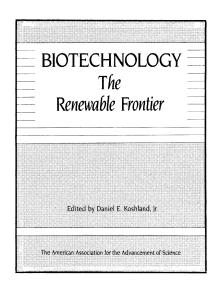
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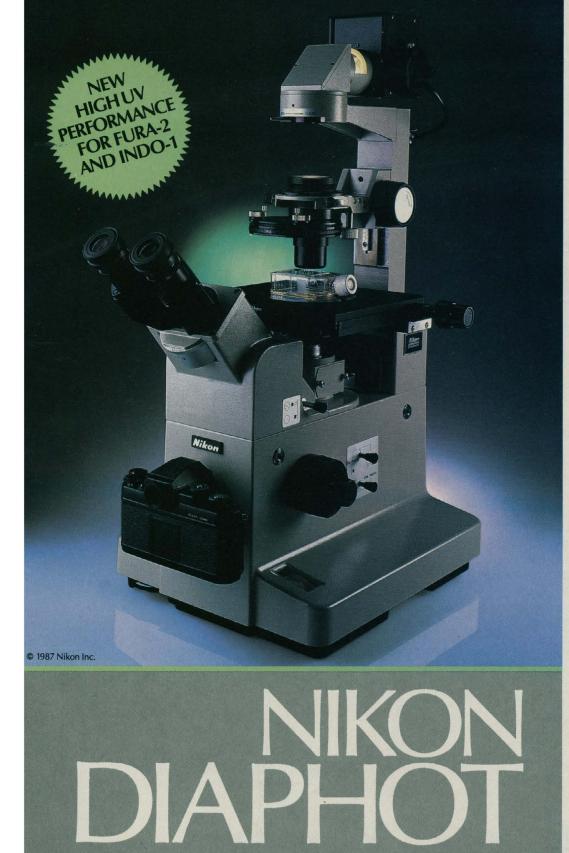
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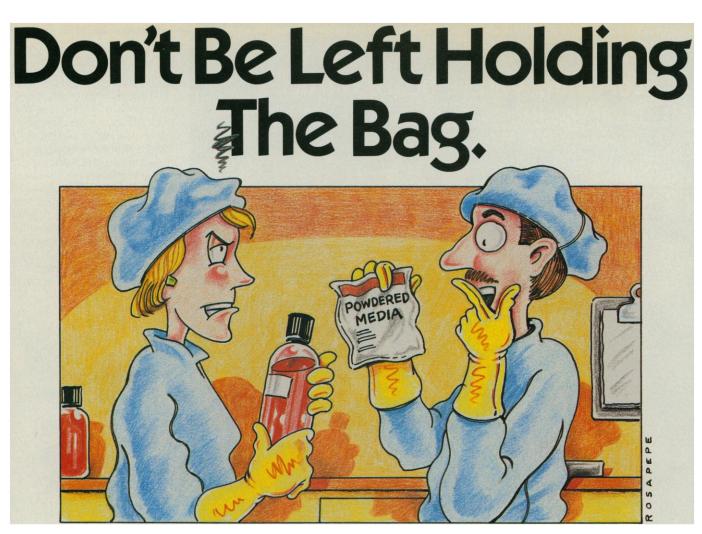


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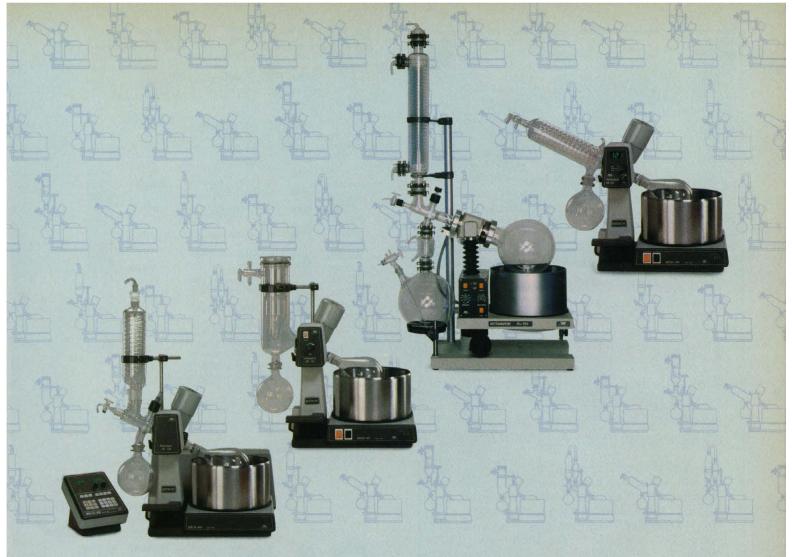
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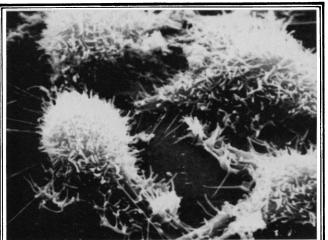
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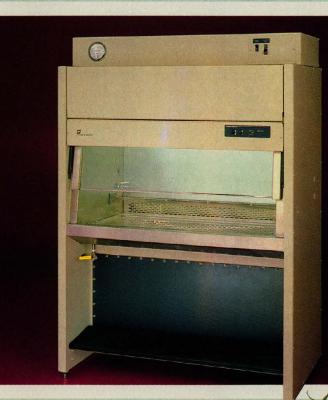
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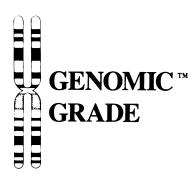


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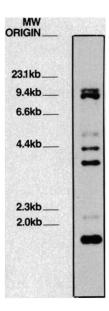


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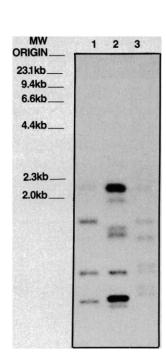
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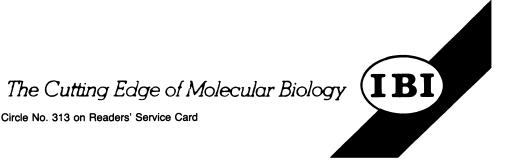


DNA was prepared from three different cultured  $\beta$ -lymphoblastoid cell lines, digested to completion with *Rsa* I (Genomic Grade) and transferred to a nylon filter membrane. A DR $\beta$  clone contained in the plasmid pBR322 was then nick-translated and used as the probe. DR $\beta$  was employed because of its highly polymorphic nature. As is evident in the autoradiogram, no signals are present in the molecular weight range >2.3 Kb, where one would expect to detect plasmid contamination.

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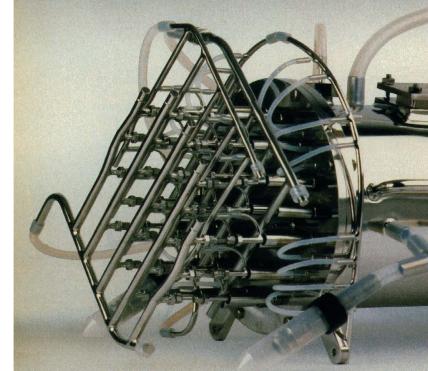
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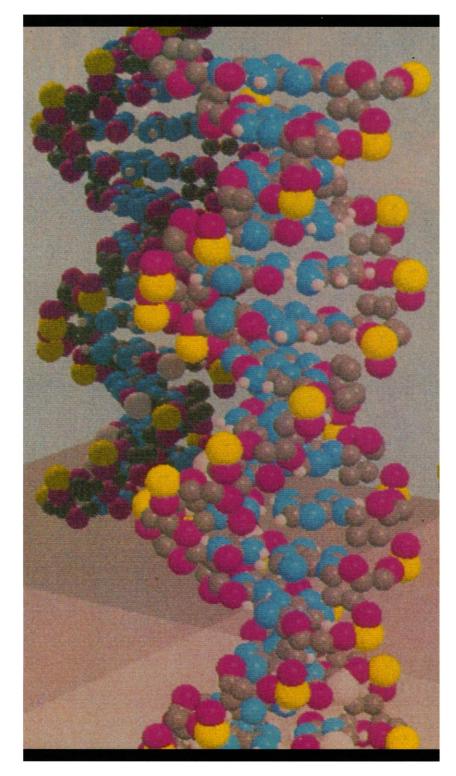
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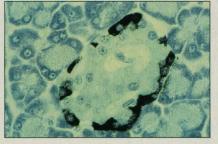
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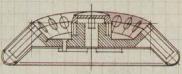
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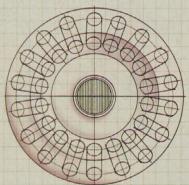
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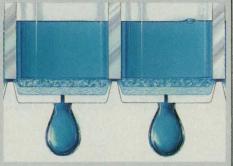
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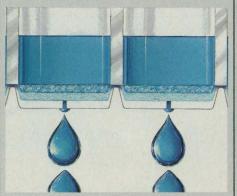
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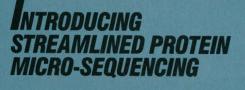
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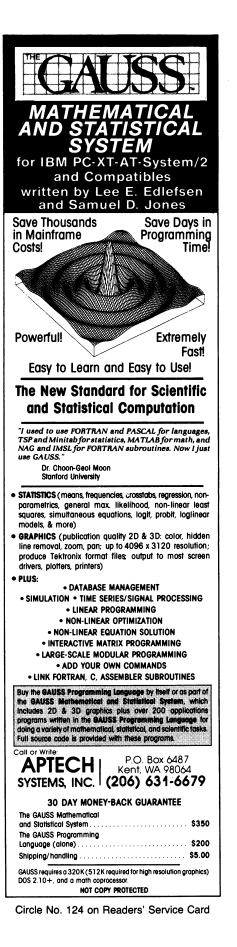
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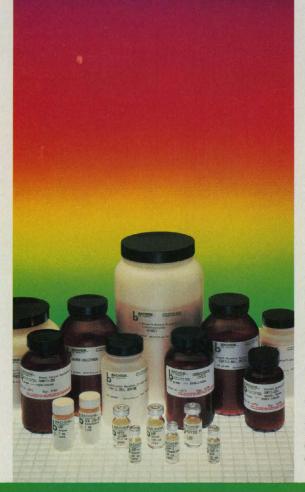
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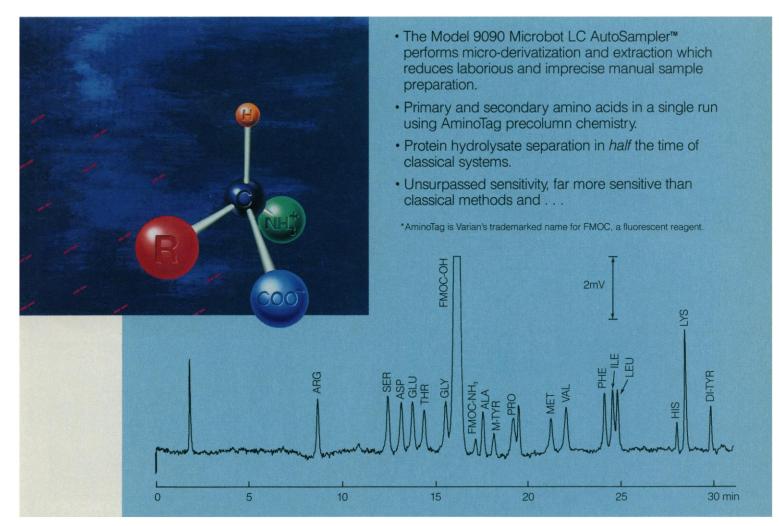
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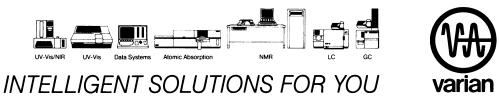


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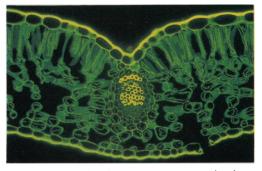
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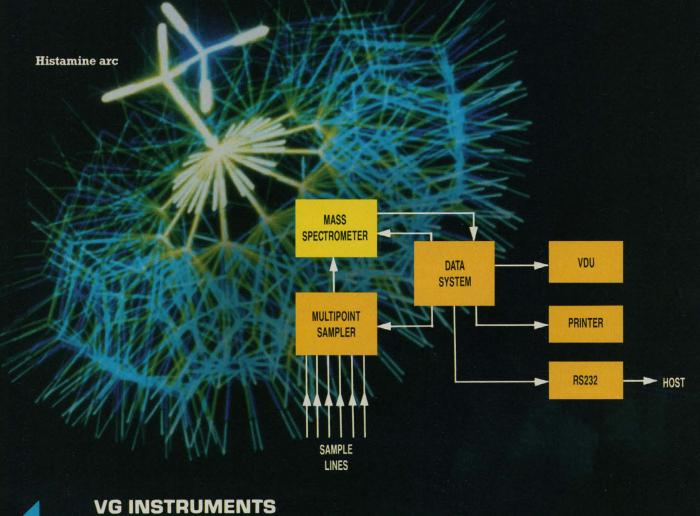
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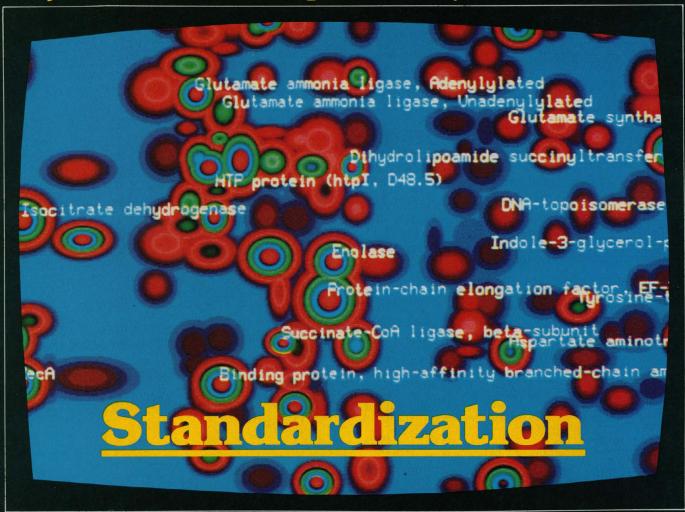


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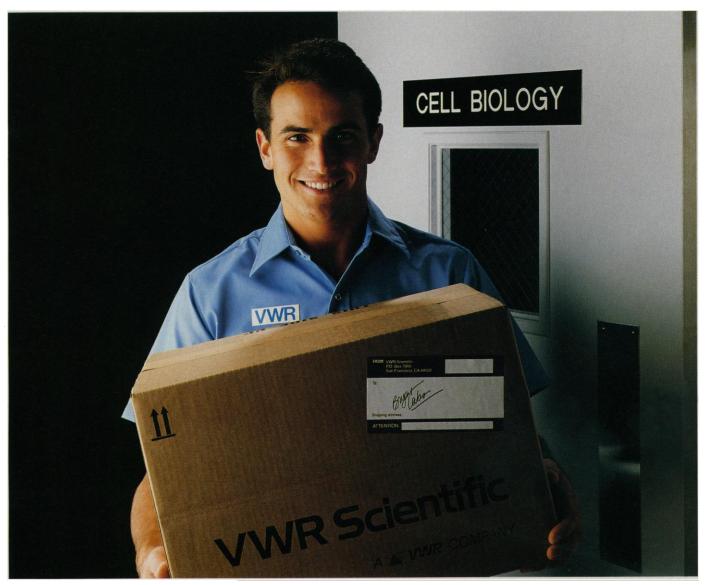
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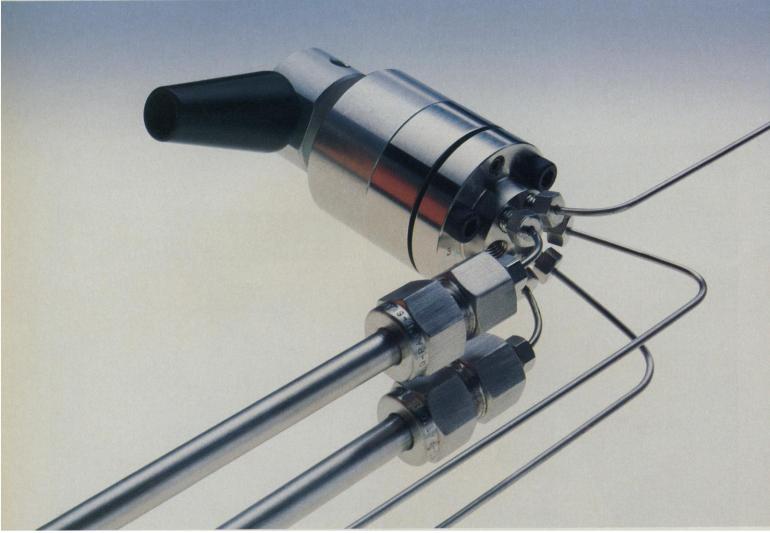
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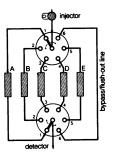


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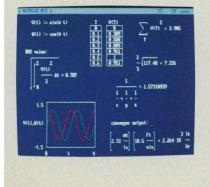
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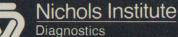
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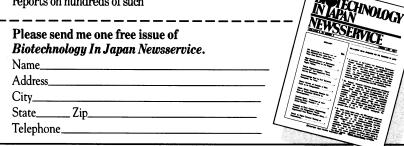
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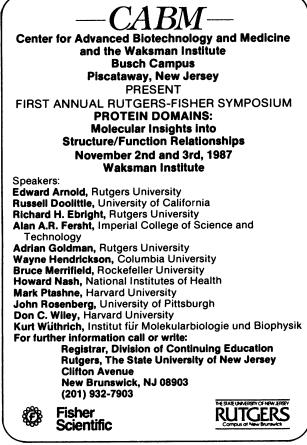
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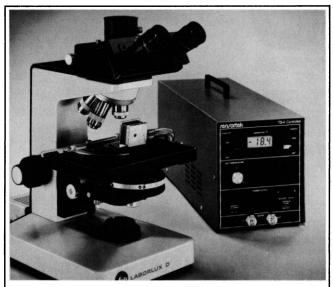
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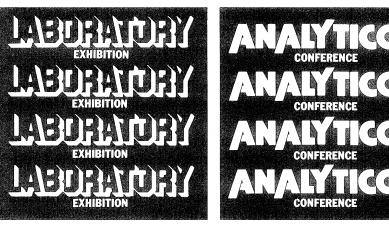
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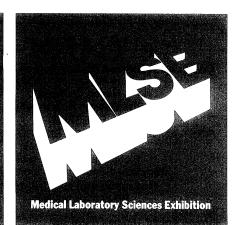
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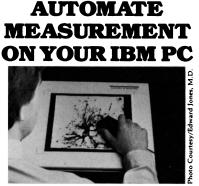
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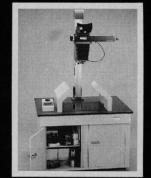
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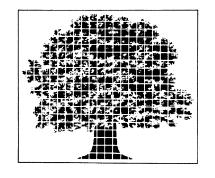
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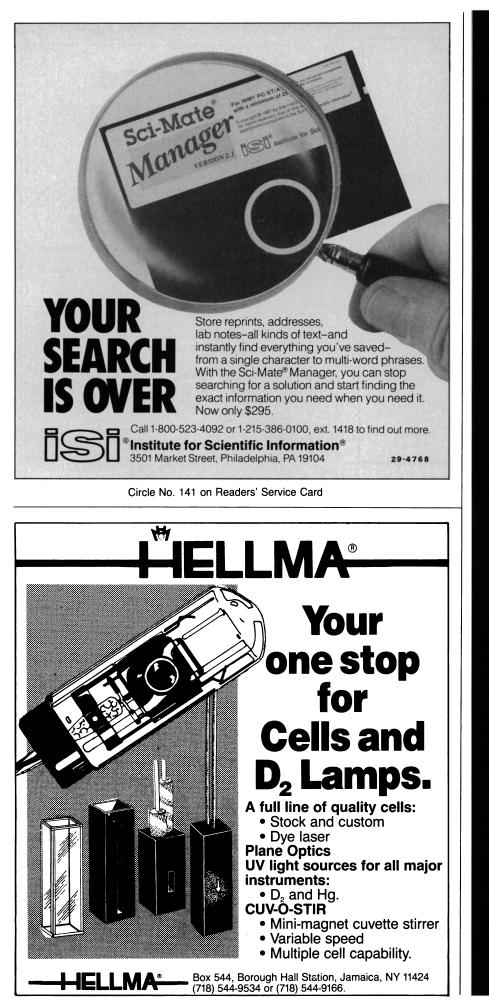
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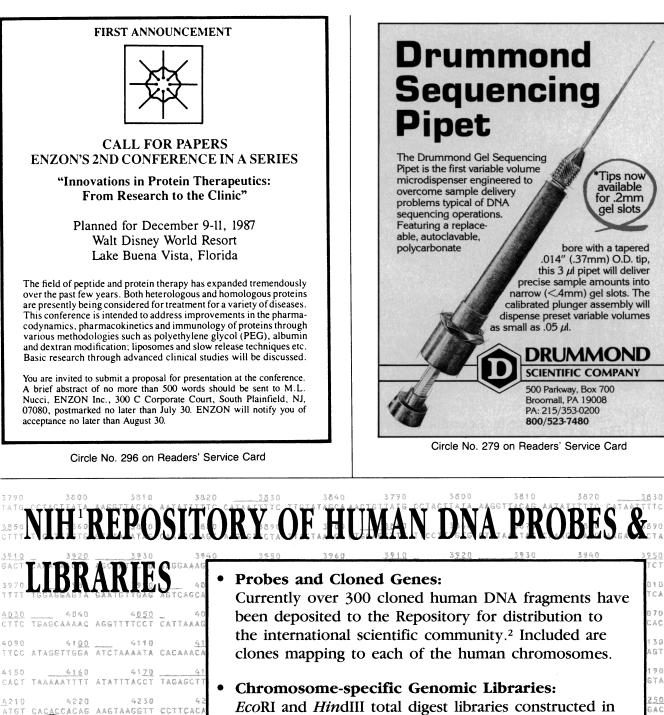
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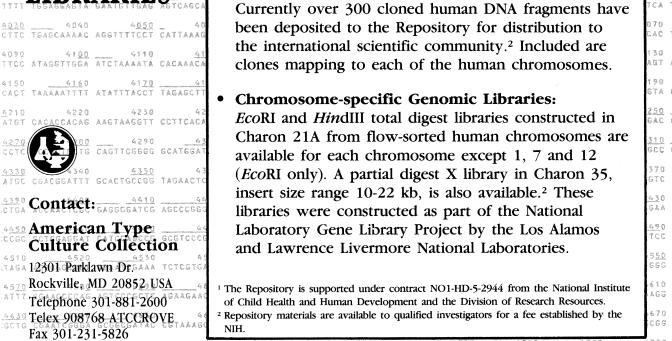


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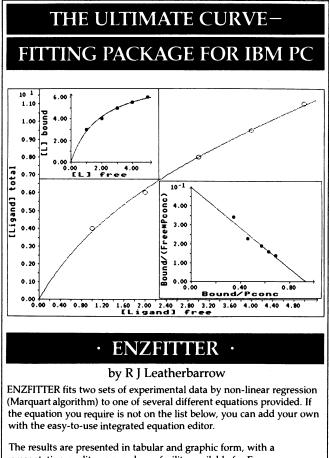
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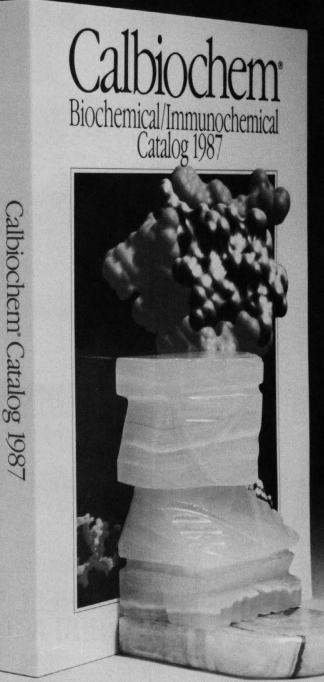
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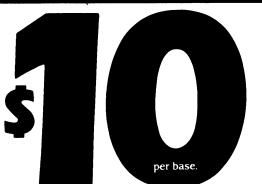
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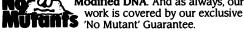
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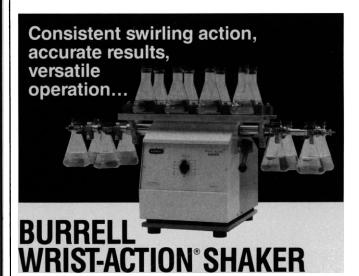


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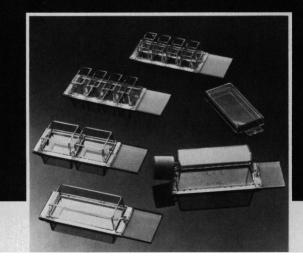
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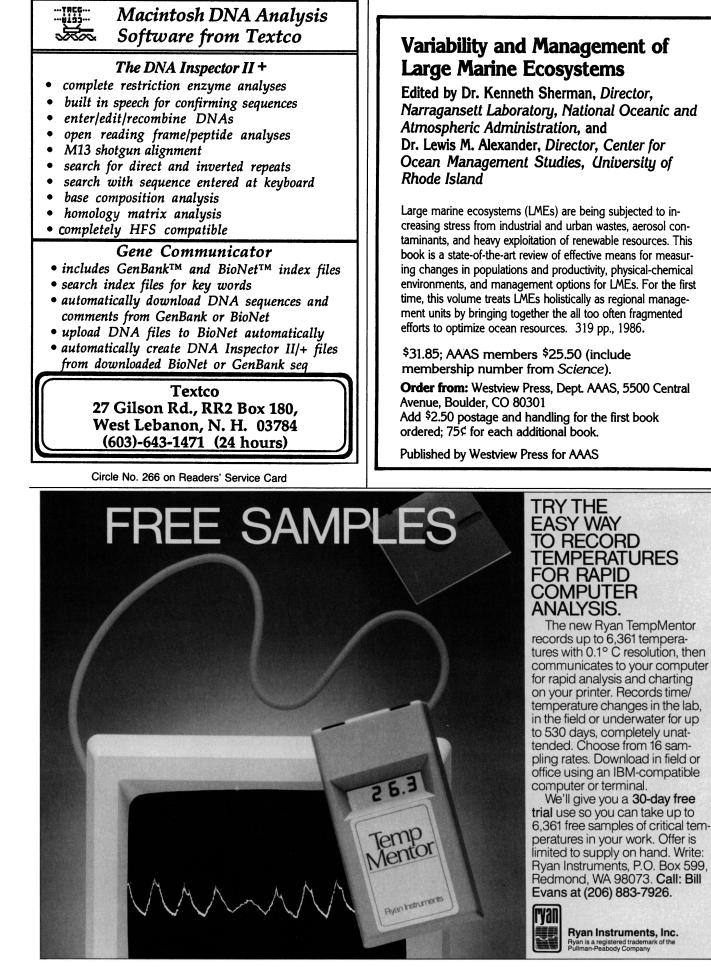
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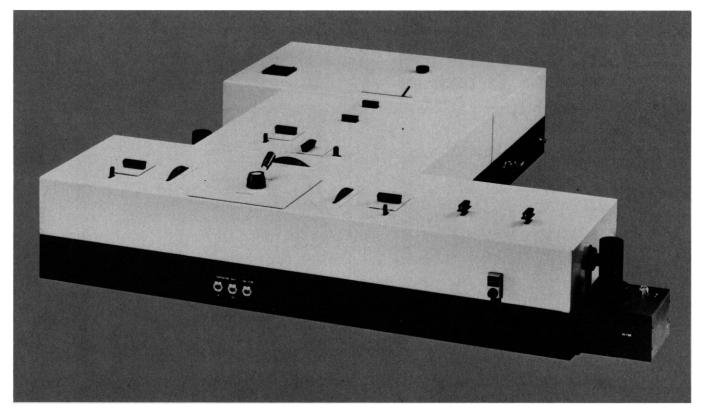
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KENT K. STEWART, Virginia Polytechnic Institute and State University, Blacksburg, Virginia Volume 1 (1987), 4 issues ISSN 0889-1575 Annual Subscription Rates U.S.A. and Canada: \$70.00 All other countries: \$82.00

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Volume 1 (1987), 4 issues ISSN 0889-1583

Annual Subscription Rates U.S.A. and Canada: \$100.00 All other countries: \$112.00

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