AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE

## Science

5 MARCH 1993 Vol. 259 • PAGES 1369–1508 \$6.00

special Report GOE TOXICOLOGYULAT

## 

Stratagene's 1993 Catalog features many new and exciting products. Among them are new vectors for mammalian and bacterial expression, vectors to quickly subclone amplified fragments, thermostable ligases, mycoplasma detection, *in situ* hybridization, new DNA sequencing enzymes, a benchtop cold chamber, a robotic thermocycler and much more. Please call Stratagene or the Stratagene distributor nearest you to receive your personal copy of the 1993 catalog. Also ask to be placed on the mailing list to receive our Strategies Newsletter and future product information.



#### Stratagene USA

Corporate Headquarters Phone: 800-424-5444 Telefax: 619-535-0034 Germany Stratagene GmbH Phone: (06221) 40 06 34 Telefax: (06221) 40 06 39 United Kingdom Stratagene Ltd. Phone: (0223) 42 09 55 Telefax: (0223) 42 02 34 France Stratagene France Phone: (0590) 72 36 Telefax: (1) 44 28 19 00 Switzerland Stratagene GmbH Phone: 01-3657707

Australia Integrated Sciences Phone: (2) 4362 Telefax: (2) 4364 (2) 4362611 (2) 4364442 Austria Chemomedica *Creutzberg & Co.* Phone: (0222)5332666 Telefax: (535)330658 Brazii Instrucom Phone: 55-11-530-7833 11-53949 Telefax: 55-11-530-0895 Canada nal Diagnostic Inc. 403/461-8794 Professi Phone: Telefax 403/450-9293 Teletax. China Hong Man Medicine and Development Co. Phone: 852-5-417-1803 Teletax: 852-5-417-1368

Distributors

Chemos Czechoslovakia Phone: 2-7862198 2-7862198 2-7862198 2-7862198 Telefax: Denmark AH Diagn Phone: Telefax: France nostics 86101055 86161533 Ozyme Phone: (1) 30570025 Telefax (1) 30441513 Finiand Kebo Lab Phone: 0y (9)0-43542003 (9)0-43543011 Telefax: Greece Biometria 01/88 37166 01/88 36757 01/88 34258 Phone: Telefax: Hong Kong Line Analytics, Ltd. 852-5-578-5839 852-5-807-2674 Phone Telefax:

Czechosiovakia

Hungary Inovex (Wien) Phone: (0222) 94-8509 Teletax: (0222) 94-850730 Inovex (Budapest) Phone: 36-1 166 9995 Teletax: 36-1 166 9995 India Genetix 5413401 091-11-5433906 Telefax: iran Farda Beh Co. Phone: 00 0098-21-893292, 0098-21-894591 0098-21-8801251 Telefax: ireland MedLabs Limited 353-1-2952101 353-1-2952165 Phon Telefax: Israei Getter (2000) Ltd Phone 03-576 1520 Telefax: 03-752 3626

Duotech SRL Phone: (2 (2) 33106630 (2) 33106640 Telefax: Japan Funakoshi Co. Ltd. Phone: (03)5684-1616 Telefax: (03)5684-1634 *Toyobo Co. Ltd. (Osaka)* Phone: 06-348-3785 Phone: or 3788 06-348-3322 Telefax: Toyobo Co. Ltd. (Tokyo) Phone: 03-3660-4819 Telefax: 03-3660-4887 Korea Koram Biotech. Corp. Phone: (2) 5560311 Telefax: (2) 5560828 Telefax: Malaysia Trans-Techno Enterprise 603-983-2212 603-984-9116 Phone: Telefax: The Netherlands and Belgium Westburg b.v. (33)950094 (33)951222 Phone: Telefax:

itaiv

New Zealand Science & Technology (NZ) Ltd. (Auckland) 09-270-3332 Phone: Phone: 00 2. (Christchurch) Phone: 03-383-1146 Phone: (Dunedin) Phone: 03-477-7860 Phone: C. (Wellington) Phone: 04-566-6096 KEBO Lab AS Phone: 47 5 28 13 10 Telefax: 47 5 28 57 96 Portugai Porcugal Biocontec Phone: 1-4533941 Telefax: 1-4533924 Russia DRG International, Inc. 253-1904 253-1082 Phone: Telefax: Singapore Research Instruments PTE Phone: 65-473-9035 65-473-9035 65-475-8565 Telefax:

South Africa Chemiab Ply. Ltd. Phone: 27-11-7893932 Telefax: 27-11-7877572 Whitehead Scientific 021-981-1560 Phone: 021-981-5789 Telefax Spain Cultek, Phone: 1 729 03 33 1 358 17 61 Telefax: Swede KFBO Lab AB Phone: Telefax: АВ (8) 6213400 (8) 6213470 Taiwan Cold Spring Biotechnology Co. Ltd. Phone: (02) 647-8855 (Rep.) Telefax: (02) 647-9160 Unimed Health Care Inc. Phone: (2) 720-2215 (2) 723-3666 Telefax: Turkey Genomed A.S. 90-1-248-2000 90-1-246-6819 90-1-247-9709 Telefax:

Circle No. 29 on Readers' Service Card

## **Certify Your LSC Results.**



#### Tri-Carb: The Only LSC Capable Of Certifying Its Own Performance.

All LSC's have some means of calibration, but only Packard Tri-Carbs actually certify their own performance. Tri-Carbs warn you of potential problems, such as changes in background or component aging, long before they can affect your results. Only with a Tri-Carb can you be sure you have correct performance every time you use it - and that you're in-line with today's GLP standards.

The fully integrated computer control and Instrument Performance Assessment (IPA) features form the basis of this unique self-testing capability. IPA monitors efficiency, background, sensitivity, and repeatability for <sup>3</sup>H and <sup>14</sup>C - automatically or on demand. The integrated computer stores the 100 most recent values for each of these critical parameters for immediate recall and output. There's no need to setup special procedures, or to labor through stacks of data. It's as easy as pressing a single key. Now you can certify your LSC performance and validate your experimental results . . . anytime!

It takes a lot of confidence to test yourself and show the results to everyone. It takes a Packard Tri-Carb to have the performance worth showing.

Packard Tri-Carb ... Certified TR-LSC Performance.



Packard Instrument Company, 800 Research Parkway, Meriden, CT 06450 U.S.A. Tel: 203-238-2351 Toll Free: 1-800-323-1891 TX: 643251 FAX: 203-235-1347

Freferen



A PACKARD

. Hannahara

- initiant

#### Packard International Offices:

Australia, Victoria 008-335638, Mt Waverley 543-4266; Austria, Vienna 43-1-302504-0; Belgium, Brussels 32-2-4668210; Canada, Ontario 1-800-387-9559; Denmark, Greve 45-42909023; France, Rungis (33) 1 46.86.27.75; Germany, Frankfurt (49-69) 663010; Italy, Milano (02) 33910796; Japan, Tokyo 81-3-3-866-5850; Netherlands, Groningen (050) 413360; Tilburg (013) 423900; Sweden, Uppsala 46-18 556900; Switzerland, Zurich (01) 481 69 44; United Kingdom, Pangbourne, Berks (44) 0734 844981.

Circle No. 24 on Readers' Service Card

# With our new range of options, monitoring your animals electronically is as easy as...

#### ONC THE POCKET SCANNER

**THE POCKET SCANNER** Introducing the world's smallest, fully integrated Pocket Scanner, the DAS-4004. It offers the simplest way to convert from the obsolete and unreliable methods of animal identification to our proven technology of pre-encoded microchip transponders that are carried on board the animal.

Direct and fool proof positive identification from cage to computer is easily facilitated with this unique Pocket Scanner. The unit is powered by a 9 volt replaceable battery and comes equipped with a communications port for downloading stored data to your own computer or printer.

Now you can enter the world of electronic monitoring at entry level, single-handed. Also available is the DAS-4004 EM with extended memory.

## THE NOTEBOOK

The DAS-4002 Portable Programmable Data Acquisition Notebook, with rechargeable battery, is easy to use.

The unit is simple to address with a menu driven system, allowing you to identify your animals using your own codes or numbers. At the time of the scan, the unit can record Date and Time, Weight and clinical observation data related to each animal. You can then transfer the recorded data to your own computer system.

This easy-to-use, state-of-the-art product improves data accuracy and efficiency in your animal research.

BioMedic

THE BENCH-TOP The DAS-4001 is a bench-top, line powered Programmable ID Data Acquisition System that can be fully integrated with the host computer. It can directly link the information gathered at a work station, such as weight, clinical observations and dosing, to your own on-line terminal based system.

Bio Medic Data Systems is the only company that offers animal monitoring systems that allow you to use your own identification codes.

There is simply no justification any longer for continuing primitive animal identification methods (tattooing, clipping, tagging) and accepting the additional errors in recording and transferring monitored information

ELAMS is a trademark of Bio Medic Data Systems, Inc.

mechanically. Today's research is too costly and errors are too expensive. Our Electronic Lab

Animal Monitoring System (ELAMS<sup>™</sup>) is economical and easily integrated with your existing methodologies. We are the only company in the

microchip (shown 1.5x) is the focus of the unique ELAMS system. industry with proven, task-oriented systems solutions

training, service and longterm R & D. Bio Medic Data Systems has tomorrow's answer to today's problems. Please call us and let's talk.



a bio Medic company Bio Medic Data Systems, Inc. 255 West Spring Valley Ave., Maywood, N.J. 07607 Phone: 201-587-8300

Toll free: 1-800-526-2637 Fax: 201-843-8816 Circle No. 14 on Readers' Service Card © Copyright BioMedic Corporation, 1992 Patents pending, US and International

ISSN 0036-8075 5 MARCH 1993 VOLUME 259 NUMBER 5100



AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE



1402 & 1442 Parasitic virulence varies with host population structure

STATISTICS.
1388
1389
1390
1391
1392
1393
1394
1398
1400
The second

What Might Cause Parasites to Become More Virulent	1402
Long Search for Sea Urchin Sperm Receptor Pays Off	1403
Mathematics: If You're Stumped, Try Something Harder	1404
PERSPECTIVE	
Cells in Stress: Transcriptional Activation of Heat Shock Genes R. I. Morimoto	1409
ARTICLES	
Radiative Climate Forcing by the Mount Pinatubo Eruption P. Minnis, E. F. Harrison, L. L. Stowe, G. G. C F. M. Denn, D. R. Doelling, W. L. Smith, Ju	1411 Bibson, r.
Molecular Matchmakers A. Sancar and J. E. Hearst	1415
RESEARCH ARTICLE	
Sea Urchin Egg Receptor for Sperm: Sequence Similarity of Binding Domain and hsp70 K. R. Foltz, J. S. Partin, W. J. Lennarz	1421

The Origins of Agriculture, reviewed by J. McCorriston • The Savage Within, F. Spencer • Natural Selection, S. C. Stearns • The History and Development of Human Genetics, A. G. Motulsky • Vignettes: Highs

#### DEPARTMENTS

1387

Stephen P. Goff Corey S. Goodman Stephen J. Gould Ira Herskowitz

Eric F. Johnson Stephen M. Kosslyn

Michael LaBarbera

Harvey F. Lodish Richard Losick Anthony R. Means

THIS WEEK IN SCIENCE	1377
EDITORIAL Basic Research (III): Priorities	1379
LETTERS Pondering Greenhouse Policy: S. H H. Dowlatabadi and L. B. Lave; M. C W. D. Nordhaus • Priority Envy: J. L.	<b>1381</b> I. Schneider; Oppenheimer; . Sutton

SCIENCESCOPE

John Abelson Frederick W. Alt

Henry R. Bourne James J. Bull

Kathryn Calame C. Thomas Caskey Dennis W. Choi

Prederick W. Alt Don L. Anderson Stephen J. Benkovic David E. Bloom Floyd E. Bloom

#### Board of Reviewing Editors

RANDOM SAMPLES

**BOOK REVIEWS** 

Mortimer Mishkin Roger A. Nicoll William H. Orme-Johnson III Stuart L. Pimm Yeshayau Pocker Dennis A. Powers Ralph S. Quatrano V. Ramanathan Douglas C. Rees Charles S. Levings III Erkki Ruoslahti Ronald H. Schwartz

and Lows • Books Received **PRODUCTS & MATERIALS** 

> Terrence J. Sejnowski Thomas A. Steitz Richard F. Thompson Robert T. N. Tjian Emil R. Unanue Geerat J. Vermeij Bert Vogelstein Harold Weintraub Zena Werb George M. Whitesides Owen N. Witte Keith Yamamoto

1405

1473

1479

AAAS Board of Directors

F. Sherwood Rowland Retiring President, Chairman Eloise E. Clark President Francisco J. Ayala President-elect

Robert A. Frosch Florence P. Haseltine William A. Lester, Jr. Alan Schriesheim Jean'ne M. Shreeve Chang-Lin Tien Warren M. Washington Nancy S. Wexler

William T. Golden Treasurer Richard S. Nicholson Executive Officer

SCIENCE • VOL. 259 • 5 MARCH 1993

John M. Coffin Bruce F. Eldridge Paul T. Englund Richard G. Fairbanks Douglas T. Fearon Harry A. Fozzard Victor P. Euchs

Victor R. Fuchs Theodore H. Geballe Margaret J. Geller John C. Gerhart Roger I. M. Glass

Victor R. Fuchs

1374

#### COVER

Zonally averaged latitudinal spreading of the increase (yellow, orange, and red) in reflected shortwave radiation measured by the Earth Radiation Budget Experiment after the eruption of Mount Pinatubo in June 1991. The vertical scale is latitude from 40° S to 40°N, and the

horizontal scale is time from May to November 1991. Volcanic aerosols reflect some of the sun's energy back to space; as a result, Earth's climate is cooled. See page 1411. [Image: P. Minnis et al., Atmospheric Sciences Division, NASA Langley Research Center]

1453

Antagonism of Catecholamine Receptor

#### **REPORTS**

Suble Compounds of Helium and Neon:       1428         Fighe Cos and NegCos       1457         M. Saunders, H. A. Jiménez-Vázquez, R. J. Cross,       S. L. Reiner, ZE. Wang, F. Hatam, P. Scott,         Do the Application of the Minimal       1430         Principle to Solve Unknown Structures       1460         M. Weeks, H. A. Hauptman       1430         A 2000-Year Tree Ring Record of Annual 1433       Inhibition of Human Colon Cancer       1463         Temperatures in the Sierra Nevada Mountains       A. Scuderi       Optical Time-of-Flight and Absorbance       1463         Copy of North American Ozone       1433       D. P. Arrish, J. S. Holloway, M. Trainer,       Optical Time-of-Slight and Absorbance       1463         Cossilization of Soft Tissue in the       1439       D. Parrish, J. S. Holloway, M. Trainer,       Neither Scheckman         Cossilization of Soft Tissue in the       1439       D. C. Briggs and A. J. Kear       1440         Condition for Virulence in Nematode Parasize of Fig Wasps       1442       The Legray Density of Water and Le       1459         R. K. Shotcher, R. M. Stroud, D. V. Santi, D. Kunz, K. M. Perry       1450       S. J. Whittaker and       1. Luzzatro, A. Bazarbachi, D. Bazatbachi, D. Brageul, J. Pérès, H. de The', W. W. Hall       1. Luzzatro, A. Bazarbachi, D. Kunz, K. M. Perry       1. Mathemator A. Bazarbachi, D. Perrisk, J. S. Thapseer, H. Weintraub       1. Luzzatro,	Crystal Structure and Optical Properties 1426 of Cd <sub>32</sub> S <sub>14</sub> (SC <sub>6</sub> H <sub>5</sub> ) <sub>36</sub> ·DMF <sub>4</sub> , a Cluster with a 15 Angstrom CdS Core N. Herron, J. C. Calabrese, W. E. Farneth, Y. Wang	Domains of the Receptors L. M. Luttrell, J. Ostrowski, S. Cotecchia, H. Kendall, R. J. Lefkowitz	
On the Application of the Minimal       1430         On the Application of the Minimal       1430         Principle to Solve Unknown Structures       6         R. Miller, G. T. DeTitta, R. Jones, D. A. Langs,       LA Cells in SCID Mice         A 2000-Year Tree Ring Record of Annual 1433       Competentiation of North American Ozone       1436         Converted Function       1430         Properatives in the Sierra Nevada Mountains       D. A. Benaron and D. K. Stevenson       1466         Poolution to the North Atlantic Ocean       D. A. Benaron and D. K. Stevenson       1466         Poolution of Soft Tissue in the       1439       Thokibitors of Human Colon Cancer       1466         Poolution of Virulence in Nematode       1439       Requirement for a GTPase-Activating       1466         Poolution of Virulence in Nematode       1442       The Energy Density of Water and Ice       1469         Subcrater Bace Discovery of Inhibitors       1442       The Singeldes       J. Wilen; M. Lahav, M. Eisenstein,       L. Leiserowitz         HTLV-1 Provirus       1470       Mycosis Fungoides       J. Wilen; M. Lahav, M. Eisenstein,       E. Leiserowitz         Officiency in Rhabdomyosarcomas       1450       S. J. Wilttaker and       L. Lucartor, A. Bazarbachi,       F. Saal, L. Laroche,       B. Flageul, J. Pérics,         B. J. Agesort, M.	Stable Compounds of Helium and Neon:       1428         He@C <sub>60</sub> and Ne@C <sub>60</sub> M. Saunders, H. A. Jiménez-Vázquez, R. J. Cross, R. J. Poreda	T <sub>H</sub> 1 and T <sub>H</sub> 2 Cell Antigen Receptors1457in Experimental LeishmaniasisS. L. Reiner, ZE. Wang, F. Hatam, P. Scott, R. M. Locksley	
A 2000-Year Tree Ring Record of Annual 1433         Competentives in the Sierra Nevada Mountains L. A. Scuderi         Export of North American Ozone       1436         Pollution to the North Atlantic Ocean       1437         D. D. Partish, J. S. Holloway, M. Trainer,       1439         P. C. Murphy, G. L. Forbes, F. C. Fehsenfeld       1439         Fossilization of Soft Tissue in the       1439         Aboratory       1442         Population Structure and the       1442         Volution of Virulence in Nematode       1442         Parasites of Fig Wasps       1444         F. A. Here       1442         Structure-Based Discovery of Inhibitors       1445         of Fator Required for MyoD Activity       1450         M Myogenesis       5. J. Tapscott, M. J. Thayer, H. Weintraub         Policiates accompanying feature       1         South Myogenesis       1         S. J. Tapscott, M. J. Thayer, H. Weintraub       1         StelEMCE (658M 0036-6075) la published weeky on Friday, except the last week in       2       2         Moderator       1       1         S. J. Tapscott, M. J. Thayer, H. Weintraub       2       2         Medicates accompanying feature       2       2         Moderato request. Authorization to	On the Application of the Minimal Principle to Solve Unknown Structures R. Miller, G. T. DeTitta, R. Jones, D. A. Langs, C. M. Weeks, H. A. Hauptman	Inhibition of Human Colon Cancer 1460 Growth by Antibody-Directed Human LAK Cells in SCID Mice H. Takahashi, T. Nakada, I. Puisieux	
Export of North American Ozone 1436   Export of North American Ozone 1436   Pollution to the North Atlantic Ocean 1436   D. D. Partish, J. S. Holloway, M. Trainer, P. C. Murphy, G. L. Forbes, F. C. Fehsenfeld   Fossilization of Soft Tissue in the 1439   Jaboratory D. E. G. Briggs and A. J. Kear   Population Structure and the 1442   Colution of Virulence in Nematode 1442   Parasites of Fig Waps 1442   E. A. Herre 1442   Structure-Based Discovery of Inhibitors 1445   B. K. Shoichet, R. M. Stroud, D. V. Santi, 1. Luzzatro; A. Bazarbachi,   J. D. Kuntz, K. M. Perry Structure, M. Stroud, D. V. Santi,   Deficiency in Rhabdomyosarcomas 1450   of a Factor Required for MyoD Activity 1450   of a Factor Required for MyoD Activity 1450   M Myogenesis S. J. Tapscott, M. J. Thayer, H. Weintraub   SciENCE (USM 9036-8075) is published weekly on Friday, sccept the last week in   Bus rates on request. Authorization to photocopy material for internal or period for internal or period for internal or period by material for internal or perio	A 2000-Year Tree Ring Record of Annual 1433 Temperatures in the Sierra Nevada Mountains	Optical Time-of-Flight and Absorbance1463Imaging of Biologic MediaD. A. Benaron and D. K. Stevenson	
Fossilization of Soft Tissue in the laboratory       1439         Laboratory       D. E. G. Briggs and A. J. Kear         Population Structure and the laboratory       1442         Population of Virulence in Nematode       1442         Value arisites of Fig Wasps       1442         E. A. Herre       1442         Structure-Based Discovery of Inhibitors       1445         Mycosis Fungoides       S. J. Whittaker and         L. Luzzatto; A. Bazarbachi,       F. Saal, L. Laroche,         B. Flageul, J. Périès,       H. de Thé; W. W. Hall         Deficiency in Rhabdomyosarcomas       1450         of a Factor Required for MyoD Activity       1450         of a Factor Required for MyoD Activity       1450         M Myogenesis       S. J. Tapscott, M. J. Thayer, H. Weintraub         Indicates accompanying feature       Indicates accompanying feature	Export of North American Ozone 1436 Pollution to the North Atlantic Ocean D. D. Parrish, J. S. Holloway, M. Trainer, P. C. Murphy, G. L. Forbes, F. C. Fehsenfeld	Requirement for a GTPase-Activating       1466         Protein in Vesicle Budding from the       1466         Endoplasmic Reticulum       T. Yoshihisa, C. Barlowe, R. Schekman         TECHNICAL COMMENTS       1466	
Population Structure and the       1442         Evolution of Virulence in Nematode       1442         Parasites of Fig Wasps       E. A. Herre         Structure-Based Discovery of Inhibitors       1445         of Thymidylate Synthase       S. J. Whittaker and         B. K. Shoichet, R. M. Stroud, D. V. Santi,       I. Luzzatto; A. Bazarbachi,         I. D. Kuntz, K. M. Perry       Flageul, J. Péries,         Deficiency in Rhabdomyosarcomas       1450         of a Factor Required for MyoD Activity       1450         M Myogenesis       S. J. Tapscott, M. J. Thayer, H. Weintraub         SciENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in       Bulk rates on request. Authorization to photocopy material for internal or percenters	Fossilization of Soft Tissue in the Laboratory1439D. E. G. Briggs and A. J. Kear	The Energy Density of Water and Ice 1469 Nucleation	
<ul> <li>B. K. Shoichet, R. M. Stroud, D. V. Santi, I. D. Kuntz, K. M. Perry</li> <li>Deficiency in Rhabdomyosarcomas</li> <li>1450</li> <li>fa Factor Required for MyoD Activity and Myogenesis</li> <li>S. J. Tapscott, M. J. Thayer, H. Weintraub</li> <li>Indicates accompanying feature</li> </ul> SciENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in Bulk rates on request. Authorization to photocopy material for internal or personal or p	Population Structure and the I 1442 Evolution of Virulence in Nematode Parasites of Fig Wasps E. A. Herre Structure-Based Discovery of Inhibitors 1445 of Thymidylate Synthase	HTLV-1 Provirus 1470 and Mycosis Fungoides S. J. Whittaker and L. Luzzatto; A. Bazarbachi, F. Saal, L. Laroche,	
Deficiency in Rhabdomyosarcomas       1450         of a Factor Required for MyoD Activity       1450         and Myogenesis       S. J. Tapscott, M. J. Thayer, H. Weintraub         Indicates accompanying feature       semi         SCIENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in       Bulk rates on request. Authorization to photocopy material for internal or personal o	B. K. Shoichet, R. M. Stroud, D. V. Santi, I. D. Kuntz, K. M. Perry	B. Flageul, J. Périès, H. de Thé; W. W. Hall	
SCIENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in Bulk rates on request. Authorization to photocopy material for internal or personal states and the last week in Bulk rates on request.	Deficiency in Rhabdomyosarcomas       1450         of a Factor Required for MyoD Activity       1450         and Myogenesis       1         S. J. Tapscott, M. J. Thayer, H. Weintraub       1		A
SCIENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in Bulk rates on request. Authorization to photocopy material for internal or personal structure in the structure in		Indicates accompanying feature	semic
recember, by the American Association for the Advancement of Science, 1333 H Street, circumstances not failing within the fair use provisions of the Copyright Act is gran	SCIENCE (ISSN 0036-8075) is published weekly on Friday, exc December, by the American Association for the Advancement of Sc	cept the last week in Bulk rates on request. Authorization to photocopy material for circumstances not falling within the fair use provisions of the Copy	r internal or perso yright Act is grant

December, by the American Association for the Advancement of Science, 1333 H Street, NW, Washington, DC 20005. Second-class postage (publication No. 484460) paid at Washington, DC, and additional mailing offices. Copyright © 1993 by the American Association for the Advance-ment of Science. The title SCIENCE is a registered trademark of the AAAS. Domestic individual membership and subscription (51 issues): \$87 (\$47 allocated to subscription). Domestic institutional subscription (51 issues): \$205. Foreign postage extra: Mexico, Caribbean (surface mail) \$50; other countries (air assist delivery) \$95. First class, airmail, student and emeritus rates on request. Canadian rates with GST available upon request, GST #1254 88122. Change of address: allow 6 weeks, giving old and new addresses and 11-digit account number. Postmaster: Send change of address to *Science*, P.O. Box 2033, Marion, OH 43305-2033. Single copy sales: \$6.00 per issue prepaid includes surface postage; Guide to Biotechnology Products and Instruments, \$20.

anal use under ed by AAAS to circumstances not falling within the fair use provisions of the Copyright Act is granted by AAAS to libraries and other users registered with the Copyright Clearance Center (CCC) Transactional Reporting Service, provided that the base fee of \$1 per copy plus \$0.10 per page is paid directly to CCC, 27 Congress Street, Salem, MA 01970. The identification code for *Science* is 0036-8075/83 \$1 + .10. *Science* is indexed in the *Reader's Guide to Periodical Literature* and in several specialized indexes. The American Association for the Advancement of Science was founded in 1848 and incorporated in 1874. Its objectives are to further the work of scientists, to facilitate cooperation among them, to foster scientific freedom and responsibility, to improve the effectiveness of science in the promotion of human welfare, to advance education in science, and to increase public understanding and appre-ciation of the importance and promise of the methods of science in human progress.

ciation of the importance and promise of the methods of science in human progress.

SCIENCE • VOL. 259 • 5 MARCH 1993

1426 chip of a onductor

#### **Restriction Enzymes from Promega...**





#### THIS WEEK IN SCIENCE

.....

edited by PHIL SZUROMI

#### Aerosol cooling by Mount Pinatubo

AND THE REAL PROPERTY AND A REAL PROPERTY OF

The eruption of Mount Pinatubo in June 1991 injected gases and ash into the stratosphere. Aerosols produced from this eruption acted to cool global climate, primarily by acting to reflect more of the incoming solar radiation. Minnis et al. (p. 1411) use measurements from the Earth Radiation Budget Experiment to quantify these and other changes in Earth's radiation budget that occurred after the eruption. Cooling increased through September 1991 and was related to both an increase in albedo over clear areas resulting directly from the presence of the aerosol as well as an increase in the albedo of clouds as the aerosols acted as nucleation sites for cloud droplets.

#### Creating a piece of a crystal core

A cadmium sulfide (CdS) complex containing a large number of Cd and S atoms has been made that has a structure resembling bulk CdS, a semiconductor. Herron et al. (p. 1426) found that recrystallizing  $Cd_{10}S_4(SC_6H_5)_{12}$  with dimethylformamide (DMF) formed  $Cd_{32}S_{14}(SC_6H_5)_{36}$ ·DMF<sub>4</sub>. The core of this large cluster is similar to a 12 angstrom spherical piece of CdS in the sphalerite form. The cluster has luminescence and exciton properties similar to those of extended solids.

#### Helium and neon inside C<sub>60</sub>

Fullerenes can be prepared by arc heating graphite rods in low pressures of helium. Saunders *et al.* (p. 1428) show that heating  $C_{60}$  to high temperatures releases helium. The amount corre-

#### Receptor domains as receptor antagonists

When many hormones or neurotransmitters bind to their cellular receptors, they cause activation of heterotrimeric guanine nucleotide binding proteins (G proteins). The G proteins couple the receptors to enzymes that are activated in response to the bound ligand. Luttrell *et al.* (p. 1453) expressed a gene encoding a fragment of the  $\alpha_{1B}$ -adrenergic receptor in cells along with the wild-type receptor. The fragment contained the region of the receptor primarily responsible for interaction with G proteins, and its presence specifically inhibited signaling by the full-length receptor. Even signals mediated by the same G protein through interaction with other receptors were not affected. The results suggest that it may be possible to design drugs that act as very specific antagonists of such receptors.

sponds to about 1 in  $10^6$  molecules incorporating a helium atom and is consistent with a simple model based on cavity volume. The activation energy for release suggests a reversible bond-breaking event in the fullerene cage. Neon can be incorporated into  $C_{60}$  by using a neon atmosphere.

From soft tissue

to fossil

Some of the most spectacular

and valuable fossils in the geo-

logic record preserve intricate

details of soft tissues of extinct

organisms. Such fossils often

consist of calcium phosphate,

and it has been thought that el-

evated levels of phosphate in

sediment pore waters were

needed for this process. Briggs

and Kear (p. 1439) show that

the calcium and phosphate pres-

ent in tissue itself can be suffi-

cient to create a fossil. They

studied the decay of shrimp un-

der simulated fossilization conditions and found that the shrimp became partially mineralized with amorphous calcium phosphate, especially under anaerobic conditions. This mineralization process preserved morphological details of the soft tissue.

#### Antigen receptors and infection response

Certain diseases caused by parasitic infection, such as leprosy, tuberculosis, and leishmaniasis, progress in a highly variable way; some infected individuals are asymptomatic, and others manifest most or all of the symptoms of the disease state. Reiner et al. (p. 1457) analyzed the types of T cell receptors (TCRs) used by mice that were infected with the disease-causing agent of leishmaniasis, Leishmania major. Mice that responded by mounting an inappropriate T helper 2 CD4<sup>+</sup> attack developed progressive leishmaniasis and yet expressed the same TCRs as the cells from mice that produced a healing T helper 1 response. Because the same epitope can drive both immunity and progression of disease, candidate vaccine antigens from immune individuals might actually exacerbate the disease.

#### **Targeting killers**

And the second second

Interleukin-2 (IL-2) can activate a killer cell response when it is added to peripheral blood in vitro. The lymphokine-activated killer (LAK) cells that are generated are cytotoxic to colon cancer cells in vitro but appear to have little effect when used in clinical trials against advanced human colon cancer. One potential problem is that an insufficient number of LAK cells arrive at the tumor mass. Takahashi et al. (p. 1460) show that the effectiveness of LAK cells can be improved by targeting them to the tumor cells with monoclonal antibodies (MAbs). They conjugated LAK cells to MAbs to a cell surface antigen of the tumor cells with polyethylene glycol. Such cell conjugates were more effective against human colon adenocarcinomas established in the livers of severe combined immunodeficient (SCID) mice than either the LAK cells or the MAbs alone.

#### Vesicle budding and GTPase activation

Proteins are transported in vesicles that bud from the membranes of the endoplasmic reticulum. The formation of such transport vesicles in the yeast Saccharomyces cerevisiae requires binding and hydrolysis of guanosine triphosphate (GTP) by a GTP-binding protein called Sar1p. Yoshihisa et al. (p. 1466) examined the interaction of Sar1p with another protein, Sec23, that is also required for vesicle budding. They found that Sec23 activated the guanosine triphosphatase (GTPase) activity of Sar1p but not that of other small GTP-binding proteins. Sec23 has no sequence similarity to other GTPase-activating proteins and thus may be an example of a new class of these enzymes.

We'll be the first to admit it: You have to have a lot of faith in your DNA supplier.

To manufacture precisely the proper sequence. To provide full documentation. And to deliver your order right on time, right on the money.



It's precisely that kind of trust that has made us the fastest growing supplier of synthetic DNA in the world. And it's why our customers keep coming back for more.

At National Biosciences, we'll deliver your custom synthesized primers and probes within three business days for as low as *\$2 per base\**. With no set-up charges. We'll even give you a free oligonucleotide with your first order.

To find out more, call (800) 747-4362. Or fax us your order at (800) 369-5118.

We've got the right stuff.



\*Based on estimated annual usage

#### WHILE YOU SPENT ONE HOUR DOING THIS,



#### LOOK AT ALL WE'VE BEEN UP TO.



We began by running the separation at several pHs.



Tested gradient slopes for improved resolution.





Confirmed the optimum pH at 6.0.



Selected 30 column volume gradient as optimum.









Studied the effect of loading at 2 mg and 5 mg.



Checked biological activity and purity.

Here's the challenge. Take your protein sample, map a range of pHs, optimize gradient and loading, purify 10 mg, analyze fractions, and check purity—all in about an hour. How? Using the BioCAD Workstation, you can run 3-minute separations, examine more variables than ever before, and get the highest purity products in hours...not weeks, not months. BioCAD employs Perfusion Chromatography," a patented new technology enabling high resolution separations on any of 25 surface chemistries. It's the one tool you need to accelerate the pace of discovery and development in your lab. Find out more. Call

Toll Free (800) 899-5858.



**PerSeptive Biosystems** 38 Sidney Street, Cambridge, MA 02139





©1993 PerSeptive Biosystems, Inc. BioCAD is a trademark of PerSeptive Biosystems, Inc. Poros and Perfusion Chromatography are registered trademarks of PerSeptive Biosystems, Inc

Circle No. 16 on Readers' Service Card



#### **TSG-**P53<sup>™</sup> AND **PIM<sup>™</sup>** TRANSGENIC MICE. EXPLORE NEW OPTIONS IN DRUG DISCOVERY AND TOXICOLOGY.

Examine the role of tumor suppression in vivo. Screen chemopreventive and chemotherapeutic compounds in the absence of tumor suppressor function. Accelerate carcinogenicity testing to save time and expense.

GenPharm<sup>™</sup> TSG-p53 mice have an increased susceptibility to a broad range of tumors arising from inactivation of the p53 tumor suppressor gene function. By using a mouse with inactivation of either one or both p53 alleles, you can study a compound's carcinogenicity as well as tumor prevention and treatment.

GenPharm PIM mice, with an additional copy of the pim-1 oncogene, develop T-cell lymphomas in response to a carcinogen in significantly less time than nontransgenic mice.

Transform your ideas into results through GenPharm TSG-p53 and PIM transgenic mice. For technical information and availability of models, contact GenPharm sales at 415-964-7024 or fax 415-964-3537.



#### GENPHARM INTERNATIONAL

P.O. Box 568, Los Altos, CA 94023 1. Cell 56, 673-682 (1989). 2. Nature 340, 61-63 (1989). 3. Cancer Research 51, 958-963 (1991). GenPharm, PIM and TSG-p53 are trademarks of GenPharm International, Inc.

Circle No. 20 on Readers' Service Card

## **Infrared DNA Sequencing...**

## Now you have a reason to accelerate your automation plans!



If you've been considering an automated system,

only the new Model 4000 DNA Sequencer from LI-COR can give you the accuracy, flexibility and affordability that characterize automated *infrared* DNA sequencing.

- 500 Bases per Sample
- Typical Accuracy of 99%
- Solid-State Reliability
- Application Flexibility
- Affordable Expansion
- Reduced Operation Costs

With the Model 4000, you prepare samples using standard Sanger dideoxy protocols. You can load up to 11 samples. Novel, infrared fluorescent dyelabeled primers provide excellent sensitivity for detection. A high contrast image of the DNA bands is produced within 4 to 7 hours of loading.

Data are presented in the most easily interpreted form an autoradiogram-like image. Real time image display lets you verify sample quality in under one hour. And the system's **Base ImagIR™** software typically lets you automatically call 450 bases per sample with 99% accuracy; 500 bases or more by switching to semiautomatic base calling.

Image quality and base calling accuracy are ensured by infrared detection and solidstate components. Solid-state technology increases reliability while reducing the initial purchase price, operating costs and instrument size. Infrared detection also allows you to use affordable reagents and consumables.

Focusing infrared optics and variable scan speeds give you the flexibility to use gels of different height, thickness and composition for faster throughput and new applications. Even greater flexibility comes from the Model 4000's multitasking software which lets you expand the system. Two additional sequencers can be added to one computer at very reasonable prices. LI-COR has been a leader in instrumentation for plant biological sciences for over 20 years. Now it's time for new ideas in biotechnology!

Why wait? Call for a free video or technical information.



1-800-645-4267 (Toll Free, U.S. & Canada)

LI-COR, Inc., 4421 Superior Street, Lincoln NE 68504, USA • 1-800-645-4267 (Toll Free U.S. & Canada) • Phone: 402-467-3576 • FAX: 402-467-2819 Distributors & Tel - Australia: John Morris Scientific Pty Limited (02) 417 8877 • Germany: MWG-BIOTECH 80 92 2 40 71 Italy: F.A.S.T. Canovai SRL (06) 8862246 • Switzerland, Austria: DMP AG (01) 946 06 76 • Taiwan: San Kwang Instruments Co. Ltd. (02) 393-4231 Circle No. 21 on Readers' Service Card



### Scientific Excellence in Reference Publishing



#### Academic Press Dictionary of Science and Technology

"The Academic Press Dictionary of Science and Technology captures the terminology of today and gives modern definitions. This dictionary is highly recommended for all public, academic, and high school libraries."

—BOOKLIST/REFERENCE BOOKS BULLETIN

**One Volume** 1992, 2432 pp., \$115.00/ISBN: 0-12-200400-0

#### Encyclopedia of Immunology

Edited by Ivan M. Roitt and Peter J. Delves

"A most impressive accomplishment. Its organization is extremely user-friendly, with a complete list of entries at the beginning of each volume....This is a good buy. Its value will be evident for years."

> —Kent T. Hayglass UNIVERSITY OF MANITOBA

Three-Volume Set 1992, 2032 pp., \$450.00/ISBN: 0-12-226760-5

#### Encyclopedia of Human Biology

Editor-in-Chief Renato Dulbecco

"The format and illustrations are excellent and encourage browsing; and the whole series has clearly been assembled with great labour and dedication. Articles are longer than those in most enclyclopaedias."

-NATURE

Order from your local bookseller or directly from

ACADEMIC PRESS

Eight-Volume Set 1991, 6722 pp., \$1995.00/ISBN: 0-12-226747-8



#### Encyclopedia of Physical Science and Technology SECOND EDITION

Edited by Robert A. Meyers

"Five years ago, *The Chicago Tribune* called the First Edition of this set 'awesome.' The Second Edition is even better....While there will always be a lag between the discoveries of scientists and researchers in the physical sciences and the dissemination of information on them, you can depend on seeing the first complete presentation in the Encyclopedia of Physical Science and Technology."

-INTERNET WORLDWIDE

Eighteen-Volume Set 1992, 14,080 pp., \$2500.00/ISBN: 0-12-226930-6

#### Encyclopedia of Earth System Science

Editor-in-Chief William A. Nierenberg Foreword by Robert M. White

"This excellent encyclopedia of the earth sciences stresses human interaction with all environments on and surrounding the earth....The articles are extremely well-written so that In most cases the educated layperson as well as the researcher are accommodated."

-BOOKLIST/REFERENCE BOOKS BULLETIN Four-Volume Set

1991, 2629 pp., \$950.00/ISBN: 0-12-226719-2



#### Encyclopaedia of Food Science, Food Technology and Nutrition

Edited by

Robert Macrae, Richard Robinson, and Michéle Sadler

#### **Key Features**

- An eight-volume set including a complete subject index with more than 500,000 entries
- More than 1500 contributors from 50 different countries
- 500 subject entries covering all aspects of food science, food technology, and nutrition
- 1000 articles written by the world's leading scientists and approved by an independent review procedure

Eight-Volume Set June 1993, c. 6114 pp., \$2100.00 (tentative) ISBN: 0-12-226850-4 Prepublication Price: \$1795.00\* \*Expires on final day of month of publication.

#### Encyclopedia of Microbiology

Editor-in-Chief Joshua Lederberg

HBJ Order Fulfillment Department DM17915 | CALL TOLL FREE

6277 Sea Harbor Drive, Orlando, FL 32887

Hailed by leading scientists and researchers around the world as "excellent," "outstanding," and "impressive," the **Encyclopedia of Microbiology** will be a standard text for finer academic, industrial, and personal libraries for years to come.

> 1-800-321-5068 FAX 1-800-336-7377

Four-Volume Set 1992, 2518 pp., \$695.00/ISBN: 0-12-226890-3

Circle No. 22 on Readers' Service Card

Prices subject to change without notice. © 1993 by Academic Press, Inc. All Rights Reserved. LK/CEP/BTK - 20033



## It's Time You Took A Second Look At CE Sensitivity

New levels of sensitivity and reproducibility have been achieved by Applied Biosystems' Capillary Electrophoresis System (Model 270A-HT). We've now eclipsed our own industry-leading standard by more than 30-fold, ensuring results you can't attain from any other system.

Whether you need high-efficiency separations for research or quality control, the automated 270A-HT provides the flexibility to handle a range of chemistries and applications. And now we've developed a new,



10 pmol/mL digest of β-lactoglobulin Top graph: High Sensitivity option. Bottom graph: Standard Instrument

proprietary optical technology especially for our CE systems-increasing sensitivity by more than an order of magnitude. You can achieve this unprecedented sensitivity without modifying applications or protocols you currently use on our systems. Additional sensitivity-enhancing strategies, such as stacking techniques, can increase performance even further.

With the 270A-HT you can measure reaction rates-instead of evaporation rates-because our specially designed sample vials and thermostatted sample handling system eliminate evaporation for precise, run-to-run reproducibility.

There are more reasons to take another look at Applied Biosystems capillary electrophoresis systems. In addition to providing the highest performance CE instrumentation, we're committed to keeping our customers on the leading edge with a full selection of specialty application kits, responsive service and



Antihypertensive drug formulation analysis with impurity sensitivity capability of < 0.01%.

expert technical assistance. For free product information phone Applied Biosystems today at: **Australia** (03) 808-7777, **Benelux** (0) 3465-74868, **Canada** (800) 668-6913, **France** (1) 48 63 24 44, **Germany** (0) 6150/101-0,

Italy (02) 8912 6011, Japan (03) 3699-0700, U.K. (0925) 825650, U.S. (800) 345-5ABI. ©1993 Applied Biosystems, Inc.



A Division of Perkin-Elmer Corporation

Circle No. 17 on Readers' Service Card

## The Fastest Confocal You Can Buy... 30 Confocal Images in One Second

0.5

In the time it takes you to read this sentence, you could have acquired and stored 240 confocal images using the **INSIGHT** Laser Scanning Confocal Microscope system.\*

0.4

Isn't it time you took a look at the **INSIGHT** confocal system from Meridian Instruments? Call or FAX us today for more information.

0.3



0.7

0.6

\* This rapid speed is made possible by the patented Bilateral Scanning technology, developed by Dr. G. J. Brakenhoff at the University of Amsterdam.

Image series and graph showing calcium response during a single beat of a continuously beating rat heart cell stained with Fluo-3. (Images acquired and analyzed using the INSIGHT-IQTM system; cells courtesy of Dr. Scott Henry, Parke-Davis, Ann Arbor, MI.)



2310 Science Parkway Okemos, Michigan 48864 Phone: 800-247-8084 • 517-349-7200 Fax: 517-349-5967 Meridian Instruments Europe, Inc. Industriepark—West 75 B-9100 St. Niklaas BELGIUM Phone: 32-3-7801760 • Fax: 32-3-7781727 Meridian Instruments Far East K.K. The Sonehara Building (2nd Floor) 2-23-3 Higashi Nihonbashi Chuo-ku, Tokyo 103 JAPAN Phone: 03-5820-3315 • Fax: 03-5820-3316

Distributors located in Australia, Austria, Benelux, France, Germany, Hong Kong, Israel, Italy, Korea, New Zealand, Portugal, Spain, Sweden, Switzerland, Taiwan, and the United Kingdom.

Circle No. 13 on Readers' Service Card

#### **Materials Science and Technology**

A Comprehensive Treatment A Series of 18 Self-Contained Handbooks

#### **C**ONTRIBUTORS

"Materials Science and Technology" is the first indepth, topic-oriented publication devoted to this enormous interdisciplinary field. It is truly international in scope: More than 200 highly-qualified authors from throughout the world have contributed state-of-the-art information.

#### COVERAGE

Each volume is a comprehensive reference work. Volumes are devoted to the most important classes of materials: metals, ceramics, glasses, polymers, semiconductors, and composites. They are presented in detail according to materials properties

■ materials processing

■ specific applications (e.g. in nuclear energy and medicine)

■ general phenomena (e.g. phase transformations, characterization of materials, plastic deformation and fracture of materials).

#### APPEAL

Materials scientists, physicists, chemists, and chemical and electrical engineers, metallurgists and ceramists, whether in industry or academia will find this series indispensable. In addition, the individual volumes serve as excellent graduate-level texts.



#### **EDITORS-IN-CHIEF**

R.W. Cahn, Cambridge, U.K. P. Haasen, Göttingen, Germany E.J. Kramer, Cornell , U.S.A.

#### **ORDER FORM**

Please invoice me for

- the marked volume(s)
   (\$ 325.00 per vol.)
- □ the whole series (\$ 270.00. per vol.)
- Please send me more information

Name	 	 
Address	 	
Signature		

#### THE VOLUMES

□ Vol.1: Structure of Solids (Ed: V. Gerold, MPI Stuttoart. FRG) ISBN 3-527-26814-6 O Vol. 2A/B (Two parts, part B mid-1993): Characterization of Materials (Ed: E. Lifshin, GEC, USA) ISBN 3-527-26815-4 Vol. 3A/B (two parts, part B) mid-1993): Electronic and Magnetic Properties of Metals and Ceramics (Ed: K.H.J. Buschow, Philips, Eindhoven, NL) Vol. 3A ISBN 3-527-26816-2 Vol. 3B ISBN 3-527-28264-5 □ Vol. 4: Electronic Structure and Properties of Semiconductors (Ed: W. Schröter, Göttingen, FRG) ISBN 3-527-26817-0 □ Vol. 5: Phase Transformations in Materials (Ed: P. Haasen, Göttingen, FRG) ISBN 3-527-26818-9 □ Vol. 6: Plastic Deformation and Fracture of Materials (Ed: H. Mughrabi, Erlangen, FRG) ISBN 3-527-26819-7 □ Vol. 7: Constitution and Properties of Steels (Ed: F.B. Pickering, Sheffield, UK) ISBN 3-527-26820-0 Vol. 8 (1994) Structure and Properties of Nonferrous Alloys (Ed: K.H. Matucha, Frankfurt, FRG) □ Vol. 9: Glasses and Amorphous Materials (Ed: J. Zarzycki, Montpellier, France) ISBN 3-527-26822-7 Vol. 10 (May 1993): Nuclear Materials (Ed: B.R.T. Frost, Argonne, USA) ISBN 3-527-26823-5 D Vol. 11 (April 1993): Structure and Properties of Ceramics (Ed: M. Swain, Sydney, Australia) ISBN 3-527-26824-3 □ Vol. 12 (April 1993): Structure and Properties of Polymers (Ed: E.L. Thomas, MIT, USA) ISBN 3-527-26825-1 Vol. 13 (May 1993): Structure and Properties of Composites (Ed: T.W. Chou, Delaware, USA) ISBN 3-527-26826-X □ Vol. 14: Medical and Dental Materials (Ed: D.F. Williams, Liverpool, UK) ISBN 3-527-26827-8 □ Vol. 15: Processing of Metals and Alloys (Ed: R.W. Cahn, Cambridge, UK) ISBN 3-527-26828-6 □ Vol. 16 (1994): Processing of Semiconductors (Ed: K.A. Jackson, Arizona, USA) □ Vol. 17 (1994): Processing of Ceramics (Ed: R.J. Brook, Oxford, UK)

VCH, 220 East 23rd Street, New York, NY 10010-4606, USA VCH, P.O. Box 10 11 61, D-6940 Weinheim, Germany VCH. Hardstrasse 10, P.O. Box, CH-4020 Basel, Switzerland VCH, 8 Wellington Court, Cambridge CB1 1HZ, UK

![](_page_16_Picture_23.jpeg)