# SCIENCE 3 July 1998 SCIENCE

Vol. 281 No. 5373 Pages 1–132 \$7

M

AMERICAN ASSOCIATION FOR THE ADVANCEMENT OF SCIENCE

# PfuTurbo

Supercharge your PCR

PfuTurbo
DNA Polymerase

Taq
DNA Polymerase



Stratagene's newly discovered PCR-enhancing factor has been specially formulated with cloned *Pfu* DNA polymerase to create *PfuTurbo* DNA polymerase.

- Same high fidelity as Pfu DNA polymerase
- Enhances PCR yield, amplifies longer targets and provides higher throughput performance
- Allows the use of less DNA template, fewer PCR cycles and shorter extension times

The Ultimate New Enzyme for High Fidelity PCR

PfuTurbo

PfuTurbo DNA Polymerase 100 units CATALOG # 600250 500 units CATALOG # 600252

www.stratagene.com

Heaction (PCH) process in conjunction with an authorized thermal cycler. Stratagene's PCR products are sold under licensing arrangements with Hoffmann-La Roche Inc.,

UNITED STATES AND CANADA

Switzerland: 01 800 9045 United Kingdom: 0171 365 1056

INTERNET MAIL techservices@stratagene.com

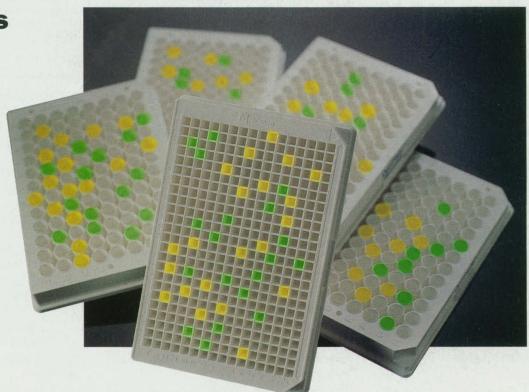
STRATAGENE EUROPE

Circle No. 27 on Readers' Service Card



### **Dual Luminescence Reporter Gene Assay Capability**

**Accelerates** the Pace of Drug **Discovery** 



### **FireLite™** from Packard

FireLite is the newest addition to Packard's Constant-Quanta™ glow luminescent product line. FireLite's homogeneous dual reporter assay system provides long-lived "glow" type signals, with a half-life up to five hours, for both firefly and renilla luciferase enzymes expressed in cells. Two reports from one well enable batch processing and the ability to screen over 100,000 assay points perday on TopCount® NXT™.

### **Double Your Cellular Expression Then Detect** with FireLite for:

- Long luminescent signal half-life (up to five hours)
- High sensitivity for both firefly (luc) and renilla (ren) luciferase enzymes
- · 96- and 384-well microplate compatible, dual reporter assays
- Batch processing on TopCount NXT
- Simple "mix-and-measure" procedures
- Elimination of injector-based methods

FireLite: The Simple, Stable and Sensitive Answer

for Luciferase Assays

Packard Instrument Company

800 Research Parkway Meriden, CT 06450 U.S.A. Tel: 203-238-2351 Toll Free: 1-800-323-1891 FAX: 203-639-2172

Web Site: http://www.packardinst.com Email: webmaster@packardinst.com

Packard International Offices

United Kingdom 44 (0)118 9844981

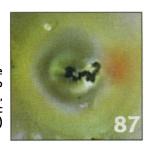
Australia 03-9543-4266 or 1 800 335 638; Austria 43-1-2702504; Belgium 32(0)2/481.85.30; Canada 1-800-387-9559; Central Europe 43 456 2230 015; Denmark 45-43909023 or 45-43907151; France (33) 1 46.86.27.75; Germany (49) 6103 385-151; Italy 39-2-33910796/7/8; Japan 81-3-3866-5850; Netherlands 31-50-549 1296; Russia 7-095-259-9632; Switzerland (01) 481 69 44;



Circle No. 17 on Readers' Service Card



COVER A composite, false-color image of Jupiter's moon to taken by the Galileo orbiter on 29 March 1998. Most of to's brilliant colors are due to sulfur compounds, but the dark features are probably silicate lava flows, many of which are associated with deposits (red) from explosive volcanic eruptions. The circular plume deposit from the volcano Prometheus (right) is ~250 kilometers in diameter. [Image processing: P. Geissler]





**26**Reconstructing the fusion program

DEPARTMENTS

NETWATCH
7
THIS WEEK IN
SCIENCE
9
SCIENCESCOPE
19
RANDOM SAMPLES
39
ESSAYS ON SCIENCE
AND SOCIETY
40
CONTACT SCIENCE
43
NEW PRODUCTS
113



150 YEARS • 1848-1998

DESCRIPTION OF THE PERSONS	FWS	NAME OF TAXABLE PARTY OF TAXABLE PARTY.	
17	NEWS OF THE WEEK	<b>▼23</b>	ARCHAEOLOGY: Eight Millennia of
16	SPENDING BILLS: U.S. R&D Budget  Becomes Political Football  Senate Bill Calls for More Spending	25	EVOLUTIONARY BIOLOGY: Successful Flies Make Love, Not War
₩ 17 82	GLOBAL CHANGE: Signs of Past Collapse Beneath Antarctic Ice		News Focus
19	SOLAR PHYSICS: Earth to SOHO, Come In Please	26	ENERGY RESEARCH: Competition Heats Up on the Road to Fusion Korea Brings U.S. Design to Life
20	GENOMICS: Canada Proposes \$175 Million Effort		Magnetic Fusion Researchers Think Small
20	SPACE: Remodeled ESA Backs	29	BIOLOGICAL WEAPONS: Arms Control Enters the Biology Lab
21	Applications Projects  SCIENTIFIC COMMUNITY: Panel Says Some UFO Reports Worthy of Study	31	HUMAN GENETICS: New Gene Found for Inherited Macular Degeneration
21	EPIDEMIOLOGY: NIH Panel Revives EMF- Cancer Link		SPECIAL FOCUS CARDIOVASCULAR DISEASE
22	MEDICAL ETHICS: No Consensus on Rules for AIDS Vaccine Trials	▼32 108	Tracking Down Mutations That Can Stop the Heart
23	WILDLIFE BIOLOGY: Fungus May Drive Frog Genocide	35	Infections: A Cause of Artery-Clogging Plaques?

	RESEARCH ARTICLE	75	Dissociative Recombination of HD+ in
<b>▼64</b> 58	Complete Structure of the 11-Subunit Bovine Mitochondrial Cytochrome bc <sub>1</sub> Complex S. Iwata, J. W. Lee, K. Okada, J. K. Lee, M. Iwata, B. Rasmussen, T. A. Link, S.		Selected Vibrational Quantum States Z. Amitay, A. Baer, M. Dahan, L. Knoll, M. Lange, J. Levin, I. F. Schneider, D. Schwalm, A. Suzor- Weiner, Z. Vager, R. Wester, A. Wolf, D. Zajfman
	Ramaswamy, B. K. Jap	78	An Inverted Hexagonal Phase of Cationic
	REPORTS		Liposome–DNA Complexes Related to DNA Release and Delivery I. Koltover, T.
₩72 23	7500 Years of Prehistoric Footwear from		Salditt, J. O. Rädler, C. R. Safinya
-	Arnold Research Cave, Missouri J. T. Kuttruff, S. G. DeHart, M. J. O'Brien	<b>82</b>	Pleistocene Collapse of the West Antarctic Ice Sheet R. P. Scherer, A. Aldahan, S. Tulaczyk, G. Possnert, H. Engelhardt, B. Kamb
	91		
	Mixer makes endoderm in the frog embry	85	Effects of Water on the α-β Transformation Kinetics in San Carlos Olivine T. Kubo, E. Ohtani, T. Kato, T. Shinmei, K. Fujino

SCIENCE (ISSN 0036-8075) is published weekly on Friday, except the last week in December, by the American Association for the Advancement of Science, 1200 New York Avenue, NW, Washington, DC 20005. Periodicals Mail postage (publication No. 484460) paid at Washington, DC, and additional mailing offices. Copyright © 1998 by the American Association for the Advancement of Science. The title SCIENCE is a registered trademark of the AAAS. Domestic individual membership and subscription (51 issues): \$108 (\$60 allocated to subscription). Domestic institutional subscription (51 issues): \$295. Foreign postage extra: Mexico, Caribbean (surface mail) \$55; other countries (air assist delivery) \$90. First class, airmail, student, and emeritus rates on request. Canadian rates with GST available upon request, GST #1254 88122. IPM #1069624. Printed in the U.S.A.

### SCIENCE'S COMPASS

### **EDITORIAL**

43 Elements of Our Design

### **LETTERS**

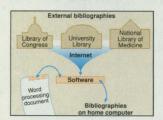
45 No Surprises? T. White; Response D. Falk; Lung Ventilation and Gas Exchange in Theropod Dinosaurs J. W. Hicks and C. G. Farmer; P. N. Nassar; R. Hengst; Response J. A. Ruben, T. D. Jones, N. R. Geist, W. J. Hillenius

### POLICY

49 SCIENCE PRIORITIES: The Scientific Investments of Nations R. M. May

### **BOOKS AND NEW MEDIA**

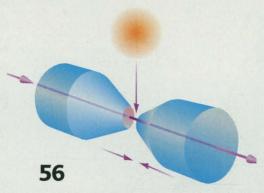
- 52 ECONOMICS AND TECHNOLOGY: The Productivity Payoff of Computers Y. Bakos
- 53 HISTORY: Los Alamos Stories H. Gusterson
- 53 Vignette: Citation Science S. Brenner
- 54 SOFTWARE: Order at the End B. Shmaefsky



54

Software for scientists ▼58 64

▼ 108



Setting the lightest atom trap

### **PERSPECTIVES**

- 55 EARTH'S INNER CORE: Is the Rotation Real?
  A. Souriau
- 56 ATOMIC PHYSICS: Under Control P. Grangier
- 57 CARCINOGENESIS: Another p53 Doppelgänger? W. G. Kaelin Jr.
  - STRUCTURAL BIOLOGY: Secret Life of Cytochrome bc<sub>1</sub> J. L. Smith

### **REVIEW**

Going the Distance: A Current View of Enhancer Action E. M. Blackwood and J. T. Kadonaga

### ONLINE PRODUCTS AND FEATURES

### SCIENCE

THE JOURNAL ONLINE www.sciencemag.org

### SCIENCENOW

DAILY NEWS SERVICE www.sciencenow.org

### **NEXT WAVE**

WEEKLY CAREER
UPDATES
www.nextwave.org

- 87 High-Temperature Silicate Volcanism on Jupiter's Moon Io A. S. McEwen, L. Keszthelyi, J. R. Spencer, G. Schubert, D. L. Matson, R. Lopes-Gautier, K. P. Klaasen, T. V. Johnson, J. W. Head, P. Geissler, S. Fagents, A. G. Davies, M. H. Carr, H. H. Breneman, M. J. S. Belton
- 91 Mixer, a Homeobox Gene Required for Endoderm Development G. L. Henry and D. A. Melton
- 96 Visualization of Specific B and T Lymphocyte Interactions in the Lymph Node P. Garside, E. Ingulli, R. R. Merica, J. G. Johnson, R. J. Noelle, M. K. Jenkins
- 99 C<sub>1</sub> Transfer Enzymes and Coenzymes Linking Methylotrophic Bacteria and Methanogenic Archaea L. Chistoserdova, J. A. Vorholt, R. K. Thauer, M. E. Lidstrom

- 103 Reproductive Dominance of Pasture Trees in a Fragmented Tropical Forest Mosaic P. R. Aldrich and J. L. Hamrick
- 105 Interaction of Human Arp2/3 Complex and the Listeria monocytogenes ActA Protein in Actin Filament Nucleation M. D. Welch, J. Rosenblatt, J. Skoble, D. A. Portnoy, T. J. Mitchison
  - Congenital Heart Disease Caused by Mutations in the Transcription Factor NKX2-5 J.-J. Schott, D. W. Benson, C. T. Basson, W. Pease, G. M. Silberbach, J. P. Moak, B. J. Maron, C. E. Seidman, J. G. Seidman

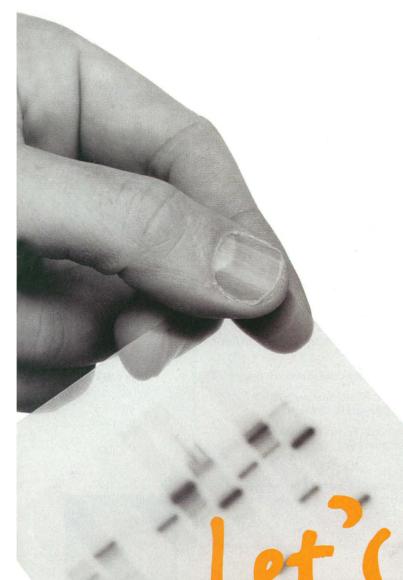
### **TECHNICAL COMMENTS**

Structure of β-iron at High Temperature and Pressure L. Dubrovinsky, S. K. Saxena, P. Lazor, H.-P. Weber; *Response* D. Andrault, G. Fiquet, M. Kunz, F. Visocekas, D. Haüsermann www.sciencemag.org/cgi/content/full/281/5373/11a



72 Sandal design in prehistoric times

Change of address: allow 4 weeks, giving old and new addresses and 8-digit account number. Postmaster: Send change of address to Science, P.O. Box 1811, Danbury, CT 06813–1811. Single copy sales: \$7.00 per issue prepaid includes surface postage; bulk rates on request. Authorization to photocopy material for internal or personal use under 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 \$4.00 per article is paid directly to CCC, 222 Rosewood Drive, Danvers, MA 01923. The identification code for Science is 0036-8075/83 \$4.00. Science is indexed in the Reader's Guide to Periodical Literature and in several specialized indexes.



Are your blotting signals getting blurred in the background noise? Are your tests as DNA-sensitive as they should be? Clearly, you could do with a little help.

### Hybond™

The advanced performance formula of Hybond-XL nylon membranes has been specifically designed to give you more information per gel.

You can look forward to outstanding signal-to-noise ratio: up to five times better than other membranes. You can detect minor bands more reliably and with shorter exposure times. So you get better results in even less time.

If you're already using a radioactive detection method for nucleic acid blotting and hybridisation, why compromise with the membranes you're using?

All clear? Now try Hybond-XL for free! Contact us today for your free sample of Hybond-XL or for more information about other Hybond products call I-800 526 3593 in the USA; in Europe (+44) (0) 1494 544550; from the rest of the world (+44) (0) 1494 544100. Or visit us on the web: www.apbiotech.com/hybond

few things (lear about Hubona-XI

Amersham Pharmacia Biotech UK Limited, Amersham Place, Little Chalfont, Buckinghamshire England HP7 9NA. All goods and services are sold subject to the terms and conditions of sale of the company within the Amersham Pharmacia Biotech group which supplies them A cory of these terms and conditions of sale is available on prequest.

Circle No. 36 on Readers' Service Card

amersham pharmacia biotech

### DNA makes protein. In a single tube.

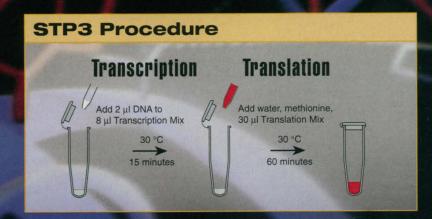
Novagen announces a *new* single-tube system for efficient *in vitro* production of proteins directly from supercoiled or linear DNA templates: Single Tube Protein™ System 3\* (STP3). Avoid 4–6 hours of lab work doing tasks required for standard methods of *in vitro* transcription and translation!

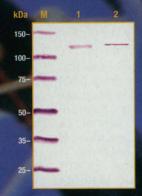
### Rapid and Versatile

- No restriction enzyme digestion, plasmid linearization, or RNA purification necessary
- Efficiently produces protein directly from PCR\*\* amplification products generated with appropriately designed primers containing T7 or SP6 promoters
- Two-step method outperforms coupled systems for PCR templates
- Use <sup>35</sup>S-methionine or non-radioactive detection with quantitative S•Tag<sup>™</sup> Rapid Assay or S•Tag Western Blot (with appropriate vectors)
- Compatible with S•Tag purification kits or other fusion tag affinity purification

### **Applications**

- Rapid testing of expression vector constructs
- Functional characterization of proteins without cloning
- In vitro analysis of protein:protein, protein:nucleic acid, protein:ligand interactions
- · Identification of open reading frames
- Screening for nonsense and frameshift mutations; protein truncation testing
- Compatible with PCR templates for colony screening, ligation PCR, RT-PCR samples, etc.





S•Tag Western Blot of crude and affinity-purified STP3 products. A standard STP3 reaction was performed with a pCITE-3 construct encoding β-gal fused with the 15 aa S•Tag peptide (used as an affinity tag for detection and purification due to its high affinity for the 104 aa S-protein). A portion of the reaction was analyzed directly, the remainder combined with S-protein agarose and incubated for 10 min at room temperature. Unbound proteins were washed away and the purified protein was eluted in SDS sample buffer. The western blot was developed with S-protein alkaline phosphatase and NBT/BCIP substrates. Lane M—Perfect Protein™ Markers (5 μl), Lane 1—crude STP3 reaction (3 μl), Lane 2—elution from S-protein agarose (20 μl).

To order introductory or standard STP3 T7 or SP6 Kits, call **800-526-7319**. Visit our web site at **WWW.NOVAGEN.COM** for more information.

\*Patent pending \*\* The PCR process is covered by patents owned by Hoffmann-LaRoche



www.novagen.com e-mail: novatech@novagen.com

800-526-7319 US & Canada

Novagen

### International Distributors

Australia · Progen Industries Ltd. 7-3375-1888

Europe (excluding UK) · Contact CalBiochem-NovaBiochem GmbH
Germany · CalBiochem-NovaBiochem GmbH 49-619663955

Hong Kong • PROTECH 886-22-3810844 Italy • M-Medical 055/5001871 Japan • Takara Shuzo Co., Ltd. 77-543-7231 Korea • BOHAN Biomedical 2-577-2002

Malaysia • BioSynTech Sdn Bnd 3-432-1357

New Zealand • Intermed Scientific Ltd. 9-443-1284

Singapore • IWAKI Glass Co., Ltd. 273-3022

Taiwan • PROTECH 22-3810844

UK • Cambridge Bioscience 1223-316855

Circle No. 31 on Readers' Service Card

### THIS WEEK IN SCIENCE

edited by PHIL SZUROMI

### **OLD SOFT SHOES**

Most of the evidence for early cultural evolution is from pottery or tools because these items are resistant to degradation and tend to be preserved, but records from more fragile items are important for providing a broader perspective. Kuttruff et al. (p. 72; see the news story by Pringle, p. 23) describe and have dated a remarkable collection of shoes preserved in deposits in Arnold Research Cave, Missouri. Eighteen shoes and sandals are complete or nearly complete and another 17 specimens are fragmentary. Together, the shoes provide a record of construction styles extending back to about 8000 years ago. Construction styles did not appear to become more complex with time; some earlier shoes were quite intricate, and all were made from grasses or woody fibers.

### SELECTIVE VIBRATIONS

Dissociative recombination of ions with free electrons is an important reaction in astrophysics and the upper atmosphere as well as in plasma processing and combustion. Experimental investigation of such reactions in molecular beams is hampered by the difficulty in generating sufficiently strong beams of vibrationally relaxed ions. Ion storage rings overcome this problem and allow molecular ions to relax to their vibrational ground state. Amitay et al. (p. 75) have extended this technique to allow determination of the product distribution as a function of vibrational excitation of the reactant ion. They studied the dissociative recombination of HD+ with an electron and show that rate coefficients generally increase for high vibrational excitations, where new dissociation routes become accessible. However, for isolated vibrational states, very low rate coefficients were observed that could not be reproduced by theoretical calculations, which suggests that the process is not fully understood.

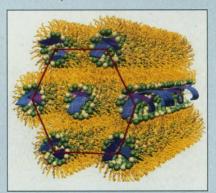
### SHIFTING ICE SHEET

Melting of the West Antarctic Ice Sheet would raise sea level by 5 to 6 meters. One clue to the stability of the ice sheet in response to current climate change is its past behavior during interglacial periods. To examine this question, Scherer et al. (p. 82; see the news story by Kerr, p. 17) drilled several holes through the ice sheet and collected glacial sediments from its bed. In several holes, the bed material contained Quaternary marine diatoms; these samples also had high con-

centrations of beryllium-10, a cosmogenic isotope with a half-life of 1.5 million years. Together, these data imply that the ice sheet receded greatly sometime during the last 1.3 million years and probably during the past 600,000 years. During that time, sediments containing the diatoms and <sup>10</sup>Be could be deposited upstream of the location of the drill holes.

### GETTING LIPOSOMES TO GIVE UP DNA

Recent studies have shown that complexes of DNA with univalent cationic liposomes (CLs) used for gene delivery can adopt a well-defined structure in which DNA is aligned between lamellar liposome sheets. Koltover et



al. (p. 78) used x-ray scattering to show that for DNA-CL ratios most favorable for gene transfer, a different structure forms in which the DNA molecules are encapsulated in liposome tubules. Optical microscopy revealed that this latter form rapidly fuses with anionic membranes to release DNA, while the lamellar complexes bind stably and retain DNA.

### TO SOFTEN, ADD WATER

Many minerals in the mantle can contain some water; one of these minerals is wadsleyite, which is abundant in the lower part of Earth's upper mantle between depths of 440 and 660 kilometers. In experiments simulating mantle pressures and temperatures, Kubo et al. (p. 85) show that even the addition of a small amount of water to wadsleyite greatly reduces its strength. Thus, even if some water is present in the mantle at this depth, the mantle could be weak, and considerably weaker than the strengths indicated from experiments conducted under dry conditions.

### SIZZLING SILICATES

The highest temperatures estimated for the surface of Io, a moon of Jupiter, by Voyager 1 in 1979 was about 650 kelvin (K). This temperature is not high enough to allow silicate volcanism as is observed on Earth but is within the temperature range for liquid sulfur; thus, sulfuric lava flows, lava lakes, and plumes were assumed to dominate the brilliant yellowish landscape of this volcanically active moon. The search for silicates on lo continued after Voyager with some observations of higher temperature "hot spots," but now McEwen et al. (p. 87) have used infrared wavelength observations from Galileo to estimate that at least a dozen hot spots have minimum temperatures exceeding 1700 K and the Pillan hot spot has a maximum temperature in excess of 2000 K (see the cover, which shows some of these hot spots colored in red). These high-temperature regions on lo indicate that silicate volcanism is prevalent on sulfur-covered Io and that some of these silicates are extremely hot compared to basaltic volcanism on Earth.

### **RELATIONSHIP FRAGMENTATION**

What effect does forest fragmentation have on the relationship between trees and their pollinators? Aldrich and Hamrick (p. 103) have discovered dramatic changes in plant fecundity (the shade tree Symphonia globulifera) and pollinator (hummingbird) behavior in the rain forest that had been fragmented during the past 10 to 30 years. The study used genetic analysis to determine the parentage of a large number of saplings and seedlings. Certain trees isolated in pasture greatly increased in fecundity and dominated the production of seedlings in the remnant forest. Hummingbird behavior was altered, resulting from increased flower production in pasture trees and leading to increased self-fertilization of these trees. These changes have led to a genetic bottleneck that has markedly constricted the plant donor pool.

### **ELECTRIC PUMPS**

Multisubunit enzyme complexes, the energy generators of the mitochondrion, use the downhill flow of electrons from NADH (the reduced form of nicotinamide adenine dinucleotide) to oxygen to pump protons across the mitochondrial membrane. Iwata et al. (p. 64; see the Perspective by Smith, p. 58) present the refined structure of the complete 11 subunit—complex III, also known as cytochrome

CONTINUED ON PAGE 11

### Within budget. Without compromise.

### Now available with MAX MODE

**Faster ramping and** higher sample volume.

Exceptional value Small footprint (1.3 square feet of bench space) Interchangeablesample blocks Fast, uniform heating and cooling Easy, intuitive programming

Circle No. 25 on Readers' Service Card

### The New GeneAmp® PCR System 9700

Until now, deciding on the best thermal cycler for your lab often meant choosing between superior performance and affordability.

Not any more.

Because now, there's an entirely new thermal cycler that gives you more of what you're looking for.

That thermal cycler is the GeneAmp® PCR System 9700.

> The GeneAmp PCR System 9700 combines the proven quality and reliability of the GeneAmp PCR System 9600 with unprecedented performance, the versatility of interchangeable sample blocks and a graphical user interface that streamlines operation.

> > But perhaps the most attractive feature of the GeneAmp PCR System 9700 is that it packs all of this into one compact instrument that fits easily on your lab bench-and into your budget.

> > > Find out how the GeneAmp PCR System 9700 makes choosing

the right thermal cycler easier than ever. In the U.S., call PE Applied Biosystems at 1-800-345-5224. Outside the U.S., contact your local sales representative. Or visit us on the Internet at www.thermalcycler.com.

### **PE** Applied Biosystems

Europe Langen, Germany Tel: 49 (0)6103 708 301 Fax: 49 (0)6103 708 310 Japan Tokyo, Japan Tel: (047) 380-8500 Fax: (047) 380-8505 Latin America Mexico City, Mexico Tel: 52-5-651-7077 Fax: 52-5-593-6223 Australia Melbourne, Australia Tel: 1 800 033 747 Fax: 61 3 9212-8502

Perkin-Elmer PCR reagents are developed and manufactured by Roche Molecular Systems, Inc., Branchburg, New Jersey, U.S.A.





The PCR process is covered by patents owned by Hoffmann-La Roche, Inc. and F. Hoffmann-La Roche Ltd. Perkin-Eimer is a registered trademark and PE Applied Blosystems, PE, and Applied Blosystems are trademarks of The Molecular Systems, Inc. PE Applied Blosystems per a registered trademark of Roche Molecular Systems, Inc. PE Applied Blosystems products are developed and produced under the quality requirements of ISO 9000.

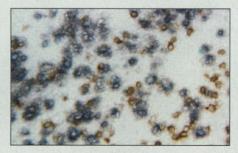
### THIS WEEK IN SCIENCE

CONTINUED FROM PAGE 9

bc<sub>1</sub>, which reveals how the electrons initially deposited by ubiquinone are transported to the acceptor cytochrome  $c_1$  by a 35° rotation of the "Rieske" subunit.

### LYMPH NODE RENDEZVOUS

T cells and B cells that are specific for an antigen must somehow contact each other in the lymph nodes, where the antigens



are usually found. Garside et al. (p. 96) have developed a system in which cognate interactions between antigen-specific lymphocytes can be visualized in situ. This approach has clarified the role of the CD40 ligand (CD154) in clonal expansion.

### **METHANE METABOLISM**

Anaerobic methanogenesis by Archaea and aerobic methane oxidation by methylotrophs were thought to be totally unrelated pathways. Chistoserdova *et al.* (p. 99) have found that these two pathways share some enzymes that are involved in processing single-carbon molecules, which is surprising because of the highly different ecological niches that these organisms occupy. The results are useful in understanding the evolution of oxidation/reduction pathways.

### **ACTIN TAIL NUCLEATION**

The pathogenic bacterium *Listeria mono-cytogenes* invades cells and then acquires a tail of actin that moves the bacterium

around the cell. A bacterial protein known as ActA is crucial for the formation of the actin tail. Welch et al. (p. 105) now describe in detail the role of a host cell protein complex that acts in concert with ActA to promote actin tail assembly. These findings shed light not only on the mechanism of *Listeria* invasion, but also suggest that endogenous host ActA-like proteins will be important in forming other polarized actin structures.

### **PATHWAY TO HEARTBREAK**

The four-chambered human heart initially develops as a two-chambered tube consisting of one atrium and one ventricle. Partitioning (septation) of these primordial chambers is critical for normal heart function. Mistakes are common, however; atrial septal defects (ASDs) occur in about 1 in 1500 live births. Schott et al. (p. 108; see news story by Barinaga, p. 32) show that a subset of ASDs, conduction defects, and other heart abnormalities are caused by mutations in the gene encoding the heart-specific transcription factor NKX2-5. Homologs of this gene have been implicated in heart development in fruit flies and mice.

### **DETERMINING ENDODERM**

Early in vertebrate embryonic development, the three germ layers are specified. These layers, ectoderm, mesoderm, and endoderm, each give rise to specific organs and tissues. Henry and Melton (p. 91) have identified a homeobox-containing gene, Mixer, that specifies the identity of the endodermal germ layer. The more specific anteroposterior pattern is specified by other downstream factors. These studies in Xenopus show that endoderm development follows a hierarchy of global definition followed by further specification that gives rise to endodermally derived organs.

### TECHNICAL COMMENT SUMMARIES

### Structure of $\beta$ -Iron at High T and P

The full text of these comments can be seen at www.sciencemag.org/cgi/content/full/281/5373/11a

D. Andrault *et al.* studied (31 Oct., p. 831) the structure of iron under high temperature (T) and pressure (P) in a laser-heated, diamond-anvil cell. They found that iron underwent a phase transformation and exhibited "an orthorhombic lattice."

L. Dubrovinsky et al. see "two problems with this conclusion." They state that the "method of applying structural refinement for the purpose of 'quantitative assessment of a structural model' is invalid." They also "question whether phase analysis of collected x-ray patterns can be interpreted as a mixture of known iron phases ..., iron oxide, and pressure medium."

Andrault *et al.* respond that they "favor an orthorhombic-iron explanation of the experimental features, which seems ... the most parsimonious," and they discuss each of the criticisms in turn.

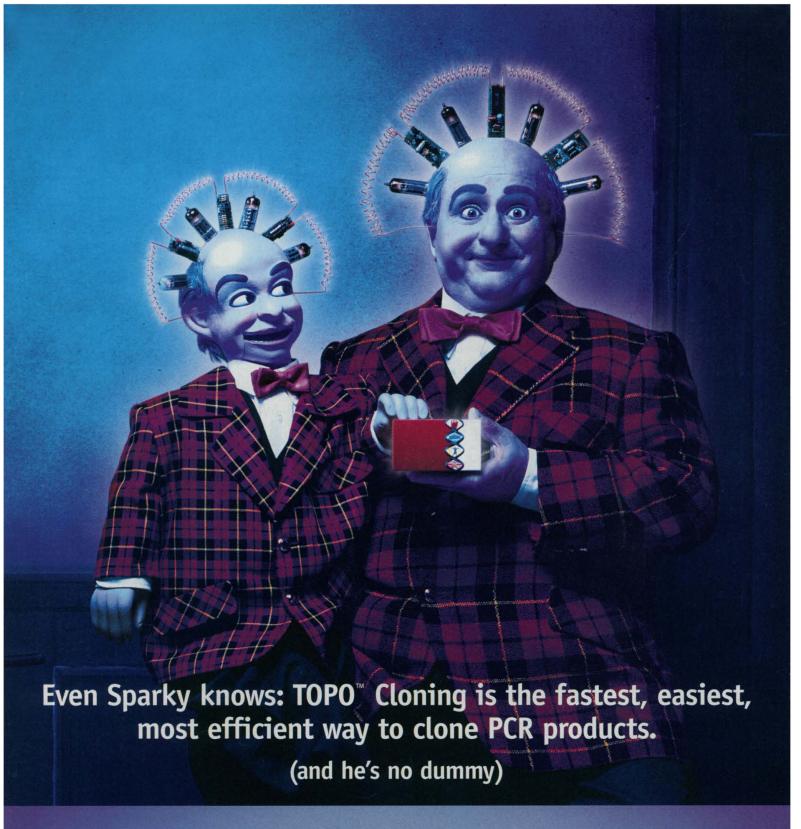


Use our new Online Reader Service card when you need information on products and services advertised in any issue of SCIENCE. Online Reader Service requests are sent instantly via the Internet to all the companies you select. This means you receive detailed product information...fast.

### www.sciencemag.org

Go to Electronic Marketplace & select Reader Service Card.





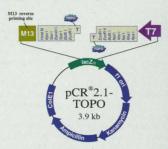
TOPO<sup>®</sup> Cloning is the only 5-minute, trouble-free method available to clone PCR products. TOPO<sup>®</sup> Cloning takes advantage of the unique activity of topoisomerase I to eliminate the hassles of conventional ligation methods. With TOPO<sup>®</sup> Cloning you can clone PCR products in just 5 minutes right on your bench top. TOPO<sup>®</sup> Cloning not only saves you time, it yields more recombinants than conventional ligation methods. Whether you need to just clone your PCR product, or clone and express it in *E. coli* or mammalian cells, Invitrogen offers a specially-designed TOPO<sup>®</sup> Cloning Kit so you can get great results quickly and easily.

### **CHOOSE YOUR POLYMERASE**

TOPO™ Cloning can be used to clone PCR products amplified from Tag, proofreading polymerases, or long polymerase mixtures. Choose your polymerase, then choose a TOPO™ Cloning Kit for 5-minute, high-efficiency cloning.

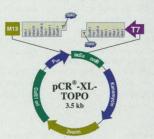
### Fast TA Cloning®

The TOPO™ TA Cloning® Kit is the fastest kit for cloning *Taq*-amplified PCR products. The pCR<sup>®</sup>2.1-TOPO vector contains 3'-T overhangs and is activated with topoisomerase so you can ligate PCR products in just 5 minutes and get up to 95% recombinants!



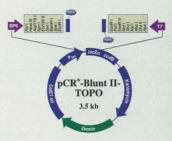
### Efficient Long PCR Cloning

The TOPO™ XL PCR Cloning Kit is the first kit specifically designed for efficient cloning of long (3-10 kb) PCR products. The pCR®-XL-TOPO vector and novel protocol let you clone long PCR products with efficiencies as high as 80% and low background! If you're cloning long PCR products, this is the kit for you!



### Easy Blunt-End PCR Cloning

Use the Zero Blunt™ TOPO™ PCR Cloning Kit to clone your blunt-end PCR products with 95% efficiency and low background. This unique kit combines Invitrogen's TOPO™ Cloning and Zero Background™ technologies to give you the easiest, most efficient way to clone blunt-end PCR products. No other blunt-end PCR cloning method even comes close.



### ONE-STEP CLONING AND EXPRESSION

TOPO™ Cloning and Expression Kits contain expression vectors that are TOPO™ Cloning ready to save you hours of ligation and cloning time. Now you can go straight from PCR cloning to expression without subcloning!

### **Express in Mammalian Cells**

The Eukaryotic TOPO™ TA Cloning® Kit contains the unique pcDNA3.1/V5-His-TOPO vector. This vector has all the elements you need to: clone your Tag-amplified PCR product in 5 minutes, efficiently express your protein in mammalian cells, rapidly purify your protein on nickel-chelating resin, and easily detect expression by western blot or ELISA. For direct expression of PCR products in mammalian cells, the Eukaryotic TOPO™ TA Cloning® Kit is the best choice!



### **Express in Bacterial Cells**

For TOPO™ Cloning and bacterial expression, you've got a choice. The pTrcHis2 TOPO™ TA Cloning® Kit is ideal for cloning and expressing non-toxic or soluble proteins. If your protein is toxic or tends to form inclusion bodies, try the pBAD TOPO™ TA Cloning® Kit. It uses the unique araBAD promoter to allow the tightest regulation of expression. Both kits offer 5-minute cloning of PCR products and direct expression in E. coli.



### TAKE IT FROM SPARKY

For the fastest, easiest, most efficient way to clone PCR products, get your hands on a TOPO™ Cloning Kit. With so many kits to choose from, there's guaranteed to be one that fits your needs. To find out more about these kits and our continually expanding line of TOPO™ Cloning products contact Invitrogen today. Ask for Sparky.



### represents covalently bound topoisomerase

### European Headquarters:

Invitrogen BV De Schelp 12, 9351 NV Leek The Netherlands Tel: +31 (0) 594 515 175 Fax: +31 (0) 594 515 312

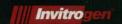
Email: tech\_service@invitrogen.nl

### Toll Free Phone Numbers:

### Distributors:

From all other countries, contact our European headquarters at +31 (0) 594 515 175.

United States Headquarters:



1600 Faraday Avenue Carlsbad, California 92008 Tel: 1-800-955-6288 Fax: 760-603-7201 Email: tech service@invitrogen.com http://www.invitrogen.com

### Science Science

EDITOR-IN-CHIEF Floyd E. Bloom

EDITOR Ellis Rubinstein

MANAGING EDITOR
Monica M. Bradford

### EDITORIAL

DEPUTY EDITORS Philip H. Abelson (Engineering and Applied Sciences); John I. Brauman (Physical Sciences); Thomas R. Cech (Biological Sciences)

ASSISTANT MANAGING EDITOR DAWN McCoy; SENIOR EDITORS Gilbert J. Chin, R. Brooks Hanson, Pamela J. Hines, Barbara Jasny, Paula A. Kiberstis, Linda J. Miller, L. Bryan Ray, Philip D. Szuromi; ASSOCIATE EDITORS Beverly A. Purnell, Linda R. Rowan; Editorial Assistant Carolyn Kyle; Manuscript Assistants Candace Gallery, Arny Herda, Josh Lipicky, Patricia M. Moore, Anita Wynn; Administrative support Sylvia Kihara; Computer Specialist Roman Frillarte

SCIENCE'S COMPASS: SENIOR EDITOR KATrina L. Kelner; ASSOCIATE EDITOR Sherman J. Suter; CONTRIBUTING EDITORS DAVID F. VOSS, KEVIN

Ahern; assistants Brent Gendleman, Jeffrey Hearn; information specialist Janet Kegg

LETTERS AND TECHNICAL COMMENTS: EDITOR Christine Gilbert; ASSOCIATE EDITOR Steven S. Lapham; ASSISTANT Charlene King

TECH.SIGHT: CONTRIBUTING EDITORS RICHARD PETERS, ROBERT SIKORSKI EDITING: SUPERVISOR CARA TATE; SENIOR COPY EDITORS HARTY Jach, Christine M. Pearce; COPY EDITORS: Jeffrey E. Cook, Etta Kavanagh, Jason Llewellyn, Joshua Marcy

COPY DESK: SUPERVISOR Ellen E. Murphy: Joi S. Granger, Abigail Hollister, Monique Martineau, Beverly Shields; Assistant Jessica Moshell

### NEW5

NEWS EDITOR Colin Norman; FEATURES EDITOR TIM Appenzeller; DEPUTY NEWS EDITORS Elizabeth Culotta (contributing editor), Jean Marx, Jeffrey Mervis, Richard Stone; NEWS WRITERS Jennifer Couzin (intern), Constance Holden, Jocelyn Kaiser, Richard A. Kerr, David Kestenbaum, Andrew Lawler, Eliot Marshall, Elizabeth Pennisi, Robert F. Service, Gretchen Vogel Burreaus-Berkreier, ca. Marcia Barinaga (contributing correspondent); SAN DIEGO, CA Jon Cohen; CHICAGO, IL James Glanz; COPY EDITORS Linda B. Felaco, Daniel T. Helgerman; CONTRIBUTING CORRESPONDENTS BAITY A. Cipra, Ann Gibbons, Charles C. Mann, Anne Simon Moffat, Virginia Morell, Gary Taubes, Ingrid Wickelgren; Administrative Support Scherraine Mack, Fannie Groom

### PRODUCTION

DIRECTOR James Landry; MANAGER Wendy K. Shank; ASSISTANT MANAGER Lizabeth A. Harman; ASSOCIATES Clarence A. Foules, Vicki J. Jorgensen, Cynthia M. Penny, Kameaka Williams

ART

DESIGN DIRECTOR AMY DECKER HENRY; ART DIRECTOR C. Faber Smith; ASSOCIATE ART DIRECTOR Elizabeth Carroll; SCIENTIFIC ILLUSTRATOR KAtharine Sutliff; GRAPHICS ASSOCIATES HOlly Bishop, Preston Morrighan, Darcel Pugh, Patricia M. Riehn; PHOTO RESEARCHER Leslie Blizard; TECHNOLOGY MANAGER Christopher J. Feldmeier

### SCIENCE INTERNATIONAL

### EUROPE OFFICE

EDITORIAL: OFFICE HEAD AND SENIOR EDITOR RICHARD B. Gallagher; ASSOCIATE EDITORS Stella M. Hurtley, Peter Stern, Julia Uppenbrink; EDITORIAL ASSOCIATE BEIINDA HOIDEN NEWS: EDITOR Daniel Clery; CORRESPONDENT Nigel Williams; CONTRIBUTING CORRESPONDENT Michael Balter (Paris); UK EDITOR, SCIENCE'S NEXT WAVE JOHN MACFARlane; ADMINISTRATIVE SUPPORT JANET MUMFORD, LIZ Ellis

### ASIA OFFICE

JAPAN NEWS BUREAU: CONTRIBUTING CORRESPONDENT Dennis Normile; CHINA REPRESENTATIVE HAO XIN

SCIENCENOW: www.sciencenow.org

SCIENCE'S NEXT WAVE: www.nextwave.org
MANAGING EDITOR WENDY Yee; ASSOCIATE EDITOR NICOLE RUEDIGER
WRITER Melissa Mertl; CANADA EDITOR CHARLES BOULAKIA

### PUBLISHER Richard S. Nicholson

ASSOCIATE PUBLISHER
Beth Rosner

MEMBERSHIP/CIRCULATION DIRECTOR
Michael Spinella

MEMBERSHIP/CIRCULATION DEPUTY DIRECTOR Marlene Zendell

MEMBER SERVICES: MANAGER Michael Lung: SUPERVISOR Mary Curry; REPRESENTATIVES PAt Butler, Laurie Baker, Jonathan Keeler, Mari Pope, Jantell Smith

MARKETING: COORDINATOR LAURI SIROIS; EUROPE MANAGER Jane Pennington; REPRESENTATIVE BEN HOlland RESEARCH: MANAGER RENUKA Chander

BUSINESS AND FINANCE: ASSISTANT SUSAN Maxim; COMPUTER SPECIALIST Charles Munson

### FINANCE AND ADVERTISING

BUSINESS AND FINANCE: BUSINESS MANAGER Deborah Rivera-Wienhold; SENIOR ANALYST RANDY YI; FINANCIAL ANALYST CONNIE DANG PERMISSIONS: ADMINISTRATOR LINCOLN RICHMAN, ASSISTANT EMILIE DAVID MARKETING: DIRECTOR John Meyers; ASSOCIATES Allison Pritchard, Chris Harbaugh Electrophic Media: Manager David Gillikin; Computer Specialist Wendy Green; Production Associates Mark Croatti, Crystal Young

### PRODUCT ADVERTISING

ACTING NATIONAL SALES MANAGER E. COAST AND E. CANADA RICHARD TEELING: 973-904-9774, FAX 973-904-9701 • MIDWEST/SOUTHEAST ELIZABETH MOSKO: 773-665-1150, FAX 773-665-1229 • WEST COAST/W. CANADA NEIL BOYLAN: 415-673-9265, FAX 415-673-9267 • U.S. INSIDE SALES Christopher Breslin: 202-326-6544, FAX 202-682-0816 • UK/SCANDINAVIA/FRANCE/ITALY/BELGIUM/NETHERLANDS ANDREW DAVIES: (44) 1-457-871-073, FAX (44) 1-457-877-344 • CERMANY/SWITZFRIAND/ALISTRIA Tracey Peers: (44) 1-260-297-530, FAX (44) 1-260-271-022 JAPAN MASHY YOSHIKAWA: (81) 3-3235-5852 • TRAFFIC MANAGER CAROL MADDOX; SALES ASSOCIATES Sheila Myers, Sandra Walls

### RECRUITMENT ADVERTISING

SALES AND PRODUCTION OPERATIONS MANAGER TERRI Seiter AZIE U.S.: SALES MANAGER Gabrielle Boguslawski: 718-491-1607, FAX 202-289-6742; SALES SUPERVISOR DARYL Anderson; SALES REPRESENTATIVES ERIC Banks, Troy Benitez, Beth Dwyer, Bren Peters-Minnis; Assistants Erika Bryant, Kathleen Clark, Angela Panton; Production Associates Ellen McGuire, Jennifer Rankin; Copy Editor/Proofreader Chris Filiatreau U.K./
EUROPE: SALES MANAGER Debbie Cummings; SALES EXECUTIVE Sabine Lenud; Assistant Michaela Heigl; (44) 1-223-302-067, FAX (44) 1-223-576-208 AUSTRALIA/NEW ZEALANDE Keith Sandell: (61) 02-9922-2977, FAX (61) 02-9922-1100 JAPAN: Mashy Yoshikawa: (81) 3-3235-5961, FAX (81) 3-3235-5852

### **AAAS BOARD OF DIRECTORS**

RETIRING PRESIDENT, CHAIR Mildred S. Dresselhaus PRESIDENT M. R. C. Greenwood PRESIDENT-ELECT Stephen Jay Gould TREASURER William T. Golden EXECUTIVE OFFICER RICHARD S. Nicholson

Robert D. Goldman; Alice S. Huang; Sheila Jasanoff; Sally Gregory Kohlstedt; Marcia C. Linn; Michael J. Novacek; Neena B. Schwartz; Jean E. Taylor

Published by the American Association for the Advancement of Science (AAAS), Science serves its readers as a forum for the presentation and discussion of important issues related to the advancement of science, including the presentation of minority or conflicting points of view, rather than by publishing only material on which a consensus has been reached. Accordingly, all articles published in Science—including editorials, news and comment, and book reviews—are signed and reflect the individual views of the authors and not official points of view adopted by the AAAS or the institutions with which the authors are affiliated.

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

### Frederick W. Alt Children's Hospital, Boston Don L. Anderson California Institute of Technology Michael Ashburner

University of Cambridge
Frank S. Bates
Univ. of Minnesota, Minneapolis

Univ. of Minnesota, Minneapolis Stephen J. Benkovic Pennsylvania State University

Alan Bernstein
Mount Sinai Hospital, Toronto
Michael J. Bevan
University of Washington,
Seattle

Seth Blair
University of Wisconsin, Madison
David E. Bloom
Harvard Institute for

Harvard Institute for International Development Piet Borst The Netherlands Cancer Institute

Henry R. Bourne
Univ. of California, San Francisco
James J. Bull
University of Texas at Austin
Kathryn Calame

Columbia Univ. College of Physicians & Surgeons Dennis W. Choi Washington Univ. School of Medicine, St. Louis David Clapham Children's Hospital, Boston Adrienne E. Clarke
University of Melbourne, Parkville
F. Fleming Crim
University of Wisconsin, Madison
Paul J. Crutzen
Max-Planck-Institut für Chemie
Jemes E. Dahlberg
University of Wisconsin Medical
School, Madison

School, Madison
Robert Desimone
National Institute of Mental
Health, NIH
Hans Eklund
Swedish Univ. of Agricultural

Sciences, Uppsala
Paul T. Englund
Johns Hopkins University
School of Medicine
G. Ertl
Max-Planck-Gesellschaft

Max-Planck-Gesellschaft Richard G. Fairbanks Lamont-Doherty Earth Observatory Douglas T. Fearon University of Cambridge

Observatory
Douglas T. Fearon
University of Cambridge
Harry A. Fozzard
The University of Chicago
Roger I. M. Class
Centers for Disease Control
Peter N. Goodfellow
SmithKline Beecham, UK
Jack F. Greenblatt

University of Toronto
Peter Gruss
Max Planck Institute of
Biophysical Chemistry

BOARD OF REVIEWING EDITORS

Philip C. Hanawalt Stanford University Paul Harvey University of Oxford M. P. Hassell Imperial College at Silwood Park Nobutaka Hirokawa University of Tokyo Tomas Hökfelt Karolinska Institutet

Karolinska Institutet
Tasuku Honjo
Kyoto University
Susan D. Iversen
University of Oxford
Eric F. Johnson
The Scripps Research Institute
Hans Kende
Michigan State University
Elliott Kleff

Elliott Kieff Harvard University Jeffrey T. Kiehl National Center for Atmospheric Research, Boulder Judith Kimble University of Wisconsin, Madison Stephen M. Kosslyn Harvard University Michael LaBarbera

Michael LaBarbera
The University of Chicago
Antonio Lanzavecchia
Basel Institute for Immunology
Nicole Le Douarin
Institut d'Embryologie Cellulaire
et Moléculaire du CNRS

Norman L. Letvin

Beth Israel Hospital, Boston

Harvey F. Lodish
Whitehead Institute for
Biomedical Research
Richard Losick
Harvard University
Seth Marder
California Institute of
Technology
Diane Mathis
Institut de Chimie Biologis
Strasbourg
Susan K. McConnell

Jiane Maunis
Institut de Chimie Biologique,
Strasbourg
Susan K. McConnell
Stanford University
Anthony R. Means
Duke University Medical
Center
Stanley Meizel
University of California, Davis
Douglas A. Melton
Harvard University
Andrew Murray
Univ. of California, San Francisco
Elizabeth G. Nabel
The Univ. of Michigan Medical

Univ. of California, San Francisco Elizabeth G. Nabel The Univ. of Michigan Medical Center Shigetada Nakanishi Kyoto University Kin Research Institute of

Kim Nasmyth
Research Institute of
Molecular Pathology, Vienna
Roger A. Nicoll
Univ. of California, San Francisco
Staffan Normark

Swedish Institute for Infectious Disease Control Kiyotaka Okada Kyoto University Martin Ráff
University College London
Douglas C. Rees
California Institute of
Technology
T. M. Rice
FIH-Hönggerberg, Zürich
David C. Rubie
Universitä Bayveuth
Erkki Ruoslahti
The Burnham Institute, CA
Gottfried Schatz
Biozentrum, Basel
Jozef Schell
Max-Planck-Institut für
Zuchtungforschung
Ronald H. Schwartz
National Institute of Allergy
and Infectious Diseases, NiH
Errence, I. Sejnowski
Salk Institute

Bert W. O'Malley
Baylor College of Medicine
Roy R. Parker

Roy R. Parker
University of Arizona, Tucson
Stuart L. Pimm
The Univ. of Tennessee, Knoxville
Yeshayau Pocker
Univ. of Washington, Seattle

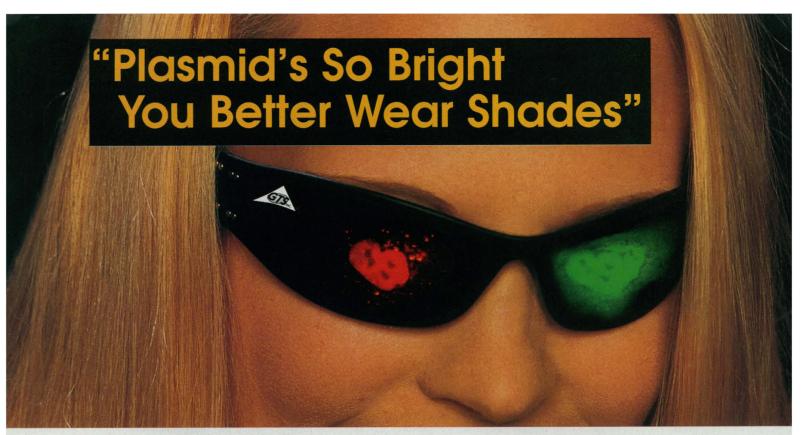
National Institute of Allergy and Infectious Diseases, NIH Ternence | Sejnowski Salk Institute Edward E. Smith Univ. of Michigan, Ann Arbor Christopher R. Somerville Carnegie Institute of Washington Michael P. Stryker Univ. of California, San Francisco Cliff Tabin Harvard Medical School John Jen Tai
Academia Sinica, Taiwan
Tomoyuki Takahashi
University of Tokyo
Masatoshi Takelchi
Kyoto University
Keiji Tanaka
RIKEN Institute
David Tilman
Univ. of Minnesota, St. Paul
Robert T. N. Tjian
Univ. of California, Berkeley
Yoshinori Tokura
University of Tokyo
Derek van der Kooy
University of Toronto
Geerat J. Vermeij
University of Galifornia, Davis
Bert Vogelstein
Johns Hopkins Oncology Center
Gerhard Wegner
Max-Planck-Institut für
Polymerforschung
Arthur Weiss
Univ. of California, San Francisco
Zena Werb
Univ. of California, San Francisco
George M. Whitesides
Harvard University
Ian A. Wilson
The Scripps Research Institute
National Institute of Child Health
and Human Development, NiH

Martin Zatz

National Institute of Mental

Health, NIH

14



### Introducing: pGeneGrip™Fluorescent vectors

- Transcriptionally active fluorescent DNA
- Conformationally intact & supercoiled
- No labeling or purification required
- Simple and ready to use

pGeneGrip™ vectors\* are a unique series of modified plasmids. Taking advantage of our innovative DNA tagging procedure, the pGeneGrip™ vector is efficiently and irreversibly labeled without changing its supercoiled structure or transcriptional activity. pGeneGrip™ vectors are a breakthrough product line for gene delivery and gene therapy research. For the first time, scientists have the opportunity to simultaneously follow biodistribution of the plasmid DNA and expression of the encoded transgene product.

Our pGeneGrip™ vectors are currently available with Fluorescein, Rhodamine or Biotin labels which encode GFP, β-Galactosidase or Secreted Alkaline Phosphatase.

### pGeneGrip™ Rhodamine/GFP vector

• 25 μg Catalog #G101040 • 100 μg Catalog #G101045

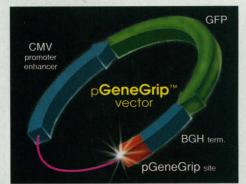


To Order Call: 888-428-0558 Fax: 619-587-1499

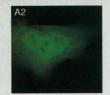
4370 La Jolla Village Drive, Suite 960 San Diego, CA 92122

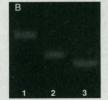
For the complete product list check out the GTS Website @ http://www.genetherapysystems.com

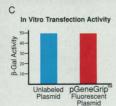
International distribution inquires welcome Circle No. 35 on Readers' Service Card











- A. Fibroblasts transfected with pGeneGrip™ Rhodamine/GFP vector: 1. Rhodamine labeled DNA
- 2. GFP expression
- B. Electrophoresis of pGeneGrip™ Rhodamine labeled fluorescent vector Lanes: 1. β-gal, 2. GFP, 3. Blank
- C. Plasmid expression with and without fluorescent label.
- \* Patent pending



# Innovation and Precision in Nucleic Acid Synthesis



Since 1987, more than 10,000 customers have come to trust IDT.
Whether you need complex modified

probes or basic primers, IDT has the expertise to deliver the quality and purity your research demands. Our experienced scientists bring a higher standard to oligonucleotide synthesis.

### IDT Can Provide You with Oligos Incorporating any Available Modification.

- Chimeric Antisense Oligos™
- Extra Long Sequences
- Dual-Labeled Fluorescent Probes
- RNA & 2'-O-Me Synthesis and More

Attention ABI & ALF Sequencer Users! IDT is the place for the full spectrum of Dye-Labeled Primers!

### Call or Click IDT Today!

Call IDT Toll Free: 1-800-328-2661

Fax: 1-319-626-8444
Website: www.idtdna.com
E-Mail: orders@idtdna.com

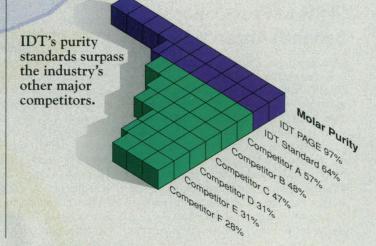
### Custom DNA Synthesis - 85¢ / Base

- No Set-up Fee/No Charge for Desalting
- 100 nmole scale
- 24 Hour Shipping

The Data Is In: For oligonucleotides longer than 40-bases, "Standard" (desalting-type) purification is not enough. You should add PAGE purification and we've made it affordable at just \$45.00.

# Capillary Electrophoresis Results Full width: 11 minutes to 23 minutes Full height: 0 to 0.9 AU Molar Purity\* IDT PAGE 97% IDT Standard 64% Competitor 28% \*Full length of product divided by the sum of all product. IDT PAGE Purified IDT Standard Competitor

Analysis of the same 40-mer sequence comparing IDT's PAGE purified synthesis to standard oligos from IDT and six major competitors. Please visit our web site to download a complete report.





### PCR optimization in one single experiment

authorized for PCR.



300 bp

The determination of the right annealing temperature is crucial for establishing a new PCR experiment. The new all-round genius Mastercycler® gradient is an innovative, compact thermal cycler for even the most demanding PCR\* applications. Its gradient function enables a temperature gradient of up to 20 °C to be generated across the block, thus optimizing the annealing, denaturation, or extension temperature in one single experiment. Its 96-position all-in-one universal block can accommodate 96 x 0.2 ml tubes, 77 x 0.5 ml tubes, or one 96-well plate - without block exchange.

The exact block homogeneity ensures reproducible results. Variable, high incubation speeds as well as time and temperature increments for hot-start, touch-down, and long PCR provide maximum programming flexibility.

The Mastercycler® personal is designed to meet personal applications. Its 25-position all-in-one universal block holds 25 x 0.2 ml tubes, 16 x 0.5 ml tubes, or one microplate in a 5x5 grid.

Personal memory cards for 10 individually developed protocols allow an easy and comfortable method transfer between both Mastercycler models.

PCR (Polymerase Chain Reaction) is protected by patent. The patent is held by Hoffmann-La Roche. Practice of the patented Polymerase Chain Reaction (PCR) process requires a license. The Eppendorf Thermal Cyclers are Authorized Thermal Cyclers and may be used with PCR licenses available from the Perkin-Einer Corporation. Their use with Authorized Reagents also provides a limit PCR license in accordance with the label rights accompanying such reagents.

Experimental determination of optimal annealing temperature. The calculated primer annealing temperature was 56.5 °C, the

of mycoplasms from H9 cell cultures was amplified.

actual annealing temperature is 63.5 °C. The ribosomal spacer region

Of course, both Mastercyclers are licensed and

1 2 3 4 5 6 7 8 9 10 11 12 13 14

53°C

Circle No. 15 on Readers' Service Card



### The Leading Edge in Neuroscience

For over 18 years, neuroscientists have depended on the quality, expertise and service of RBI. Today, our company manufactures and distributes over 1,600 chemical and biochemical products for investigating cell signaling pathways in the central and peripheral nervous systems.

### PRODUCTS FOR SIGNALING PATHWAYS

AGONISTS
ANTAGONISTS
ANTIBODIES
CHEMOKINES

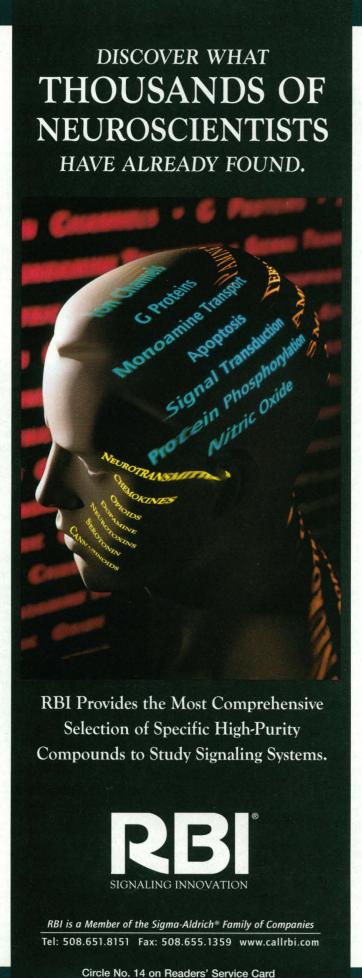
ENZYME INHIBITORS

NEUROTOXINS

RECEPTORS

### **Pure Excellence**

The consistent quality and high purity of RBI products are recognized worldwide. All are backed with detailed documentation, including structures, physical properties and references to protocols.



### APPLICATIONS

Ca2+ SIGNALING

PROTEIN PHOSPHORYLATION

ION CHANNEL MODULATION

CELL STRESS/ NO SYSTEMS

APOPTOSIS

G PROTEINS

IP3/DAG

### Expertise, Support and Service You Can Trust

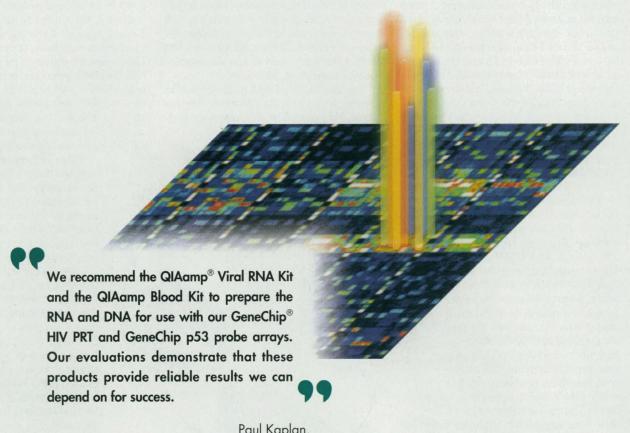
We develop and manufacture many of our products, so you can count on us for expert technical support and responsive customer service.

### Make the RBI Discovery Today

The 1998 RBI Catalog features over 1,600 research products for neuroscience, signal transduction and peripheral nervous system studies. We offer the most complete listing of high-purity neurochemicals in the world.

For a free catalog, call us at 800-736-3690 or 508-651-8151 or visit our web site at: www.callrbi.com. RBI — it could be the most important discovery you make today.

## QIAGEN® sample preparation gives Affymetrix® results they can rely on for success.



Paul Kaplan, VP Product Development, Affymetrix

High-throughput genetic analysis on GeneChip probe arrays is becoming routine, and Affymetrix is leading the way. When researchers at Affymetrix needed a reliable method to purify RNA and DNA to ensure the success of each round of analysis, they looked to QIAGEN, the leader in nucleic acid sample preparation.

Purity, reliability, service — no wonder leaders choose QIAGEN!



http://www.giagen.com



# SHAKEN STIRRED!





For further information and literature references tel: 800-424-6101

fax: 760-598-0116

www.bio101.com • info@bio101.com



FastPrep

"Ce n'est pas un centrifuge"

a closed system for a wide variety of applications. The FastPrep System employs a rapid oscillating motion and combination of proprietory matrices and chaotropic reagents to simultaneously homogenize tissues, lyse cells and stabilize nucleic acids in seconds.

The unit is used with FastRNA and FastDNA extraction kits.

- FastRNA® BLUE Kit (cat# 6020-600) Total RNA Isolation from all Bacteria including Gram positive strains.
- FastRNA® RED Kit (cat# 6030-600) Total RNA Isolation from Yeast, Fungi and Algae.
- FastRNA® GREEN Kit (cat# 6040-600) Total RNA Isolation from Plant and Animal Tissues.

FastDNA® Kit (cat# 6540-400) DNA Isolation from Any Source for PCR. Also, special FastDNA SPIN Kit for Soil Organisms (cat# 6560-200)

The Name's BIO, **BIO 101** 

### **Our Agents are No Secret**

International Distributors: Australia 800-252204 • Austria [43] 1 292 35 27 • Benelux [31] 76 579 5795 • Brazil [55] 11 663565 • Canada 800-387-8125 • Czech Republic [42] 02 758 635 • China [86] 10 6253-2114 Denmark [45] 39271777 ● France [33] 1 34 60 24 24 ● Germany [49] 40 45 06 70 ● Greece [30] 1 52 54 157 ● Hong Kong [852] 2898 3008 ● India [91] 33 472 9425 and [516] 796 2163 [in USA] ● Israel [972] 2 6520279 ● Italy: Dasif S.p.A. [39] 2 93991.1 / Stepbio S.R. L [39] 51 63 43 340 ● Japan [81] 3 5684-1622 ● Korea: BMS Co., Ltd. [82] 2 569 6902 / Koram Biotech Corp. [82] 2 556 0311 ● Malaysia [603] 432-1357 ● Mexico: Bioselec [52] 5 355 7193 / Control Tecnico y Representaciones [52] 8 371 6050 ● New Zealand [64] 09 443 5867 ● Norway [47] 22 25 5054 ● Poland [48] 58341 21 43 ● Singapore [65] 273 0898 ● South Africa [27] 21 61-5166 ● Spain [93] 490 74 40 ● Sweden [46] 31 706 3000 Switzerland (41) 41 4209636 ◆ Taiwan: Cold Spring Biotechnology Co., Ltd. (886) 22 695 9990 / Cheng Chin Trading Co., Ltd. (886) 22331 3111 ◆ Thailand (66) 2 412 5672 ◆ United Kingdom (44) 1 582 456666

\* Patents issued and pending

Circle No. 29 on Readers' Service Card

### **CAMBRIDGE: YOUR SOURCE FOR SCIENCE SCHOLARSHIP**

### Neuroimaging and the Psychiatry of Late Life

**David Ames** and **Edmond Chiu,** Editors

This book provides clinicians with a reliable reference, written by prominent figures in neuroradiology and old age psychiatry, which draws together current knowledge of late life mental disorders as revealed by neuro-imaging. A highly illustrated introductory chapter provides a useful overview of the various techniques of neuroimaging now available.

1998 252 pp. 0-521-49505-9 Hardback \$95.00

### Inclusion-Body Myositis and Myopathies

Valerie Askanas, Georges Serratrice, and W. King Engel, Editors

Inclusion-body myositis (IBM) is now understood to be an important degenerative muscle disease. The sporadic type (s-IBM) is probably the most common muscle disease among those ailments that strike first in adulthood (particularly people over 50). The hereditary type (h-IBM) affects younger patients. This book is devoted entirely to s-IBM and h-IBM. Contributors discuss what is understood about the basic scientific foundations of IBMs, the varied aspects of the pathology of IBMs, and the application of clinical treatments.

1998 416 pp. 0-521-57105-7 Hardback \$125.00

### Development of Cardiovascular Systems

Molecules to Organisms Warren W. Burggren and Bradley B. Keller, Editors

This volume is a unique overview of cardiovascular development from the cellular to the organ level across a broad range of species. The first section focuses on the molecular, cellular, and integrative mechanisms that determine cardiovascular development. The second section has eight chapters that summarize cardiovascular development in invertebrate and vertebrate systems. The third section discusses the effects of disease and environmental and morphogenetic influences on nonmammalian and mammalian cardiovascular development.

0-521-56072-1 Hardback \$74.95

Circle No. 1 on Readers' Service Card

Available in bookstores or from

### Linking Social and Ecological Systems

Management Practices and Social Mechanisms for Building Resilience Fikret Berkes and Carl Folke, Editors

Developed under the auspices of the Beijer Institute in Stockholm, this new book analyzes social and ecological linkages in selected ecosystems using an international and interdisciplinary case study approach. The chapters provide detailed information on a variety of management practices for dealing with environmental change.

1998 476 pp.

0-521-59140-6 Hardback \$80.00

### Molecular Genetics of Plant Development Stephen H. Howell

The purpose of this textbook is to present classical plant development in modern, molecular-genetic terms. This book provides a framework for integrating gene discovery and genome analysis into the context of plant development. Taking a systems approach, concepts in plant development are compared to those in animal development, and complex processes, such as flowering and photomor-phogenesis, are presented as pathways of gene action regulated by positional and environmental cues.

### The Cambridge Encyclopedia of Human Paleopathology

0-521-58784-0 Paperback about \$39.95

Arthur C. Aufderheide and Conrado Rodriguez-Martin

The Cambridge Encyclopedia of Human Paleopathology is a major reference work for all those interested in the identification of disease in human remains. Many diseases leave characteristic lesions and deformities on human bones, teeth and soft tissues that can be identified many years after death. This comprehensive volume includes all conditions producing effects recognizable with the unaided eye. Detailed lesion descriptions and over 300 photographs and diagrams facilitate disease recognition and each condition is placed in context with discussion of its history, antiquity, etiology, epidemiology, geography and natural history.

1998 496 pp.

0-521-55203-6 Hardback 100.00

### An Introduction to Vascular Biology

From Physiology to Pathophysiology Alison Halliday, Beverley J. Hunt, Lucilla Poston, and Michael Schachter, Editors

The multidisciplinary team of contributors covers topics ranging from normal and pathological aspects of endothelial function to the role of the vasculature in hemostasis, atherosclerosis and hypertension. This carefully illustrated and highly readable text provides both a valuable source of practical information and clear explanations of the impact of new techniques of cellular and molecular biology on recent and future developments.

1998 286 pp. 0-521-58998-3 Paperback \$39.95

### Comprehension A Paradigm for Cognition

A Paradigm for Cognition Walter Kintsch

In this book, Walter Kintsch presents a theory of human text comprehension and extends his analysis to related areas. In Part I, the general theory is presented and an attempt is made to situate it within the current theoretical landscape in cognitive science. The second part addresses many of the topics that are typically found in a cognitive psychology text, including how word meanings are identified in a discourse context; how words are combined to form coherent representations of texts, both at the local and global level; what the role is of working memory in comprehension; how relevant knowledge is activated during reading; and what is the distinction between remembering a text and learning from a text. 478 pp.

0-521-62986-1 Paperback \$27.95

### Handbook of Ethological Methods

Second Edition

Philip N. Lehner

This new expanded edition of the *Handbook* of *Ethological Methods* provides a complete step-by-step introduction to ethological methods from topic choice and behavioral description to data collection and statistical analysis. This book is a must for both beginning students and experienced researchers studying animal behavior in the field or laboratory.

1998 692 pp.

0-521-63750-3 Paperback \$34.95

CAMBRIDGE UNIVERSITY PRESS

40 West 20th Street, New York, NY 10011-4211 Call toll-free 800-872-7423.

The Edinburgh Building, Shaftesbury Road, Cambridge CB2 2RU, U.K.

Web site: http://www.cup.org MasterCard/VISA accepted. Prices subject to change.