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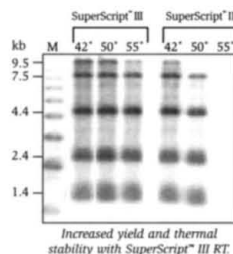
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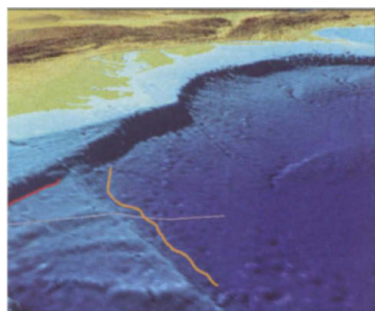
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Science

Volume 298 6 December 2002 Number 5600

1841 **SCIENCE ONLINE**
1843 **THIS WEEK IN SCIENCE**
1847 **EDITORIAL**
Kirk R. Smith
In Praise of Petroleum?

1849 **EDITORS' CHOICE**
1853 **NETWATCH**
1856 **CONTACT SCIENCE**
2021 **NEW PRODUCTS**



1877

Finding an edge

NEWS

NEWS OF THE WEEK

- 1862 **MEDICAL RESEARCH:** Inquiry Turns Into OK Corral for U.K. Primate Research
- 1863 **GENOMICS:** Sequence Tells Mouse, Human Genome Secrets
- 1865 **EUROPEAN RESEARCH:** Mystery Academy Holds First Powwow in Private
- 1865 **SCIENCESCOPE**
- ▼1866 **PLANETARY SCIENCE:** A Smashing Source of Early Martian Water?
1977
- ▼1866 **PLANETARY SCIENCE:** ... And an Icy Patch at Mars's South Pole
1841
- 1867 **MEDICAL RESEARCH:** U.K. Researchers Hope for Clarity in Tissue Use
- 1869 **CELL PROLIFERATION:** Common Control for Cancer, Stem Cells

NEWS FOCUS

- 1870 **ASTRONOMY:** Sensing the Hidden Heat of the Universe

- ▼1872 **ARCHAEOLOGY:** Oldest New World Writing Suggests Olmec Innovation
1984
- 1874 **DIGITAL INFORMATION SYSTEMS:** Mapping the Future
- 1877 **MARINE GEOLOGY:** Nations Look for an Edge in Claiming Continental Shelves
- 1879 **CANADA:** New Research Chairs Mean Brain Gain for Universities
- 1881 **RANDOM SAMPLES**



1870

Infrared eye

SCIENCE'S COMPASS

1889 LETTERS

The Ins and Outs of Organic Farming I. M. Goklany. *Response* P. Mäder, A. Fließbach, D. Dubois, L. Gunst, P. Fried, U. Niggli. **Organic Farming and Energy Efficiency** D. Ziehl. *Response* P. Mäder, A. Fließbach, D. Dubois, L. Gunst, P. Fried, U. Niggli. **Would Title IX Help Women in Science?** A. M. De Boer; M. L. Smither-Kopperl. **Atacamite in Jaws and Printed Wiring Boards** S. R. Stock and L. J. Turbini. **A National Ecological Observatory Network** J. A. Teeri and P. H. Raven.

BOOKS ET AL.

- 1894 **HISTORY OF SCIENCE:** *In Darwin's Shadow The Life and Science of Alfred Russel Wallace. A Biographical Study on the Psychology of History* M. Shermer, reviewed by T. Söderqvist
- 1895 **HISTORY OF SCIENCE:** *Lost Discoveries The Ancient Roots of Modern Science—from the Babylonians to the Maya* D. Teresi, reviewed by J. E. McClellan III
- 1896 **Browsings**

HISTORICAL ESSAY

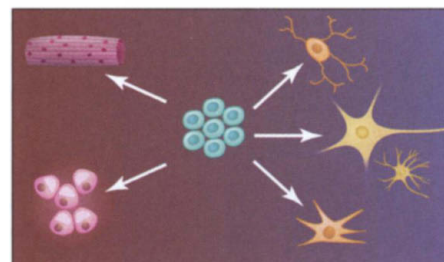
- 1899 **PORTRAITS OF SCIENCE:** Kurt Gödel—Separating Truth from Proof in Mathematics K. Devlin

PERSPECTIVES

- ▼1901 **DEVELOPMENT:** A Tail of Transdifferentiation D. L. Stocum
1993
- ▼1903 **ECOLOGY:** Looking Beneath the Surface
1987 J. A. Morgan
- 1904 **CHEMISTRY:** The Simplest "Enzyme" M. Movassaghi and E. N. Jacobsen
- 1905 **MOLECULAR BIOLOGY:** Untangling Checkpoints N. Sagata

1901

An axolotl's tale of regeneration



1899

Limitations of axiom-based systems

RESEARCH

BREVIA

- 1967 Dissolved Organic Carbon Support of Respiration in the Dark Ocean** J. Arfistegui, C. M. Duarte, S. Agustí, M. Doval, X. A. Álvarez-Salgado, D. A. Hansell

REPORTS

- 1969 Cavitation-Induced Reactions in High-Pressure Carbon Dioxide** M. W. A. Kuijpers, D. van Eck, M. F. Kemmere, J. T. F. Keurentjes
- 1971 Closed-Shell Molecules That Ionize More Readily Than Cesium** F. A. Cotton, N. E. Gruhn, J. Gu, P. Huang, D. L. Lichtenberger, C. A. Murillo, L. O. Van Dorn, C. C. Wilkinson
- 1975 Recent Earth Oblateness Variations: Unraveling Climate and Postglacial Rebound Effects** J. O. Dickey, S. L. Marcus, O. de Viron, I. Fukumori
- ▼ **1977 Environmental Effects of Large Impacts on Mars** T. L. Segura, O. B. Toon, A. Colaprete, K. Zahnle
1866
- 1980 Regulation of Oceanic Silicon and Carbon Preservation by Temperature Control on Bacteria** K. D. Bidle, M. Manganello, F. Azam
- ▼ **1984 Olmec Origins of Mesoamerican Writing** M. E. D. Pohl, K. O. Pope, C. von Nagy
1872
- ▼ **1987 Grassland Responses to Global Environmental Changes Suppressed by Elevated CO₂** M. R. Shaw, E. S. Zavaleta, N. R. Chiariello, E. E. Cleland, H. A. Mooney, C. B. Field
1903
- 1991 A General Model for Designing Networks of Marine Reserves** E. Sala, *et al.*

THE HUMAN KINOME

VIEWPOINT

- 1911 Mitogen-Activated Protein Kinase Pathways Mediated by ERK, JNK, and p38 Protein Kinases** G. L. Johnson and R. Lapadat

REVIEW

- 1912 The Protein Kinase Complement of the Human Genome** G. Manning, D. B. Whyte, R. Martinez, T. Hunter, S. Sudarsanam

See also Science's STKE on p. 1841

POLARITY: FROM CELL TO ORGANISM

- 1941 Humpty Dumpty and All That ...**

REVIEWS

- 1942 Generating and Exploiting Polarity in Bacteria** L. Shapiro, H. H. McAdams, R. Losick
- 1946 Anterior-Posterior Polarity in *C. elegans* and *Drosophila*—PARallels and Differences** J. Pellettieri and G. Seydoux
- 1950 Shaping the Vertebrate Body Plan by Polarized Embryonic Cell Movements** R. Keller
- 1955 Composition and Formation of Intercellular Junctions in Epithelial Cells** E. Knust and O. Bossinger
- 1959 Molecular Mechanisms of Axon Guidance** B. J. Dickson

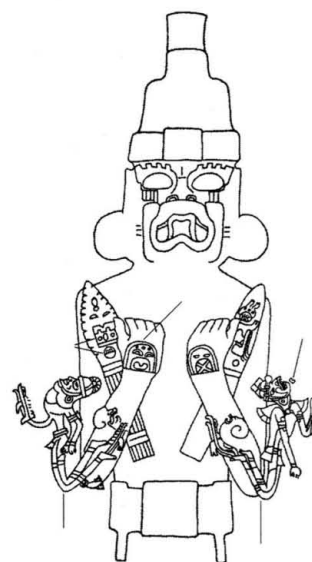
See also Science's STKE on p. 1841



COVER 1941

The *Caenorhabditis elegans* polarity regulator PAR-2, here tagged with green fluorescent protein, is uniformly distributed in a one-cell embryo (top left) but localizes to the posterior before the first cell division (bottom right). Mechanisms involved in the establishment and maintenance of polarity are featured in the special section in this issue. [Photo: Adrian Cuenca]

- ▼ **1993 Ectoderm to Mesoderm Lineage Switching During Axolotl Tail Regeneration** K. Echeverri and E. M. Tanaka
1901
- 1996 Inhibition of Excess Nodal Signaling During Mouse Gastrulation by the Transcriptional Corepressor DRAP1** R. Iratni, Y.-T. Yan, C. Chen, J. Ding, Y. Zhang, S. M. Price, D. Reinberg, M. M. Shen
- 1999 Regulation of Spermatogenesis by Testis-Specific, Cytoplasmic Poly(A) Polymerase TPAP** S. Kashiwabara *et al.*
- 2002 A Role for the Protease Falcipain 1 in Host Cell Invasion by the Human Malaria Parasite** D. C. Greenbaum, A. Baruch, M. Grainger, Z. Bozdech, K. F. Medzhradszky, J. Engel, J. DeRisi, A. A. Holder, M. Bogoy
- 2006 Signaling of Rat Frizzled-2 Through Phosphodiesterase and Cyclic GMP** A. Ahumada, D. C. Slusarski, X. Liu, R. T. Moon, C. C. Malbon, H. Wang
- 2010 Resetting the Circadian Clock by Social Experience in *Drosophila melanogaster*** J. D. Levine, P. Funes, H. B. Dowse, J. C. Hall
- 2013 Functional Neuroimaging of Speech Perception in Infants** G. Dehaene-Lambertz, S. Dehaene, L. Hertz-Pannier

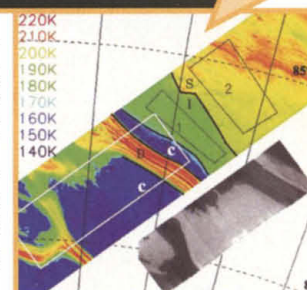


1984

First words in the New World

New on Science Express

Martian icy patch



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CONTENT HIGHLIGHTS AS OF 6 DECEMBER 2002

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SCIENCE EXPRESS

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Exposed Water Ice Discovered Near the South Pole of Mars

▼ T. N. Titus, H. H. Kieffer, P. R. Christensen

1866 Infrared radiation imaging by Mars Odyssey found exposed water ice at the south pole that provides another accessible source of water vapor for transient clouds.

Rates of Behavior and Aging Specified by Mitochondrial Function During Development A. Dillin *et al.*

Defects in the mitochondrial respiratory chain of the nematode *C. elegans* during the developmental phase (but not adulthood) result in worms with an extended life-span.

Bose-Einstein Condensation of Cesium T. Weber, J. Herbig, M. Mark, H.-C. Nägerl, R. Grimm

Bose-Einstein condensation of cesium has been achieved using a combination of optical and magnetic trapping techniques.

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career resources for scientists

GLOBAL: And Now for Something Completely Different ...

Edited by K. Urquhart

Each essayist in December's careers feature has a unique or unusual job—and a passion for what they do.

UK: A Funny Thing Happened on the Way to the Lab H. Pilcher

Working with children, animals, stem cells, and blokes in white coats provides plenty of material for lively stand-up comedy routines.

NETHERLANDS: School for T-Shaped People T. Jetten

An innovative Ph.D. program provides students with in-depth research knowledge and a wide disciplinary focus.

GERMANY: Broaden Your Horizons R. Bartz

A year in Texas was a boon to one participant in a U.S.–Germany exchange program.

SINGAPORE: The Ideal Ph.D. Mentor—A Student's Perspective

W. T. Ling

What one student expects of her supervisor—and why.

GRANTSNET: December News K. Cottingham

Funds for NIH fellows; a postdoctoral fellowship for study in the UK; and the latest biomedical funding news.

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RESEARCH FUNDING DATABASE

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NEWS SYNTHESIS: Europe Wakes Up to Aging G. Weiss

Its nations have long neglected the science of aging—but change is in the air.

NEWS FOCUS: Ageless Activist I. Chen

Paola Timiras has advocated for education about aging since the 1960s.

NOTEWORTHY THIS WEEK: Domino Effect C. Seydel

Multiple DNA-wrapping proteins might execute calorie restriction's life-extending effects in fruit flies.

science's stke

www.stke.org

signal transduction knowledge environment

KINOME

See related articles beginning on p. 1911

CONNECTIONS MAPS: Canonical Mitogen-Activated Protein Kinase Pathways R. Lapadat and G. L. Johnson

ERK1/ERK2 MAPK, JNK MAPK, and p38 MAPK pathways.

PERSPECTIVE: Hitting the Target—Emerging Technologies in the Search for Kinase Substrates B. D. Manning and L. C. Cantley

Help in the search for physiological targets of protein kinases.

PERSPECTIVE: Plasticity of the Kinomes in Monkey and Rat Tissues S. Pelech and H. Zhang

A screen of 78 mammalian kinases yields unexpected variation.

PROTOCOL: Design and Use of a Mammalian Protein-Protein Interaction Trap (MAPPIT) S. Eyckerman *et al.*

Detecting interactions and screening for novel protein partners.

POLARITY

See related articles beginning on p. 1941

PERSPECTIVE: Aurora-A in Cell Fate Control C. Gonzalez

Role for Aurora-A in asymmetric protein localization.

PERSPECTIVE: Creating Asymmetric Cell Divisions by Skewing Endocytosis Q. Shen and S. Temple

How Notch signaling is silenced in only one daughter cell by asymmetric distribution of Numb.

PERSPECTIVE: Pulsating Ion Fluxes and Growth at the Pollen Tube Tip K. R. Robinson and M. A. Messerli

Do oscillating ion fluxes direct polarized growth?

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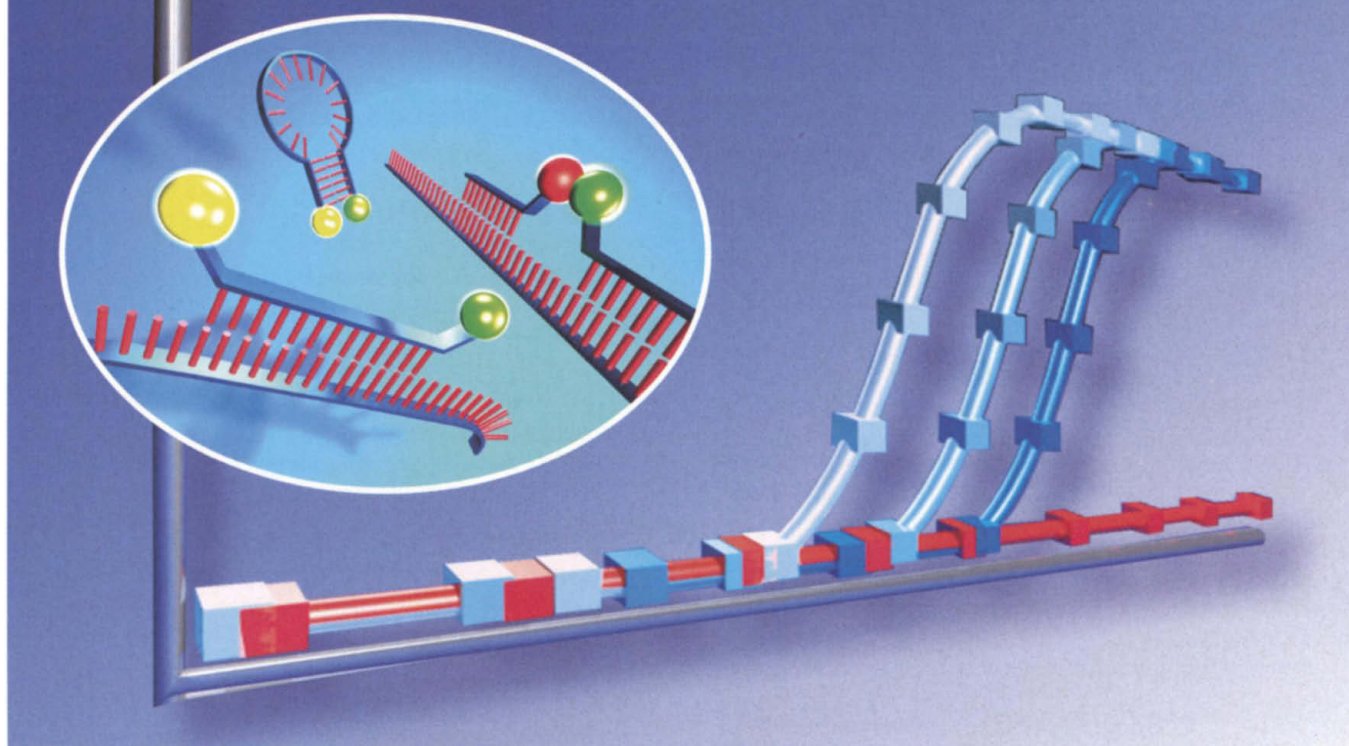
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THIS WEEK IN Science

edited by Phil Szuromi

Sonochemistry in Green Solvents

Sonochemistry takes advantage of the high temperatures that can be generated in solution by the collapse of cavitation bubbles. Kuipers *et al.* (p. 1969) show that radicals formed by cavitation can be used to induce polymerization reactions of methyl methacrylate in high-pressure liquid CO₂. High conversions result because the polymer is poorly solvated under these conditions and precipitates, thus maintaining a low-viscosity solution.

In the Original Olmec

A written language emerged in Mesoamerica sometime in the first millennium B.C. It was used in southeastern Mexico by the Olmec, who were responsible for many of the first monuments in the New World and who developed large cities, and the language formed the basis for later Mayan writing. However, the time, location, and developers of Mesoamerican writing have been widely debated. Pohl *et al.* (p. 1984; see the news story by Stokstad) have uncovered a seal and plaque holding glyphic inscriptions from near La Venta, Mexico, an ancient Olmec center. Radiocarbon dates and associated pottery imply that these artifacts date 650 B.C. These finds imply that this region was the origin of Mesoamerican writing and the calendar system, rather than cultures in southwestern Mexico.

Heavy Bombardment Formed Martian Valleys

The surface of Mars shows evidence for at least 25 large impact events about 3.5 billion years ago and for the formation of many of the extensive river valley networks at that same time. Segura *et al.* (p. 1977; see the news story by Kerr) modeled the effects of impacts on valley network formation and found that one 500-kilometer-diameter bolide or a dozen 200-kilometer-diameter bolides could have created the 50-meter-thick global layer of water needed to form the valley networks. Impact events may have created warm and wet conditions on Mars, but only for a very short time after the heavy bombardment. Thus, habitability for life would have been limited to a brief period of comfortable quiescence.

More Is Less

It is commonly assumed that plants will produce more biomass if the atmospheric CO₂ concentrations continue to rise, but CO₂ is not the only agent that affects plant growth. Shaw *et al.* (p. 1987; see the Perspective by Morgan) describe the responses of a grassland ecosystem to a suite of realistic changes in CO₂, tempera-

1975 How Our Planet Shapes Up

Recently, Cox and Chao (2 August, p. 831) determined that Earth's dynamic oblateness had increased sharply in 1998. Now Dickey *et al.* (p. 1975) show that the increase is related to oceanic mass redistribution caused by the Pacific Decadal Oscillation and the El Niño–Southern Oscillation and an increase in subpolar glacier melting. Once these short-term climatic effects are excluded, the longer-term decrease in oblateness caused by postglacial rebound can be refined to estimate the viscosity of the lower mantle.

And in Brevia ...

An upper bound of 20% for the contribution of the dissolved organic carbon flux to support respiration in the dark ocean has been estimated by Aristegui *et al.* (p. 1967) based on a data set they assembled.

ture, precipitation, and nitrogen deposition, alone and in combination. The response of net primary production (NPP) to elevated CO₂ varies strikingly, depending on the status of other global-change factors. As a single factor, elevated CO₂ increases aboveground growth, but when elevated CO₂ is combined with other global changes, elevated CO₂ tends to decrease NPP.

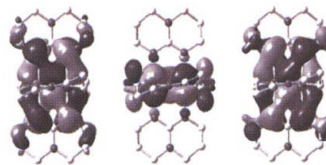
Refuge Networks

Networks of marine reserves have been suggested as a potential way to conserve biodiversity at scales larger than single isolated reserves. However, often there is not

enough information on biodiversity at medium and large scales (beyond data on species richness) to design appropriate networks. To address this problem, Sala *et al.* (p. 1991) have collected information on patterns of biodiversity, ecological processes, and connectivity in reef habitats in the Gulf of California that cover a distance of about 1000 kilometers. Using this information, as well as data on fishing pressure and socioeconomic factors, they developed an algorithm for the selection of reserves to produce a network of reserves that fulfills all conservation goals and permits coexistence with local fisheries.

Easily Lost Electrons

The extremely low ionization energy of cesium has long been exploited in photoelectric cells. Cotton *et al.* (p. 1971) now report on a molecular solid whose ionization energy beats out cesium by a few



tenths of an electron volt. This compound contains a specially ligated tungsten-tungsten quadruple bond, and it is the electron in the weakest δ bond that is emitted at such low energies. Quite unlike cesium, the molecule is relatively stable at ambient conditions and may find use as a reducing agent.

Poly(A) Tails and Spermatogenesis

A gene-regulation mechanism that operates during spermatogenesis has been identified by Kashiwabara *et al.* (p. 1999). Male mice that carry a targeted disruption of the gene for testis-specific, cytoplasmic poly(A) polymerase (TPAP) are infertile because of an arrest in spermiogenesis at the round spermatid stage. The reduced expression of a subset of haploid-specific genes demonstrates that TPAP is essential for spermiogenesis. In addition, TPAP is involved in

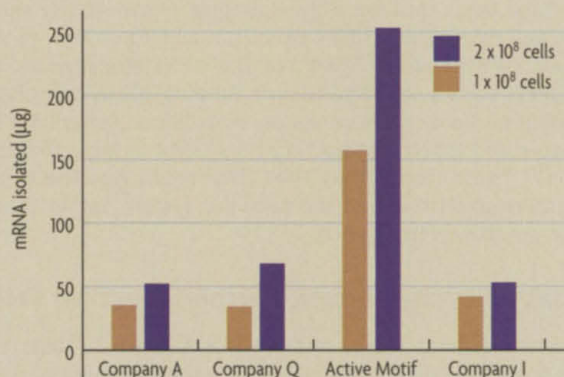


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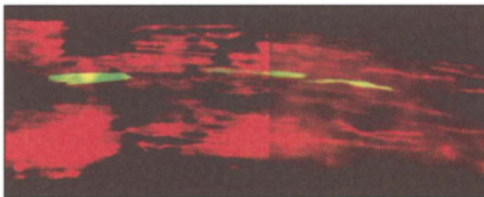


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poly(A) tail extension of particular transcription factor messenger RNAs (mRNAs) in round spermatids, and the elimination of TPAP affects nuclear transport of the TAF10 transcription factor. Hence, TPAP is critical for mRNA control during mammalian gametogenesis through posttranscriptional and posttranslational events.

Stem Cell Tails

Many recent experiments on stem cells have focused on whether their differentiative capabilities allow switching of their developmental lineages. Echeverri and Tanaka (p. 1993; see the Perspective by Stocum) now follow the fate of stem cells in the relatively normal context of a regenerating salamander tail. Using real-time, live observation of individually labeled cells, the authors find that cells labeled in the spinal cord contribute to muscle and cartilage of the regenerating tail, as well as to more neuronal cells. Although amphibians show a greater capacity to regenerate limb and tail than do mammals, these observations of amphibian cell plasticity may lead to insights into the factors that control mammalian cell plasticity.



Blocking Malarial Entry

The widespread resistance of human malaria parasite *Plasmodium falciparum* to the few effective antimalarial drugs has led to a search for new therapeutic targets. Greenbaum *et al.* (p. 2002) used a chemical proteomics screen to identify a cysteine protease, falcipain-1, that is essential for the parasite merozoites to invade human red blood cells. By screening chemical libraries, the authors identified inhibitors that specifically blocked only falcipain-1 activity and parasite invasion of erythrocytes, but not other cysteine proteases or other stages in the parasite life cycle.

Wnt Signaling Through Transducins

Wnts are signaling molecules with critical roles in development that act through a receptor called Frizzled. Although Frizzled proteins resemble heterotrimeric guanine nucleotide binding protein (G protein)-coupled receptors, Wnt signaling is best understood in cases where Wnts control transcription through a β -catenin-dependent signaling mechanism. Ahumada *et al.* (p. 2006) describe a different mechanism for Wnt signaling in which the rat Frizzled-2 protein appears to be coupled to the G protein transducin, better known for its central role in signaling in visual tissues that respond to light. In cultured cells, activation of transducin was required for Wnt-dependent activation of guanosine 3',5'-monophosphate (cGMP)-dependent phosphodiesterase, as well as subsequent decreases in the intracellular concentration of cGMP and increased release of calcium from intracellular stores. Wnt signaling during zebrafish gastrulation could be inhibited when phosphodiesterase activity was reduced after application of pharmacological inhibitors.

Wakeful Smells in the Morning

It is hard to sleep late when your household is up and about at the crack of dawn. Such social influences are seen in other animals, but how are these signals communicated? Levine *et al.* (p. 2010) have examined mixed *Drosophila* populations of wild-type (rhythmic) and short-period (arrhythmic) fly populations and found that arrhythmic flies destabilize the rhythms of wild-type flies. The information seems to be communicated by chemosensory cues—flies that lack the ability to sense chemical signals ignored their neighbors' activity.

Where Baby Talk Ends Up

How do we learn to understand speech? Dehaene-Lambertz *et al.* (p. 2013) have begun to explore the cerebral origins of language acquisition in humans by adapting the technique of functional magnetic resonance imaging for use on 3-month-old infants. They find evidence for activity in brain regions known to subserve language processing in adults, predominantly on the left side, in response to speech.

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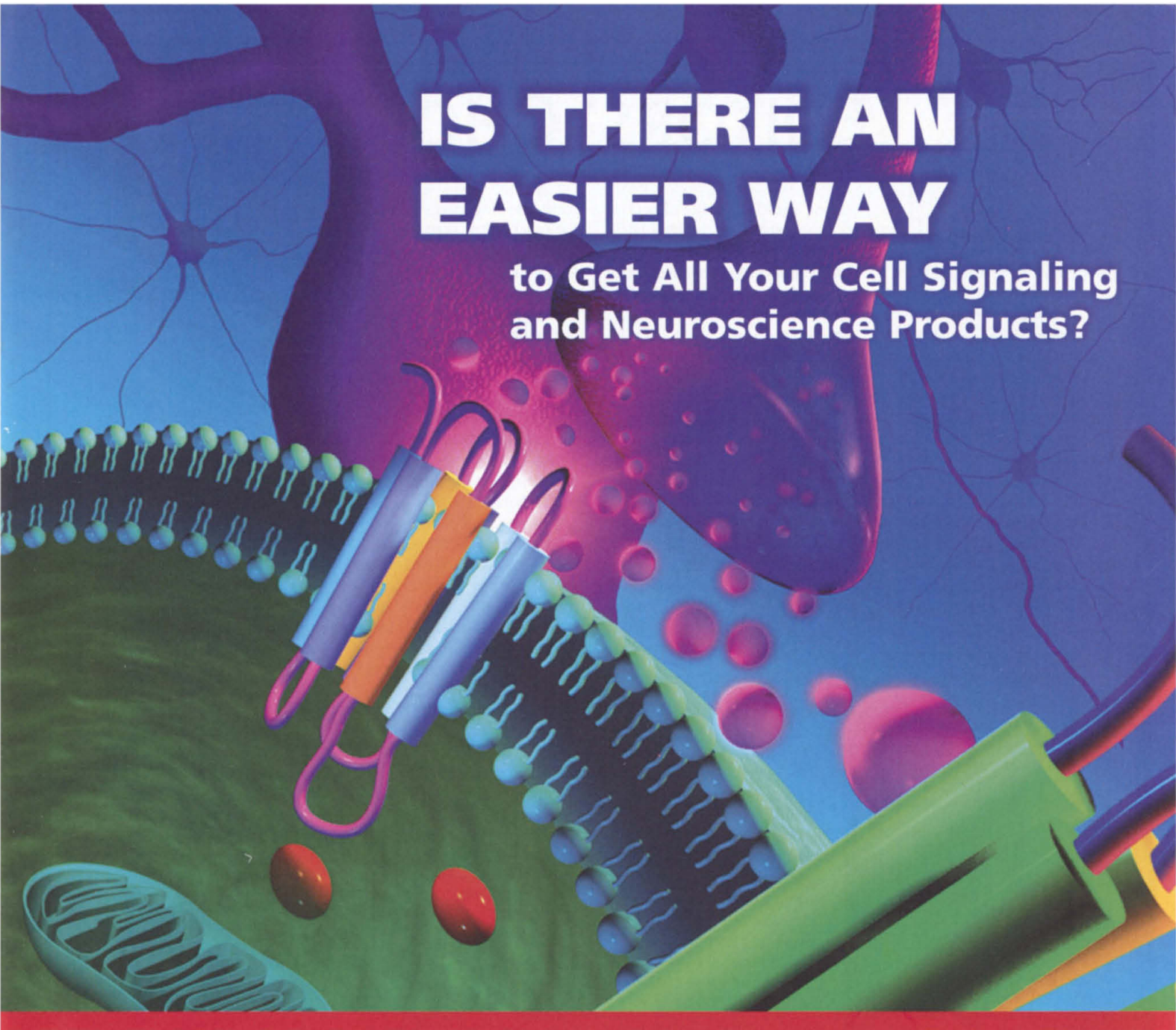
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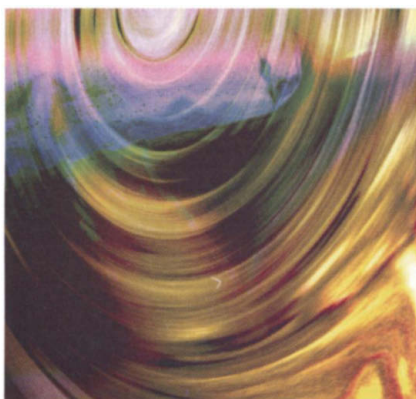
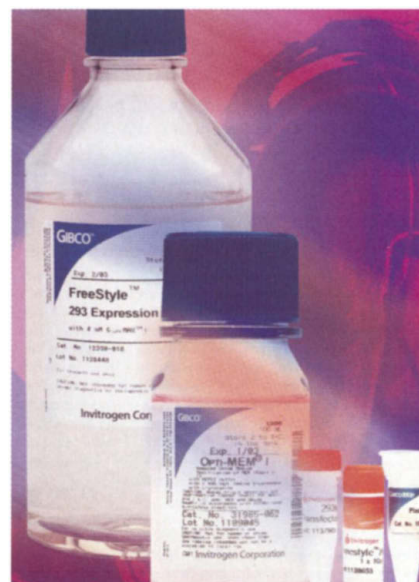
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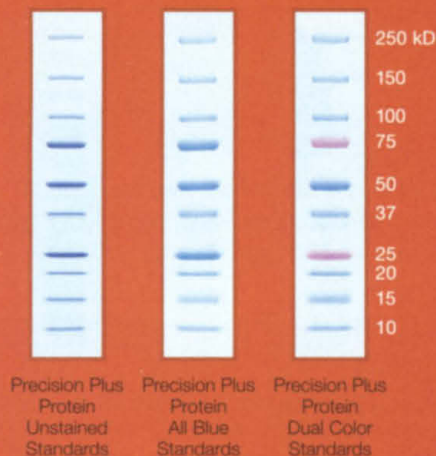
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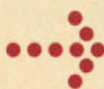
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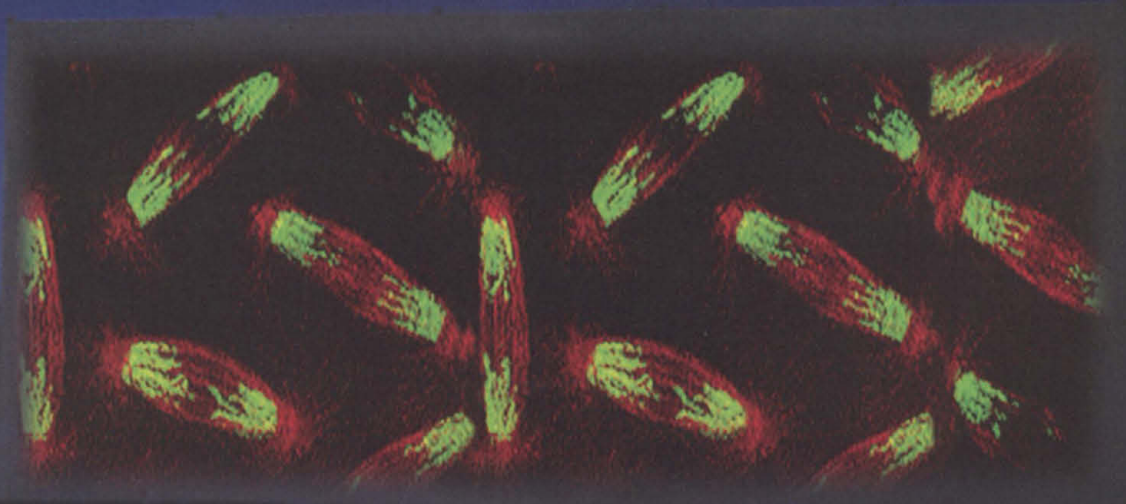
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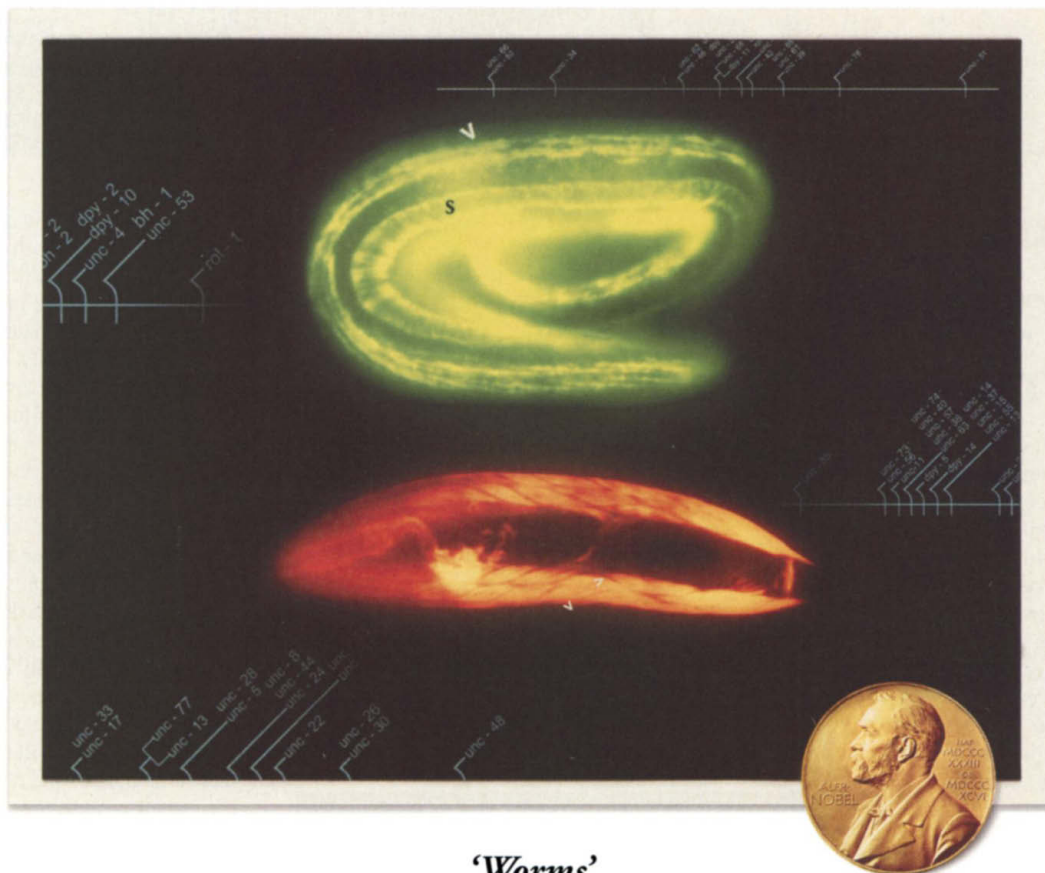
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'Worms'
Sydney Brenner, age 3



'Worms'

2002 Nobel Laureate Sydney Brenner, age 75

It just goes to show that passion grows with age.



Wit and “savage intelligence” characterise Dr Sydney Brenner, whose relationship with the worm started 40 years ago. The insightful Brenner pioneered work on the genetics and development of the humble nematode *Caenorhabditis elegans*. Dr J Sulston and Dr R Horvitz were among the scientists who later joined Dr Brenner’s lab. Now all three share the 2002 Nobel Prize in Physiology or Medicine: for research work on the nematode that has “shed new light on the pathogenesis of many diseases”.

Among his many notable discoveries, Dr Brenner established the existence of messenger RNA and demonstrated how the order of amino acids in proteins is determined. Dr Brenner also led a genomics group at Singapore’s Institute of Molecular and Cell Biology in elucidating the Fugu genome, providing an important key to understanding the complex human genome.

For over two decades, Dr Brenner played an active role in Singapore’s biomedical sciences field. He now sits on the Board of Singapore’s Agency for Science, Technology and Research (A*STAR) and is the Co-Chairman of the Biomedical Sciences International Advisory Council. He plays a key role in A*STAR’s research councils.

Singapore congratulates its Distinguished Friend and Scientific Advisor Dr Sydney Brenner for his Nobel Prize and his relentless pursuit of new knowledge.



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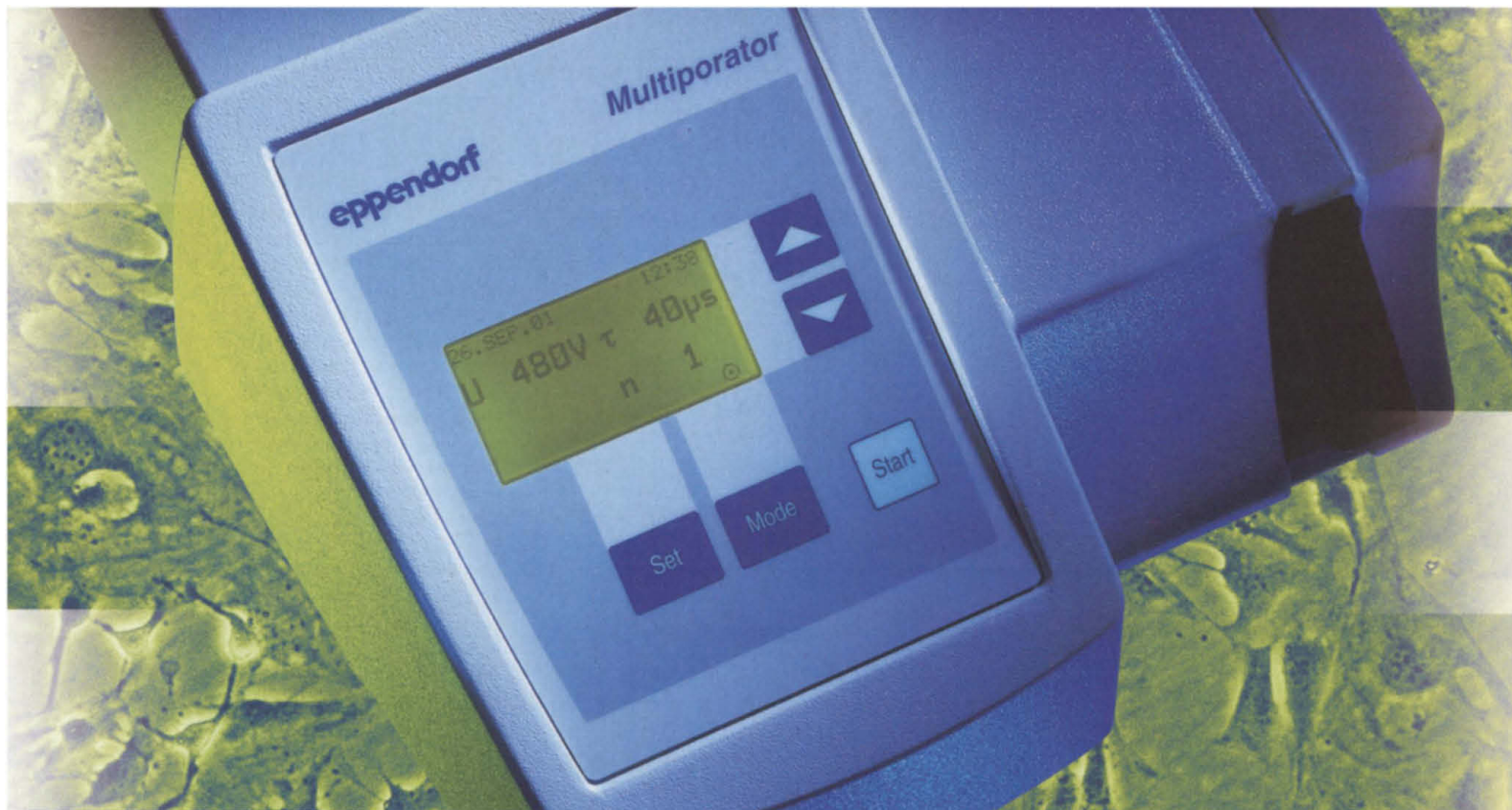
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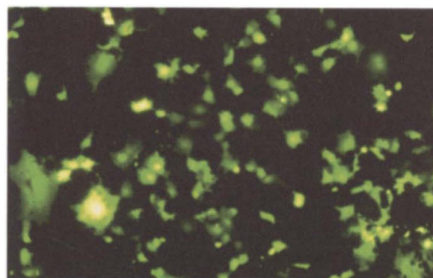
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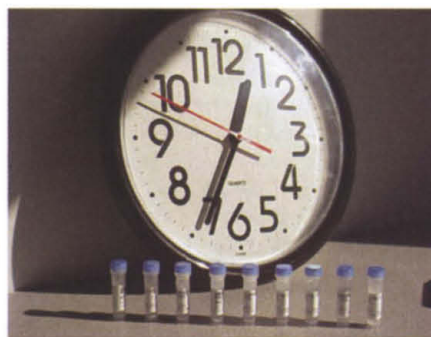
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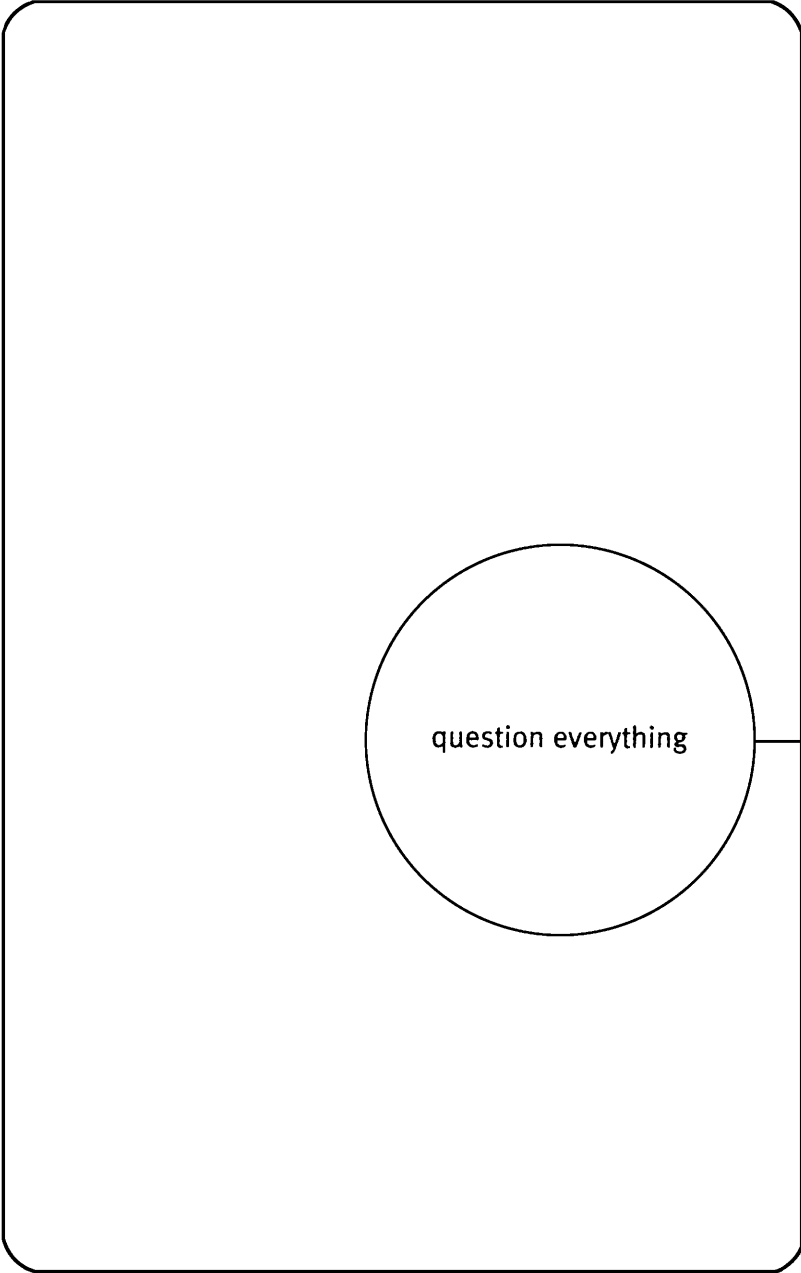
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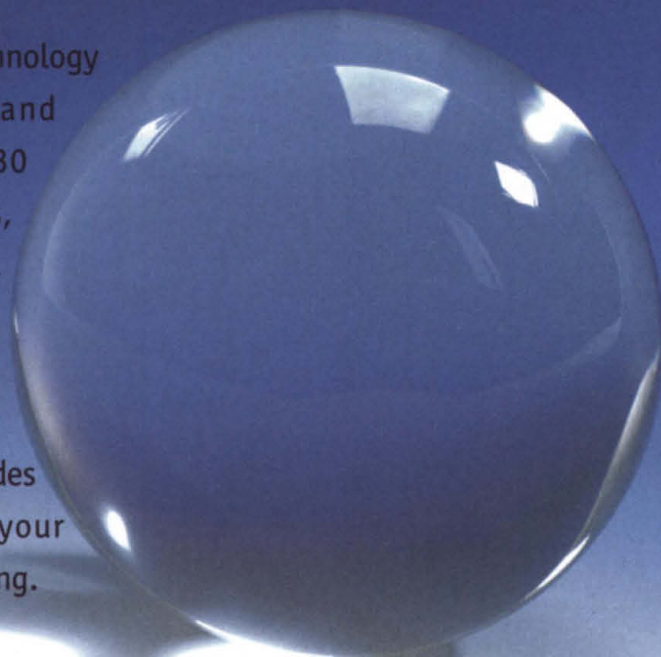
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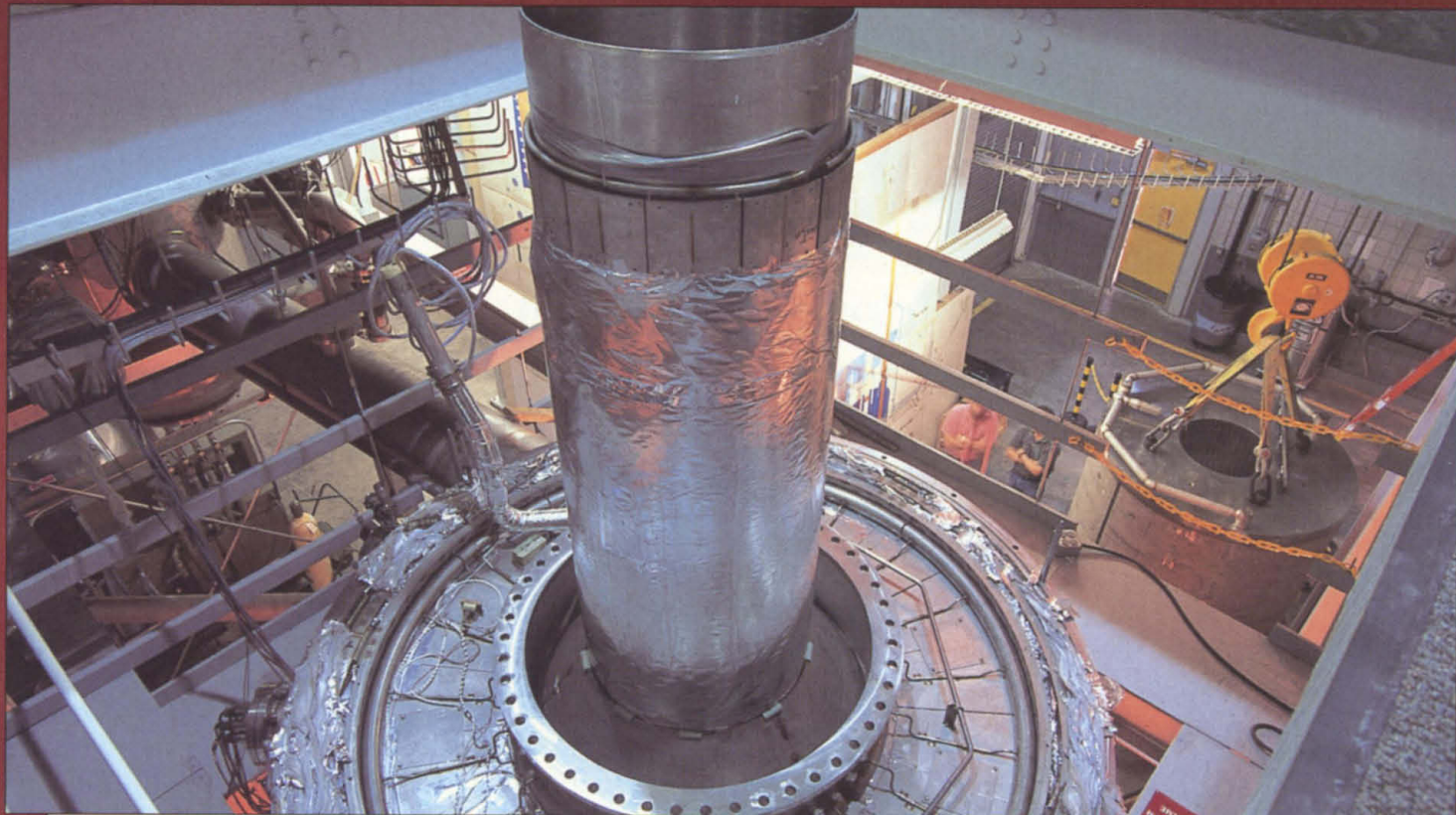
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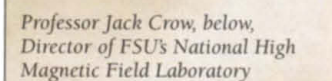
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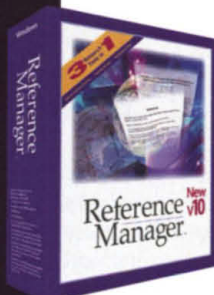
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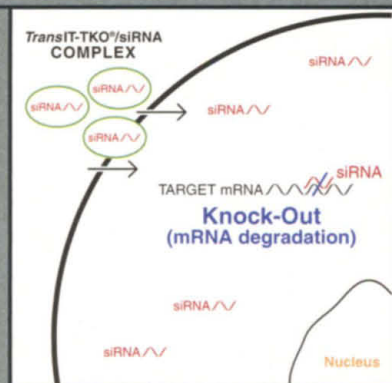
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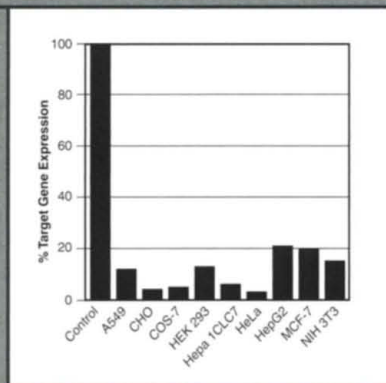
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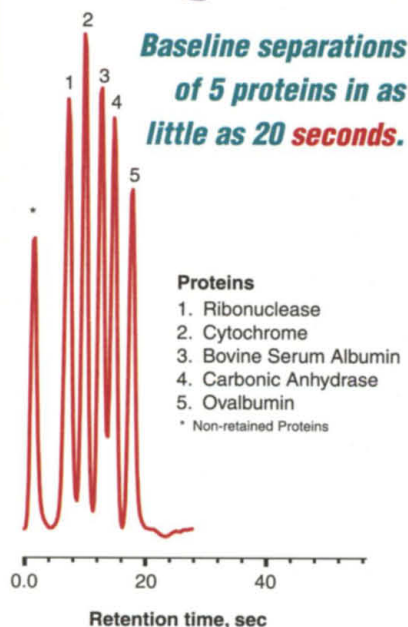
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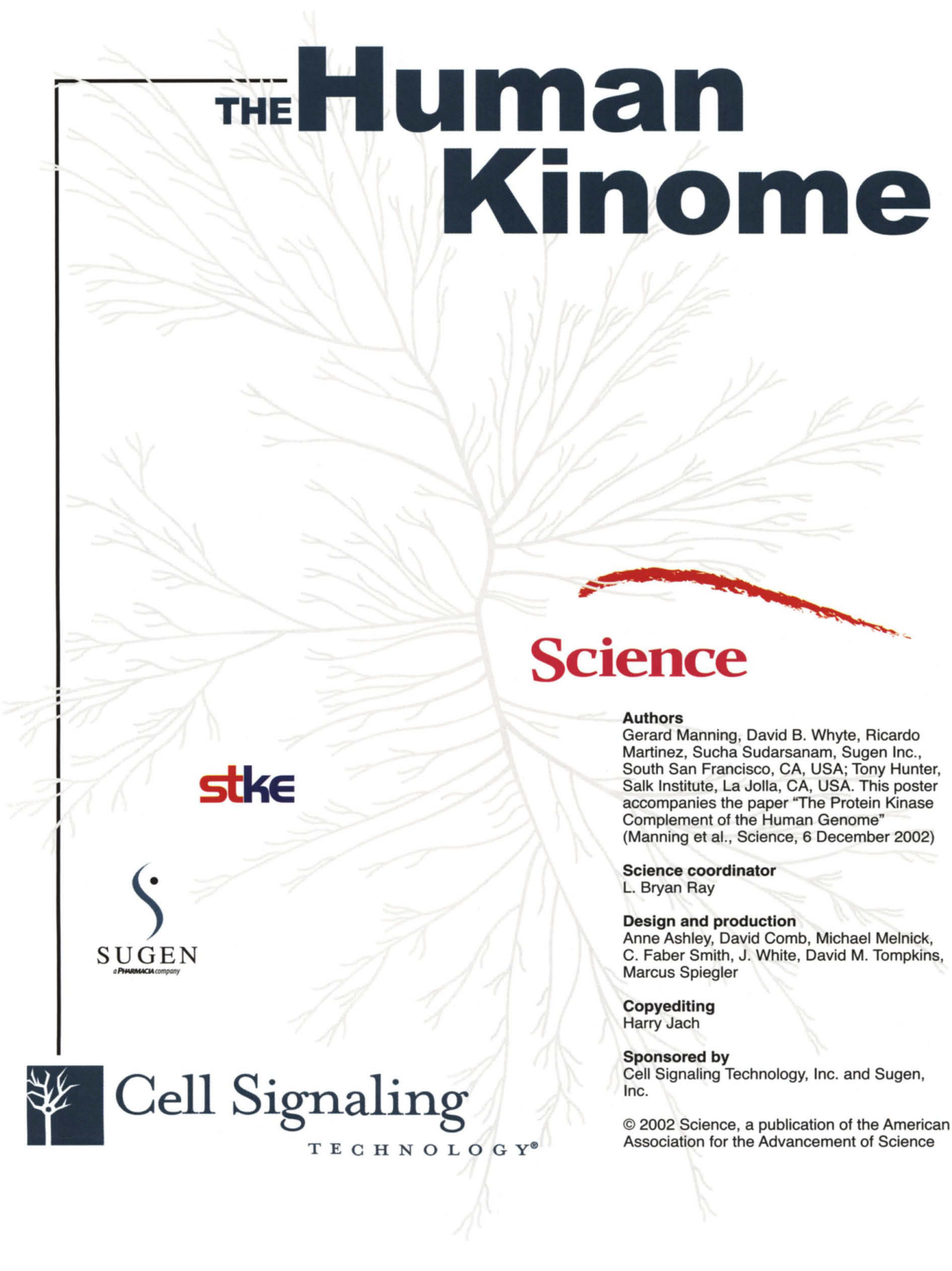
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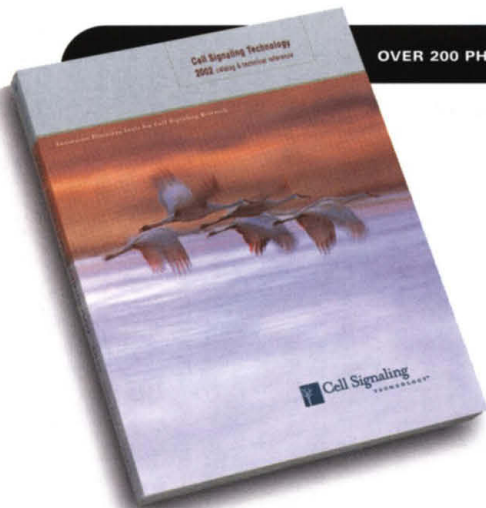
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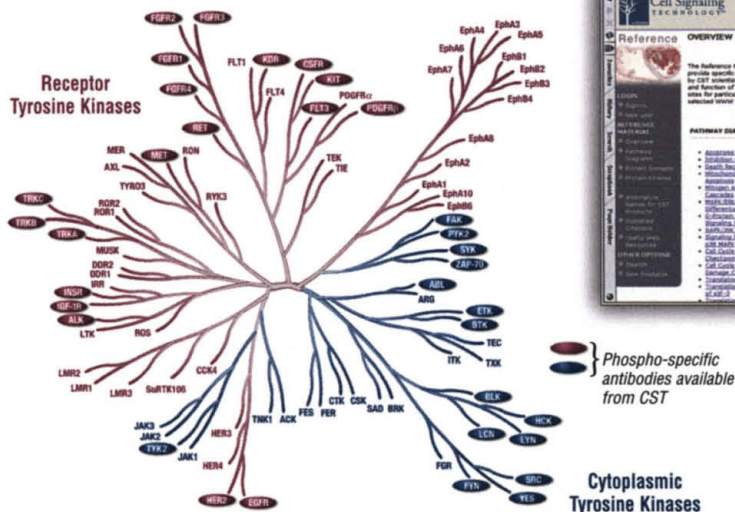
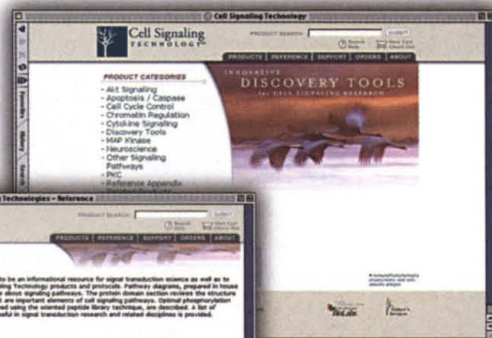
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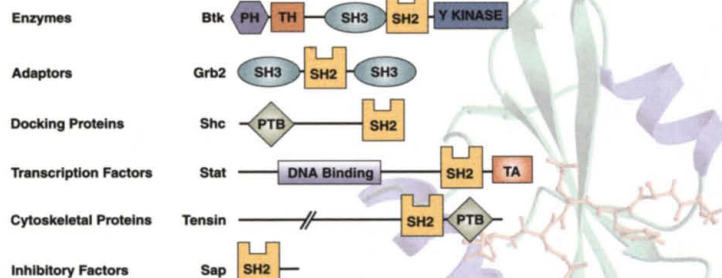
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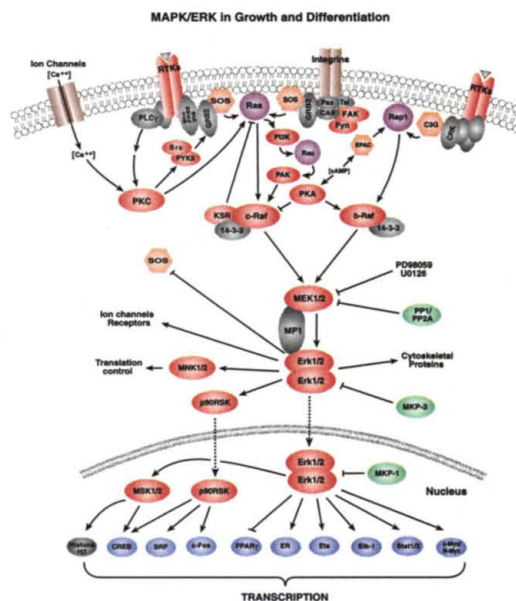
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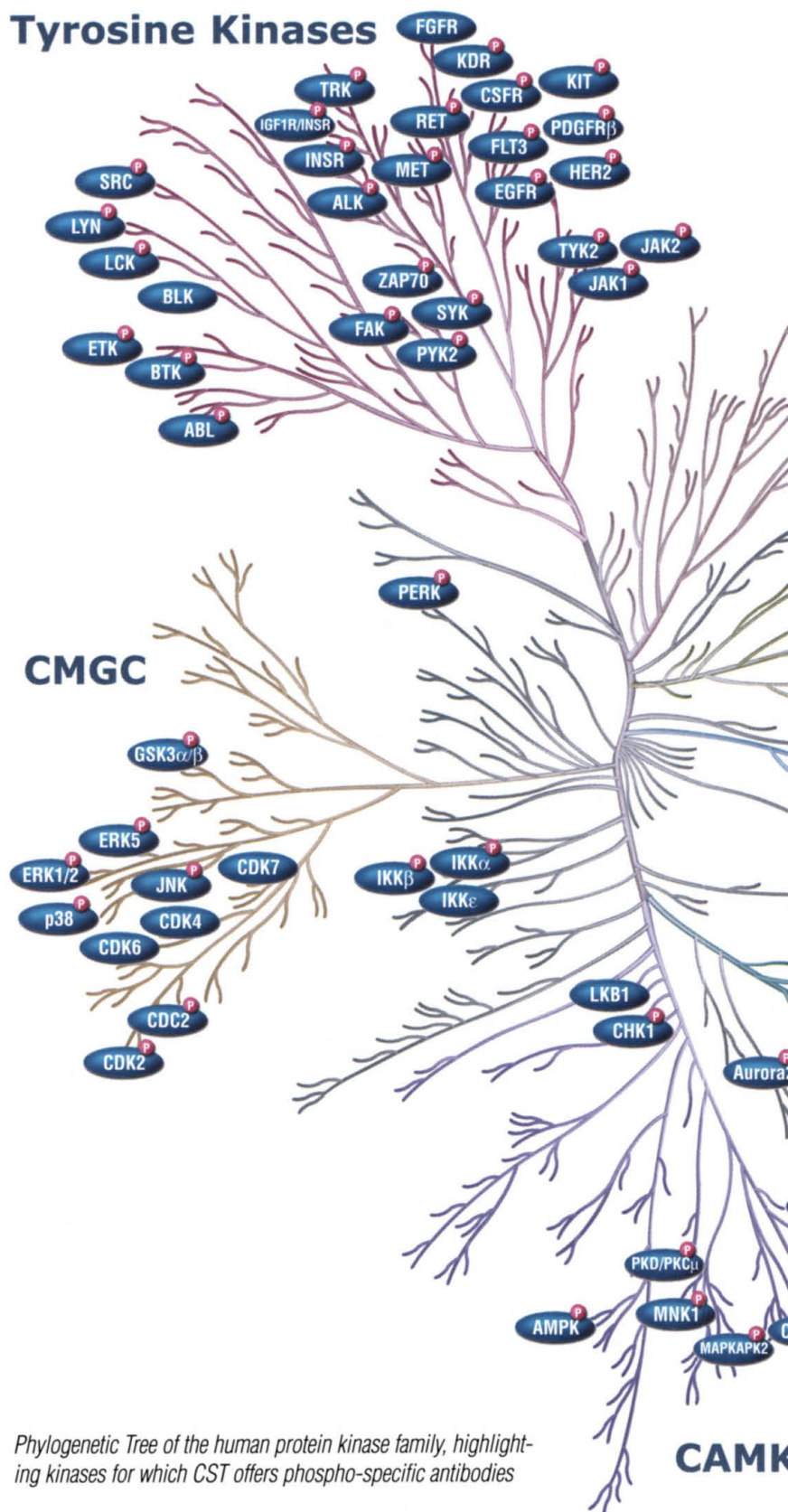
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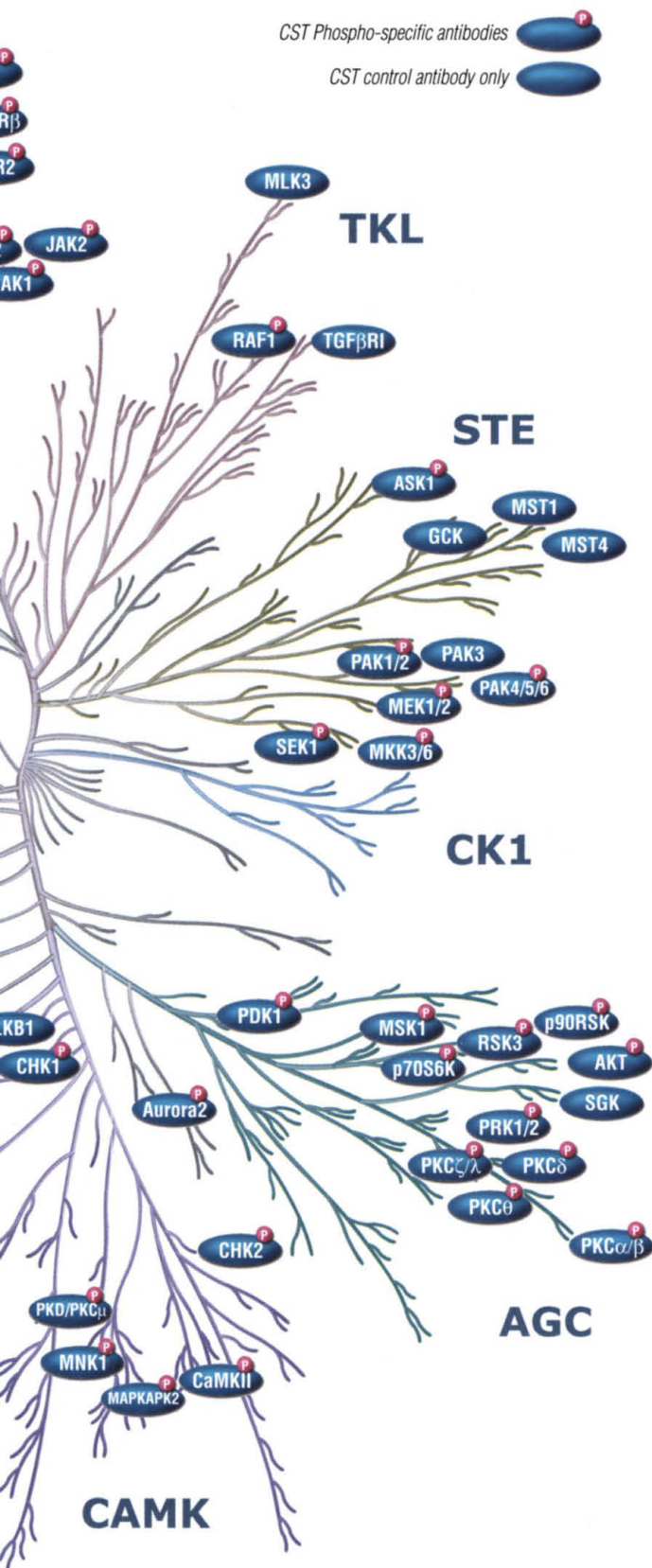
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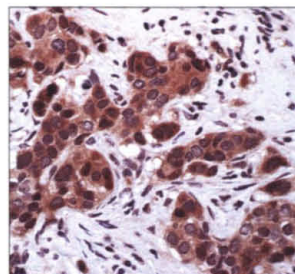


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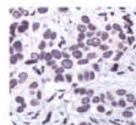
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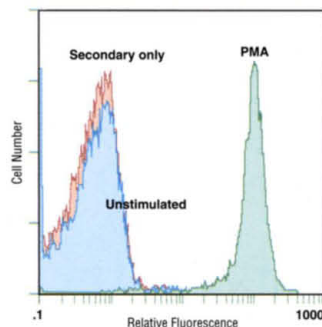


Immunohistochemical staining of paraffin-embedded human breast carcinoma using Phospho-Akt (Ser473) Antibody (IHC Specific) blocked with an antigen-specific phospho-peptide (below) or a non-specific phospho-peptide (left).



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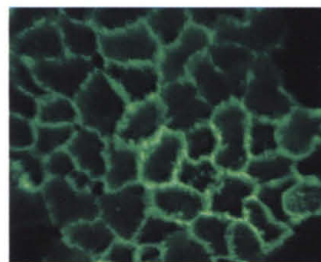
#9101 Phospho-p44/42 MAP Kinase (Thr202/Tyr204) Antibody



Phospho-p44/42 MAP Kinase (Thr202/Tyr204) Antibody was used with FITC-conjugated secondary antibody to label isolated human peripheral blood lymphocytes. Three histograms are overlaid: unstimulated lymphocytes (blue), lymphocytes stimulated with PMA (green) or PMA-stimulated lymphocytes detected with secondary antibody only (orange). (Courtesy of Dr. David Hedley, Princess Margaret Hospital, Toronto.)

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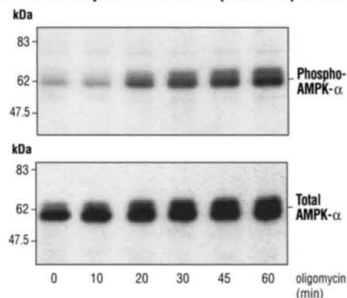


Immunofluorescent staining of EGF treated (left) or untreated (below) A431 cells using Phospho-HER2/ErbB2 (Tyr1112) Antibody, showing membrane localization.



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#2531 Phospho-AMPK-α (Thr172) Antibody and #2532 AMPK-α Antibody



Western blot analysis of HEK293 cell lysates treated with oligomycin for different time periods, using Phospho-AMPK-α (Thr172) Antibody (upper) or control AMPK-α Antibody (lower).

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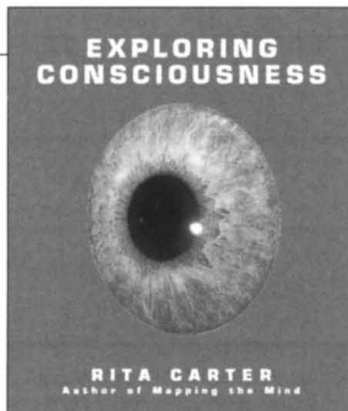
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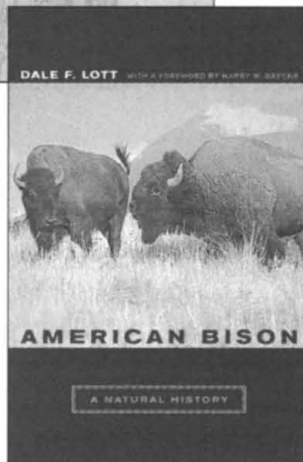
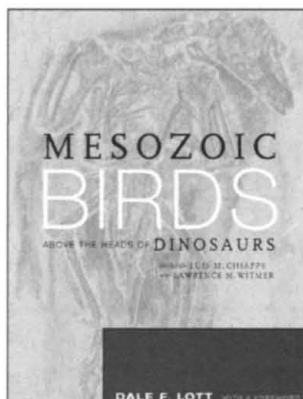
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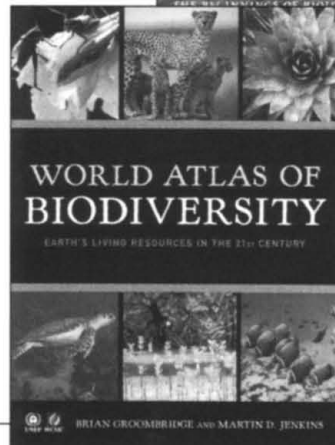
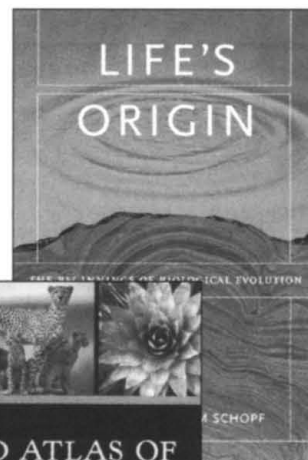
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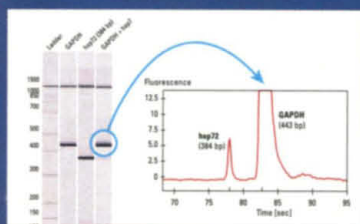
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OVERVIEW ▶ The Women's International Science Collaboration (WISC) Program is funded by the National Science Foundation (NSF) and administered by the International Office of the American Association for the Advancement of Science (AAAS). Because the application rate of women scientists and engineers to work with colleagues overseas to the NSF Office of International Programs has been disproportionately low, the goal of this Program is to increase the participation of women as PIs and co-PIs in international research projects. This program provides grants to individual US scientists who plan to establish new research partnerships with their colleagues in Central/Eastern & Western Europe, the Newly Independent States of the former Soviet Union, Near East, Middle East, Africa, the Americas, Pacific, and Asia. Each grant, up to \$4,000 or \$5,000, will provide travel and living support for a US scientist. When appropriate, an additional grant may be made to an American co-PI. The grants do not cover salary or institutional expenses (e.g. overhead). US scientists can spend up to four

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NEXT DEADLINE ▶ January 15, 2003 (notification by April 15)

INFORMATION ▶ For questions, please contact the appropriate AAAS administrator: Curtis Cook (Central/Eastern & Western Europe, Newly Independent States of the former Soviet Union), ccook@aaas.org; Suteera Nagavajara (East Asia, Pacific), snagavaj@aaas.org; John Schoneboom (Africa, Middle East, Near East, South Asia), jschoneb@aaas.org; Marina Ratchford (Americas, Caribbean), mratchfo@aaas.org. For complete guidelines and application forms please visit our website at: www.aaas.org/international/wiscnew.shtml

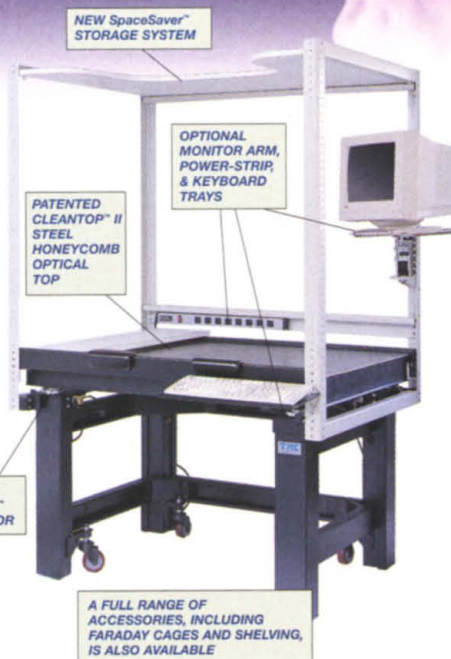


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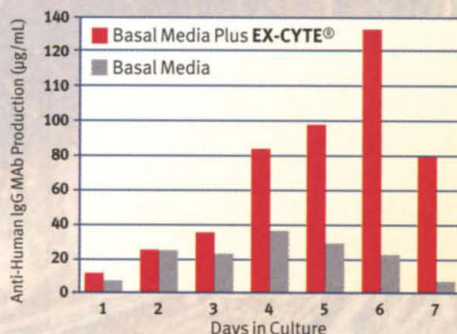
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