

## 정 동 수 박사

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### ◆ 연구관심분야

- 바이러스 및 바이로이드 복제메커니즘
- 생물정보학
- 생물산업 동향분석

### ◆ 학 력

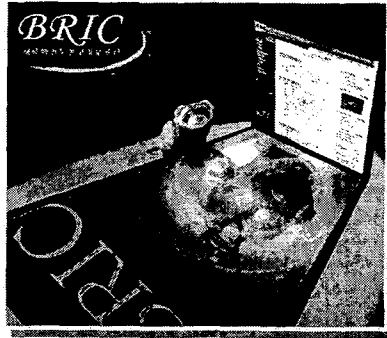
- 경북대학교 미생물학과 학사, 1990. 8.
- 경북대학교 미생물학과 석사, 1993. 2.
- 경북대학교 미생물학과 박사, 1998. 2.
- 영남대학교 미생물학과 박사후 연수 1998.3

### ◆ 주요경력

1. 영남대기초과학연구소 연구원
2. 일본이화학연구소 연구원
3. 생물학전문연구정보센터 팀장

### ◆ 연구 실적 요약

- 학술잡지 논문발표 : 10 편
- 학술컨퍼런스 논문발표 : 8 편
- 국제학회 기조연설, 초청강연, 초청세미나 : 3 회
- 저서/edited books : 1 권
- 연구과제 프로젝트 : 4 건



Dong Soo Jung

**Biological Research Information Center (BRIC)**

- What ?
- New Paradigm
- Research Field
- Application
- Bioinformatics in 2010



**1. Introduction**

2. Microbial Genome

3. Korean Bioinformatics

4. Bioinformatics Market

5. Education

6. BRIC's Bioinformatics

**Bio** + **informat** + **ics**  
 Biology Information Science

It is called as ;

**Bioinformatics, Biocomputing, Biomedical computing,  
 Computational biology, Information biology, Biological data mining,  
 in silico biology**

- apply Mathematics/Statistics/Computer Science to Biology
- storage, analysis, interpretation of biological information
- construction of biological information infrastructure
- solving problems arising from biology using methodology from computer science.

*knowledge-based discovery*

**Sequence Analysis**  
 Pairwise sequence alignment  
 Database searches  
 Multiple sequence alignment  
 Functional prediction

**Structure Analysis**  
 RNA secondary structure prediction  
 RNA tertiary structure modeling  
 Prediction of protein structural features

**Genome Analysis**  
 Comparative genomics  
 Gene expression analysis  
 Pathway databases  
 Path computations

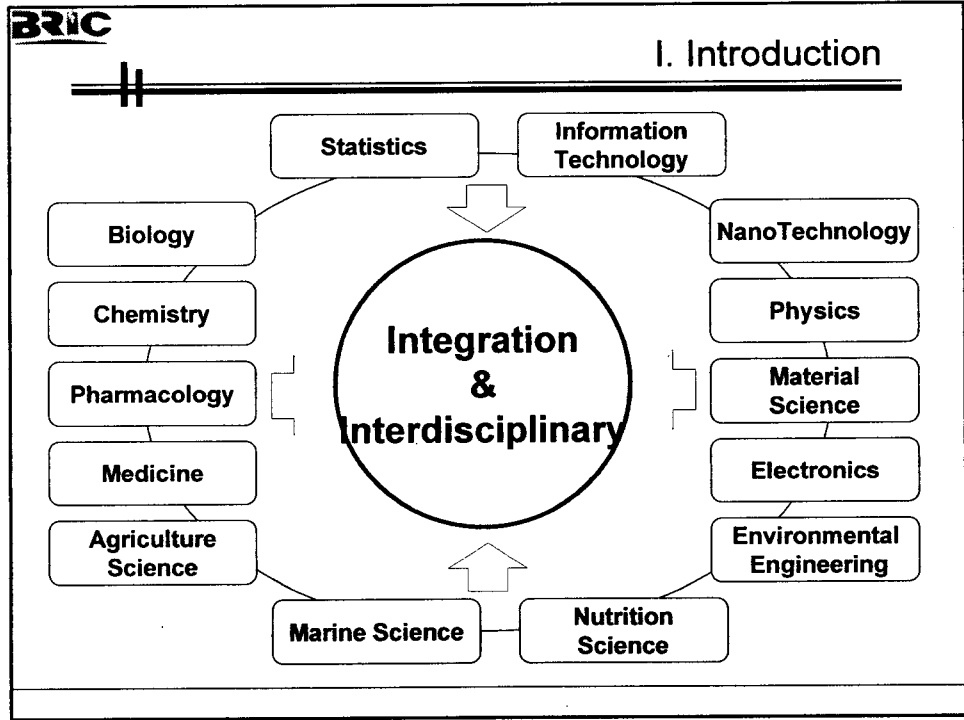
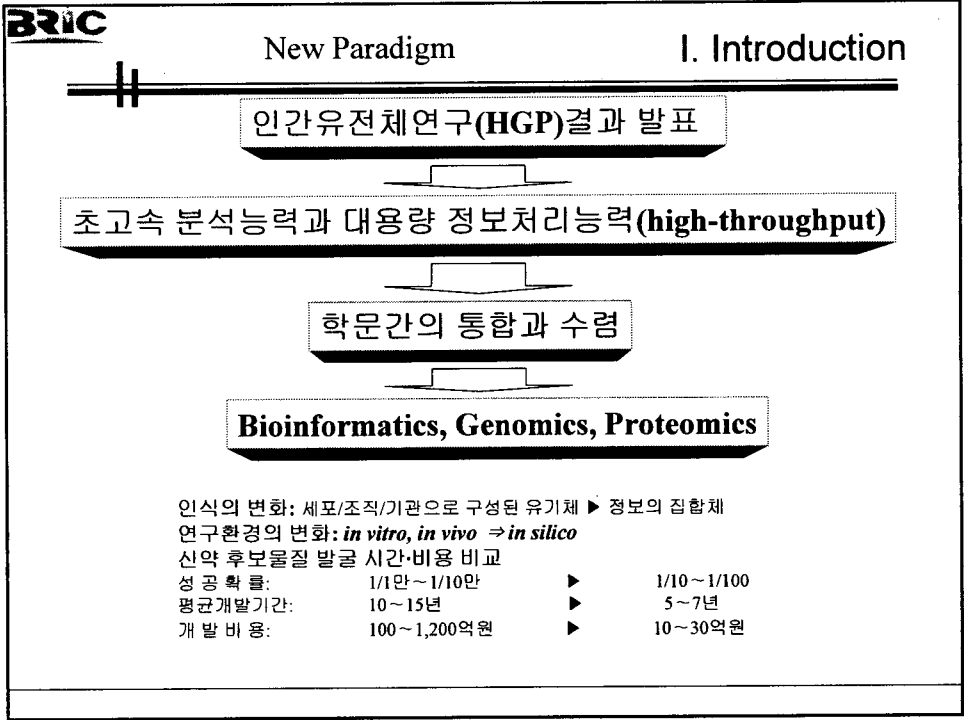
**Relevant computational techniques**  
 Data processing for DNA chip or proteomic analysis  
 Natural language processing for biology  
 Simulation of artificial cell life

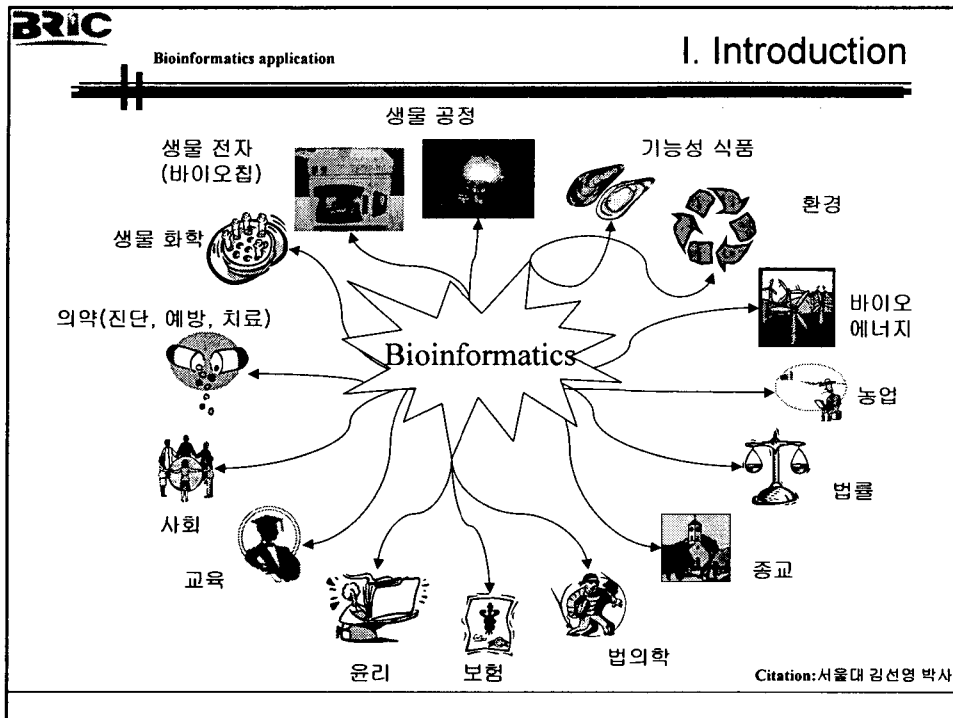
**Medical Informatics**  
 eg., Data bases  
 Telemedicine

**Biological engineering**  
 eg., Microbial processing – simulation and process control

**Bio-Chemical Informatics**  
 Metabolomics  
 Chemical-biological relationships

**Knowledge Formulation**  
 Data base formulation for biological informations  
 Hypothesis formation  
 In silico experimentation  
 New knowledge discovery: Datamining





- BRIC** Bioinformatics in 2010 I. Introduction
- 
- *In Biology*
    - *In silico* experiment
  - *In Healthcare*
    - Fast drug discovery, and personalized drug.
    - Understand diseases at the molecular level.
    - Fast diagnosis
  - *In Universities*
    - Maybe at least one bioinformatician will be needed for biological research.
  - *In Industry*
    - Biotech industries will not survive without introduction of bioinformatics tools and knowledge.
    - Many jobs will be created.

Trends in Microbial Genomics and Bioinformatics



1. Introduction

2. Microbial Genome

3. Korean Bioinformatics

4. Bioinformatics Market

5. Education

6. BRIC's Bioinformatics

Overview  
 Why Microbe?  
 Genome analysis  
 Research Institute  
 Website  
*In Silico*



Microbial Genome Project--worldwide



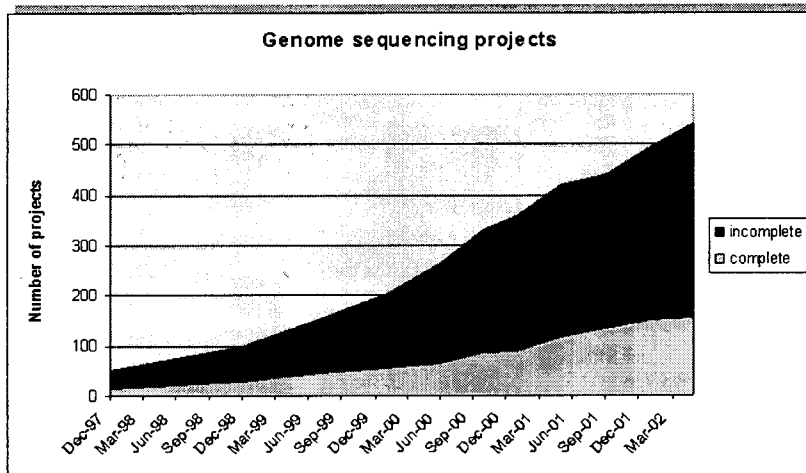
**GOLD<sup>™</sup>: Genomes OnLine Database HomePage**

|  |                                     |   |
|--|-------------------------------------|---|
| Contact:<br>GOLD   | Last Update:<br>June 5, 2002        | Sponsored by<br>Integrated Genomics Inc.  |
| Search GOLD: 570 genome projects   |                                     |   |
| Published Complete Genomes:<br>93<br><small>and 33 chromosome maps</small> | Prokaryotic Ongoing Genomes:<br>283 | Eukaryotic Ongoing Genomes:<br>194<br><small>including 32 chromosome maps</small> |

GOLD: Genomes Online Database, is a World Wide Web resource for comprehensive access to information regarding complete and ongoing genome projects around the world.

Sponsored by Integrated Genomics Inc.

## Microbial Genome Project--worldwide



자료: GOLD: Genomes Online Database

## Why Microbes?

1. Microbes make up about 60% of the Earth's biomass
2. Microbes play a critical role in natural biogeochemical cycles
3. Surviving and thriving in an amazing diversity of habitats, in extremes
4. Offer us new solutions to longstanding challenges in environmental and
5. Waste cleanup, energy production and use, medicine, industrial processes, agriculture, and other areas.
6. Biological underpinnings of climate change and the contributions of microbes to Earth's biosphere
7. Traditional commercial uses for microbes in the brewing, baking, dairy, and other industries.

## Genome Size

| Organism                                | Haploid genome size (Mb) | Predicted number of genes |
|---|--------------------------|---------------------------|
| <i>Arabidopsis thaliana</i> (plant)     | 130                      | ~25,000                   |
| <i>Caenorhabditis elegans</i> (worm)    | 100                      | 18,424                    |
| <i>Drosophila melanogaster</i>          | 180                      | 13,601                    |
| <i>Escherichia coli</i>                 | 4.7                      | 4,288                     |
| <i>Homo sapiens</i> (human)             | 30,000                   | 45,000 – 60,000           |
| <i>Saccharomyces cerevisiae</i> (yeast) | 13.5                     | 6,241                     |

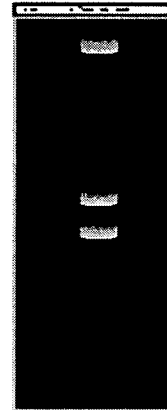
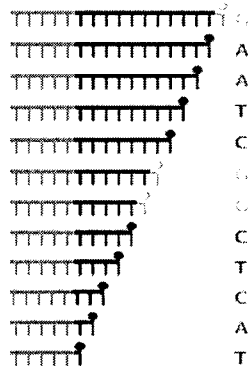
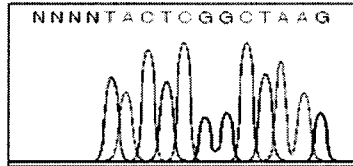
## Steps of Genome Analysis

- 1) Genome sequence assembled
- 2) Identify repetitive sequences – mask out
- 3) Gene prediction – train a model for each genome
- 4) Look for EST and cDNA sequences
- 5) Genome annotation
- 6) Microarray analysis
- 7) Metabolic pathways and regulation
- 8) Protein 2D gel electrophoresis
- 9) Functional genomics
- 10) Gene location/gene map
- 11) Self-comparison of proteome
- 12) Comparative genomics
- 13) Identify clusters of functionally related genes
- 14) Evolutionary modeling

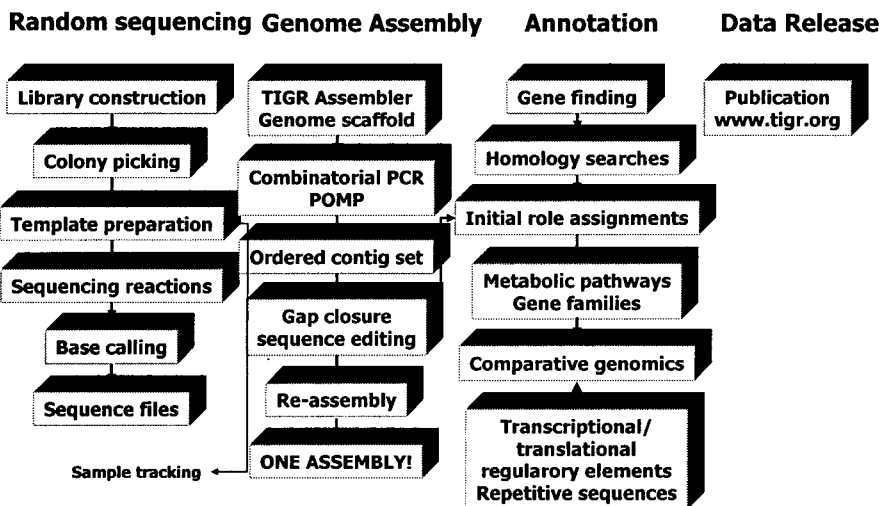


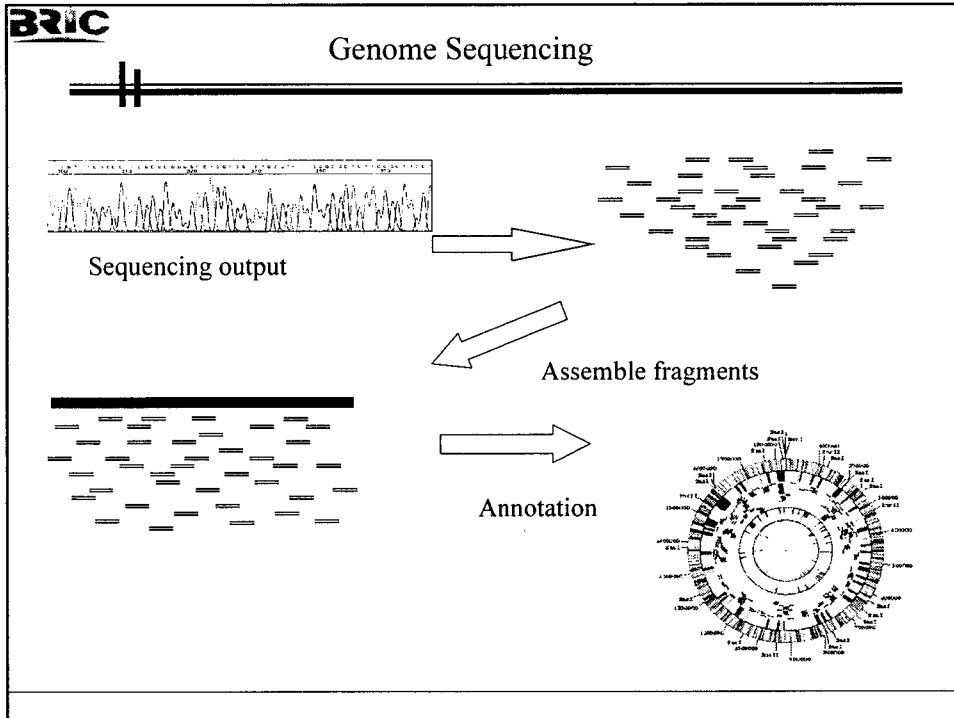
### Cycle Sequencing

The simulated gel image is read from bottom to top, starting with the smallest fragment.



### A Microbial Genome Sequencing Project at TIGR





- BRIC**
- ## Bioinformatic Analysis of Complete Genomes
- Identification/prediction of genes
  - Characterization of gene features
  - Characterization of genome features
  - Prediction of gene function
  - Prediction of pathways
  - Integration with known biological data
  - Comparative genomics

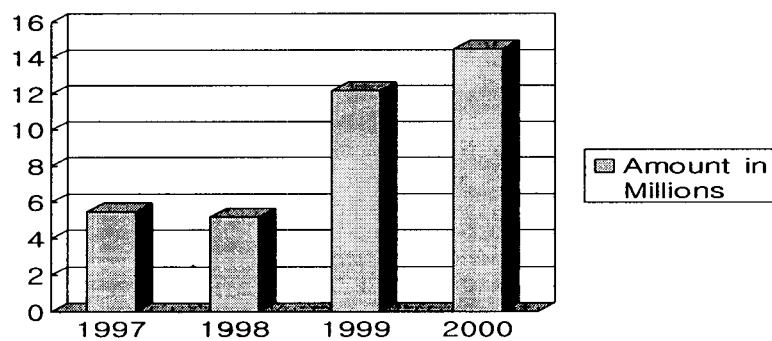
## What are we learning from genome analysis?

The complete genetic blueprint for an organism:

- ▶ Identifies novel biochemical and metabolic pathways;
- ▶ Identifies genes and pathways for bioremediation;
- ▶ Identifies organisms that have the potential to provide renewable energy sources;
- ▶ Identifies genes involved in adaptation to extreme environments;
- ▶ Identifies enzymes of industrial importance;
- ▶ Identifies disease-causing genes;
- ▶ Identifies new targets for development of antibiotics and vaccines.

## DOE Microbial Genome Program Funding

National Institute of Allergy and Infectious Diseases (NIAID)  
Department of Energy (DOE),  
National Science Foundation (NSF).



Totals include 1\$2.4M and 2\$5.8M from DOE Carbon Management Science Program (CMSP).  
CMSP is part of the federal Climate Change Technology Initiative

**DOE-Funded Microbial Genomes**

Sequencing Microbial Genomes to Uncover Potential Applications Relevant to DOE Missions

**Carbon Sequestration**

- *Chlorobium tepidum*
- *Rhodospseudomonas palustris*

**Bioremediation**

- *Deinococcus radiodurans*
- *Pseudomonas putida*

**Cellulose Degradation**

- *Clostridium thermocellum*
- *Thermobifida fusca*

**Energy Production**

- *Methanobacterium thermoautotrophicum*
- *Methylococcus capsulatus*

**Industrial Processes**

- *Halobacterium halobium*
- *Pyrococcus furiosus*

**Technology Development, Pilot Projects**

- *Mycoplasma genitalium*
- *Xylella fastidiosa*

**National Institute of Allergy and Infectious Diseases (NIAID)**

Division of Intramural Research(four)  
 Division of AIDS;  
 Division of Allergy, Immunology and Transplantation;  
 Division of Microbiology and Infectious Diseases  
 Division of Extramural Activities  
 Vaccine Research Center

**DMID Supported Large-Scale Genome Sequencing Projects**▶ **COMPLETED BACTERIAL GENOME PROJECTS**

: *Chlamydia pneumoniae* 외 16종

▶ **ONGOING SEQUENCING PROJECTS**

: *Anopheles gambiae* 46종

NIAID will give priority consideration to large scale sequencing projects for the following organisms: *Aedes aegypti*, *Anopheles gambiae*, *Brugia malayi*, *Coccidioides immitis*, *Group B streptococcus*, *Histoplasma capsulatum*, *Rickettsia rickettsii*, *Toxoplasma gondii* and *Trichomonas vaginalis*.



### Goals

The Microbe Project has three broad goals:

- To build needed infrastructure,
- To promote research,
- To develop human resources and an informed public.

Genome-enabled microbial research holds enormous promise for understanding life at its most basic level, and for enabling breakthrough applications in health, agriculture, biotechnology, the environment and national defense.

The three major components of infrastructure needed to support microbial genomics research are

- 1) Genome sequences
- 2) Tools, technologies and biological resources
- 3) Databases and bioinformatics

New Branch of Division of Cell Biology and Biophysics, The National Institute of General Medical Sciences (NIGMS)

**Purpose: support studies that take a genomics or computational approach to determining protein structures and functions**

#### Berkeley Structural Genomics Center

Focus on two bacteria with extremely small genomes to study proteins essential for independent life: *Mycoplasma genitalium* and *Mycoplasma pneumoniae*.  
Principal Investigator: Sung-Hou Kim, Lawrence Berkeley National Laboratory

#### Center for Eukaryotic Structural Genomics

This Wisconsin-based center seeks to develop high-throughput methods for protein production, characterization and structure determination from *Arabidopsis thaliana*, a plant

#### The Joint Center for Structural Genomics

Focus on novel structures from the roundworm *Caenorhabditis elegans* and on human proteins thought to be involved in cell signaling

#### The Midwest Center for Structural Genomics

unknown folds and on proteins from disease-causing organisms.

#### New York Structural Genomics Research Consortium

develop techniques to streamline every step of structural genomics.

#### Northeast Structural Genomics Consortium

target proteins from various model organisms--including the fruit fly, yeast, and the roundworm--and related human proteins.

#### The Southeast Collaboratory for Structural Genomics

two representative organisms--the roundworm *Caenorhabditis elegans* and its more primitive microbial ancestor, *Pyrococcus furiosus*.

#### Structural Genomics of Pathogenic Protozoa Consortium

This group aims to develop new ways to solve protein structures from organisms known as protozoans, many species of which cause deadly diseases

#### TB Structural Genomics Consortium

seeks to optimize the technical and managerial underpinnings of high-throughput structure determination and will develop a database of structures and functions

## Microbial Genome Projects

### Microbial Genome Projects

Bacillus subtilis Genome Database (BSORF) Bioinformatics Center, Kyoto University and Nara Institute of Science and Technology  
 Chlamydomonas Resource Center (Duke University, USA)  
 Database of Genomes Analysed in NITE (DOGAN)  
 Dictyostelium cDNA Database Dictyostelium discoideum cDNA Project (Dicty\_cDB)  
 Dictyostelium Genome Sequencing Project  
 DOE Joint Genome Institute (JGI) Microbial Genomics, US  
 E-coli genome project (K-12 and -157) (University of Wisconsin-Madison, US)  
 GenoBase: Escherichia coli Genome Database Nara Institute of Science and Technology  
 Genome Analysis Project Japan on E. coli (GenoBase)  
 Genome Database for Cyanobacteria (CyanoBase) Kazusa DNA Research Institute  
 Genome Information Broker (GIB) DNA Data Bank of Japan (84 microbes as of May 2002)  
 Genome to Proteins and Functions  
 GOLD: Genomes OnLine Database HomePage by Integrated Genomics Inc., US  
 MagnaportheDB  
 Malaria Full-Length cDNA Database (Plasmodium falciparum) (Institute of Medical Science, The University of Tokyo, Japan)  
 Microbial Genome Database for Comparative Analysis (MBGD)  
 PEDANT: Genome Analysis and Annotation by MIPS, Germany  
 Profiling of E.coli Chromosome (PEC)  
 Saccharomyces Genome Information Server  
 Synechocystis PCC6803 Gene Annotation Database (SYORF) Bioinformatics Center, Kyoto University and Cyanobacteria Research Community  
 The Institute for Genomic Research

## Microbial Genome Projects: Ex-- *Rhodospseudomonas palustris*

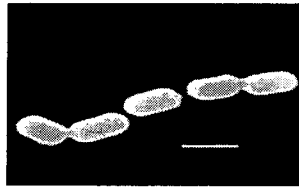
Strain: CGA009  
 Size: 5.47 Mb  
 Contigs: 2  
 Contig Sizes: 5.46 Mb (circular chromosome),  
 8.9 Kb (plasmid)  
 Reads: 117,398  
 Coverage: ~10.7X based on 500bp readlength  
 Currently: Completed 12/13/01



*Rhodospseudomonas palustris*

### Who is *Rhodospseudomonas palustris*?

1. purple non-sulfur phototrophic bacterium
2. found in soils and water
3. converting sunlight to cellular energy
4. absorbing atmospheric carbon dioxide and converting it to biomass.
5. degrade and recycle a variety of aromatic compounds
6. convert nitrogen gas into ammonia,
7. degrading a variety of carbon-containing compounds  
 (including sugars, lignin monomers, and methanol)

Microbial Genome Projects: *Lactobacillus gasseri**Lactobacillus gasseri*

Strain: ATCC33323  
 Size: 1.8 Mb  
 Release: Version 1  
 Total Scaffold Size: 2.0 Mb

1. Lactobacilli are normal inhabitants of the gastrointestinal tract of man and animals
2. Beneficial roles including immunomodulation, interference with enteric pathogens, and maintenance of a healthy intestinal microflora
3. A variety of probiotic activities and roles
  - reduction of fecal mutagenic enzymes
  - adherence to intestinal tissues
  - stimulation of macrophages
  - production of bacteriocins
4. Expected to provide key insights to the survival, roles, and potential benefits of this group of commensal organisms
5. More amenable to DNA introduction and manipulation
  - useful in the functional genomic analysis of this species

## Microbial Genome Databases / portal sites:

TIGR <http://www.tigr.org/tdb/mdb/mdbcomplete.html>  
 GOLD <http://216.190.101.28/GOLD/>  
 DDBJ <http://gib.genes.nig.ac.jp/>  
 EBI <http://www.ebi.ac.uk/genomes/>  
 MBGD <http://mbgd.genome.ad.jp/>  
 NCBI <http://www.ncbi.nlm.nih.gov/PMGifs/Genomes/bact.html>  
 PIR <http://pir.georgetown.edu/pirwww/search/genome.html>  
 STDB <http://www.stdgen.lanl.gov/>  
 InfoBIOGEN [http://www.infobiogen.fr/doc/data/complete\\_genome.htm](http://www.infobiogen.fr/doc/data/complete_genome.htm)  
 MAGPIE <http://www-fp.mcs.anl.gov/~gaasterland/genomes.htm>  
 ICBS <http://www.cbs.dtu.dk/services/GenomeAtlas/>  
 KEGG [http://www.genome.ad.jp/kegg/catalog/org\\_list.html](http://www.genome.ad.jp/kegg/catalog/org_list.html)  
 NIAID <http://www.niaid.nih.gov/dmid/genomes/genome.htm>  
 Completely Sequenced Genomes <http://linkage.rockefeller.edu/wli/seq/>  
 CyanoBase <http://www.kazusa.or.jp/>  
 ParasiteDB <http://www.ebi.ac.uk/parasites/paratable.html>



parasite-genome

## Microbial Cell Research

**1. Biological Basics Goal:** Determine and characterize the minimum set of genes and corresponding gene products necessary to sustain a simple free-living microbial cell, express the genes to produce the relevant proteins, and determine their structure.

### 2. Functional Foundations

Goal: Determine the physiological and biochemical functions of the gene and specific bioprocesses using standard biochemical techniques and structural/ computational biology.

### 3. Modeling Interactions

Goal: Use high-end computing to model gene-gene, gene-protein, and protein-protein interactions as well as the internal biochemistry of the cell.

### 4. Regulation and Manipulation

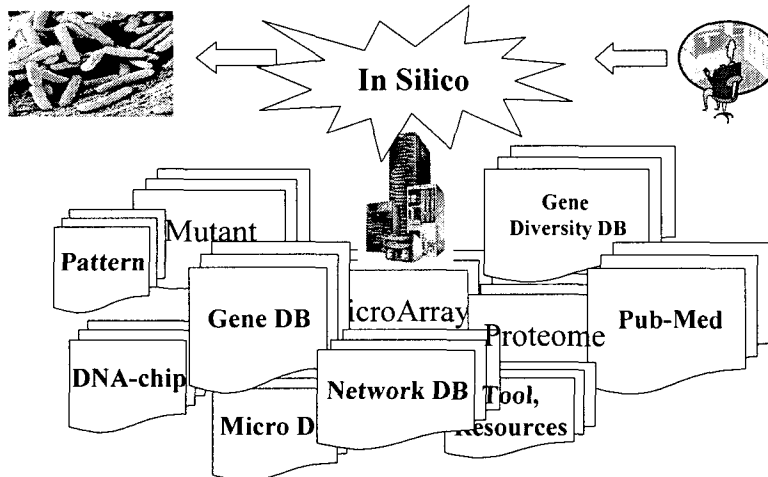
Goal: Use gene-protein manipulation to enhance or suppress various cell functions.

### 5. Functional Expression

Goal: Focus on functions that are relevant to DOE goals (e.g., bioremediation, carbon sequestration, and sustainable energy production).

## *In Silico* Microorganism

통합 검색 분석, 유전자 기능 예측 및 simulation, 원하는 미생물 디자인





***In Silico* Microorganism**Genomatica To develop "Next-Generation" *In Silico* Technology

Start: Jan 3, 2002

Organism: Microbe

Fund: Department of Energy—\$ 2 Million

Mission: Generating of integrated models of metabolism and metabolic regulation

"Cybernetic Cells" explores the movement by biologist and companies to simulate the biochemistry of living cell, with the ultimate goal of finding a way to perform virtual experiment *In silico* that can speed up the discovery of new drugs and Reduce their cost

## Microbial genome sequencing: applications and future directions

**IMMEDIATE****DNA**

IDENTIFY GENES, ORFS, ETC.  
 TRANSCRIPTION/TRANSLATION  
 START/STOPS  
 NUCLEOTIDE COMPOSITION  
 "WORDS", CONSENSUS MOTIFS  
 CODON USAGE  
 GENOME STRUCTURE

**GENES**

ORTHO-/PARA-/HOMOLOGS  
 GENE FAMILIES  
 PREDICTED CELL LOCALIZATION  
 MOTIFS

**POST-GENOMIC STUDIES****LAB EXPERIMENTS**

GLOBAL GENE EXPRESSION  
 GLOBAL PROTEIN ANALYSIS  
 GLOBAL MUTAGENESIS  
 (knock-outs, reporter genes)  
 PROTEIN OVEREXPRESSION  
 SCREENS/ASSAYS  
 (gene function, drug design,  
 virulence)

**"IN SILICO STUDIES"**

COMPARATIVE GENOMICS  
 3-D STRUCTURAL PREDICTIONS  
 BIOCHEMICAL PATHWAYS  
 GENE EXPRESSION HIERARCHY

**FUTURE**

Comprehensive understanding of:

PHYSIOLOGY  
 GENETICS  
 EVOLUTION  
 BIOCHEMISTRY



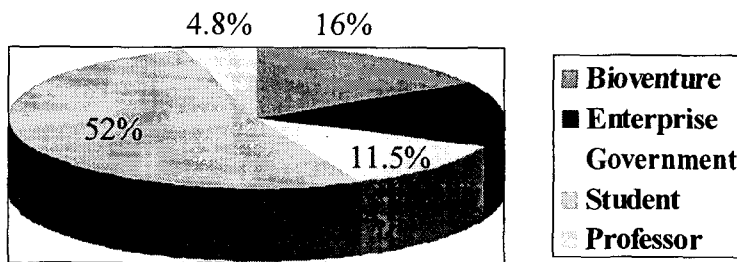
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Who...in Korea?  
Research Fund  
Korean BioTech  
R&D Program  
Korean Bioinformatics  
Government?



Who.....in Korea?

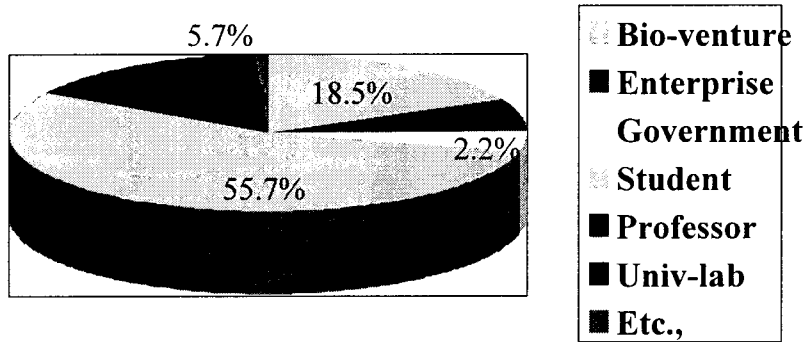
**Average number of participants in the workshops held in 2001 by Korean Society for Bioinformatics**



Provided by KSBI

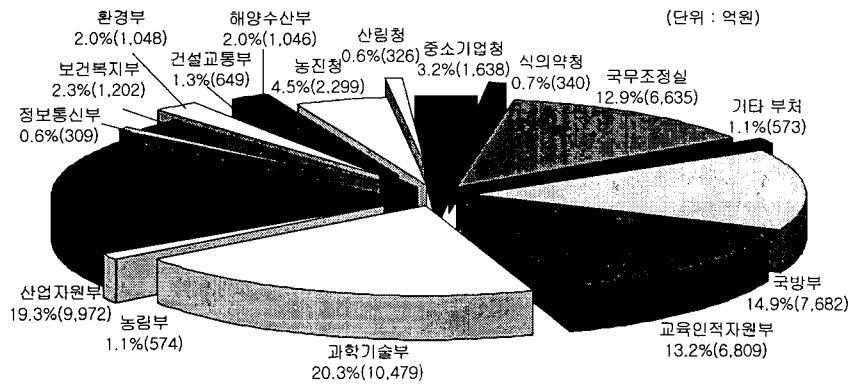
Who.....in Korea?

2002 The First workshop : Participants



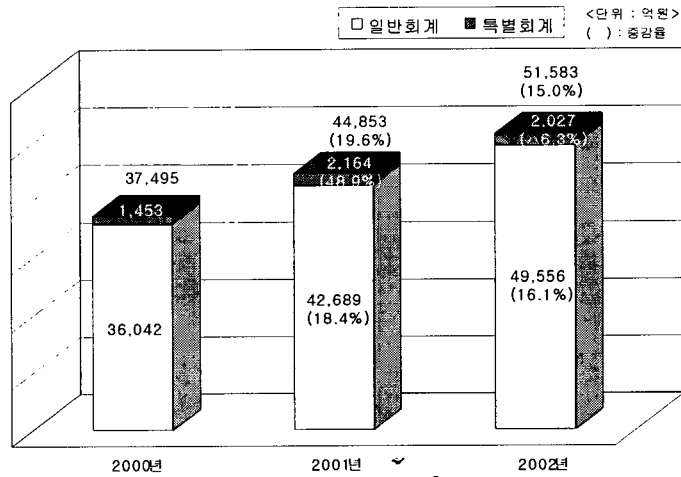
Provided by KSBI

2002년도 부처별 연구개발예산 분포



2002년 R&D 예산 : 5조 1,583억원

정부연구개발예산의 추이(2000~2002년)



자료: 과학기술부 한국과학기술기획평가원

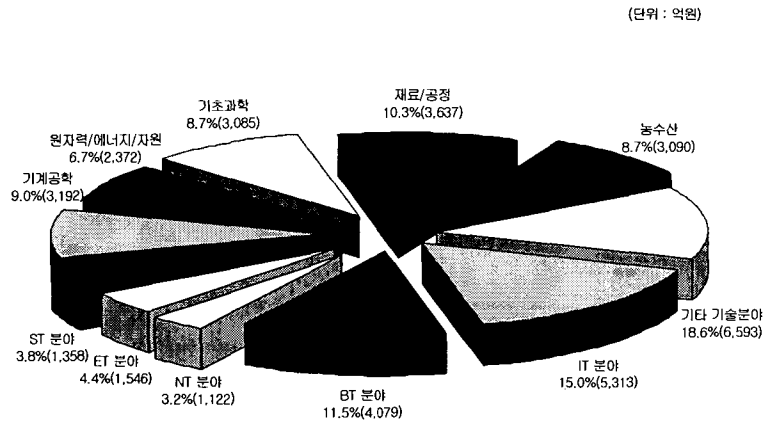
미래 유망 신기술분야 예산편성 현황

(단위:억원)

| 구분         | 2000년 | 2001년(A) | 2002년(B) | 증감 (B-A) | %     |
|------------|-------|----------|----------|----------|-------|
| 정보기술(IT)   | 4,085 | 4,536    | 5,313    | 777      | 17.1  |
| 생명공학기술(BT) | 2,462 | 3,353    | 4,079    | 726      | 21.7  |
| 나노기술(NT)   | 300   | 425      | 1,122    | 697      | 163.7 |
| 환경기술(ET)   | 955   | 1,328    | 1,546    | 218      | 16.4  |
| 우주항공기술(ST) | 721   | 1,212    | 1,358    | 146      | 12.0  |
| 합계         | 8,563 | 10,854   | 13,418   | 2,564    | 23.6  |

자료: 과학기술부 한국과학기술기획평가원

### 2002년도 기술분야별 예산분포

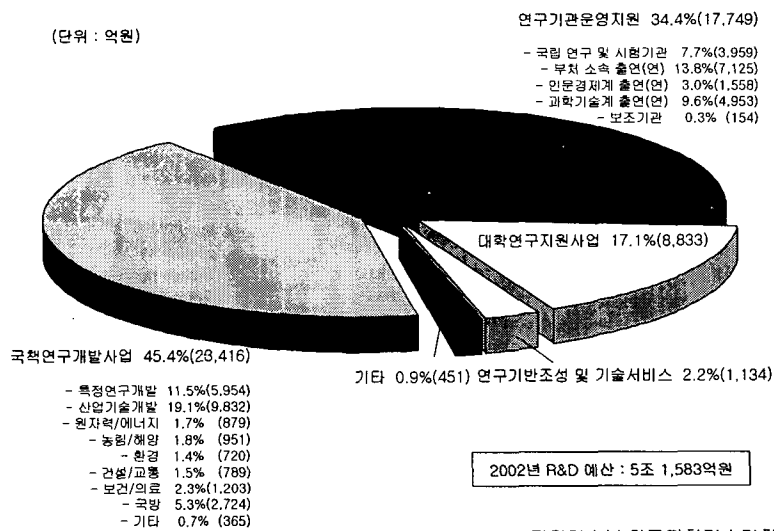


기술분류 가능 2002년 R&D 예산 : 3조 5,387억원

자료: 과학기술부 한국과학기술기획평가원

### 2002년도 기능별 연구개발예산 분포

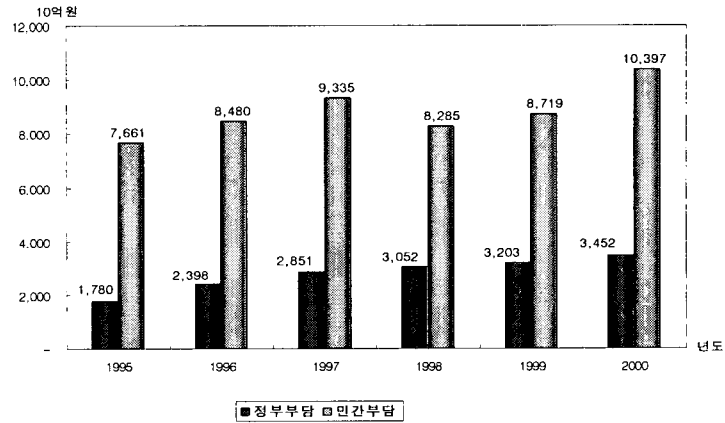
(단위 : 억원)



2002년 R&D 예산 : 5조 1,583억원

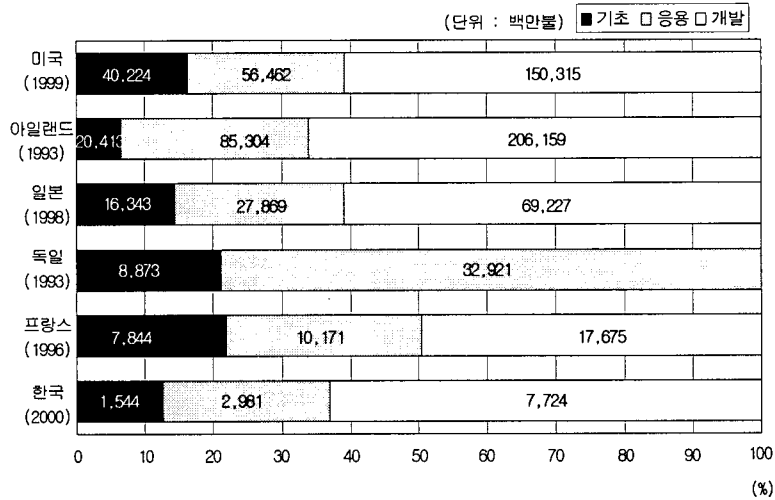
자료: 과학기술부 한국과학기술기획평가원

총 연구개발비의 자원별 추이(1995~2000년)



자료: 과학기술부 한국과학기술기획평가원, 「2001 과학기술연구활동조사보고」, 2001.

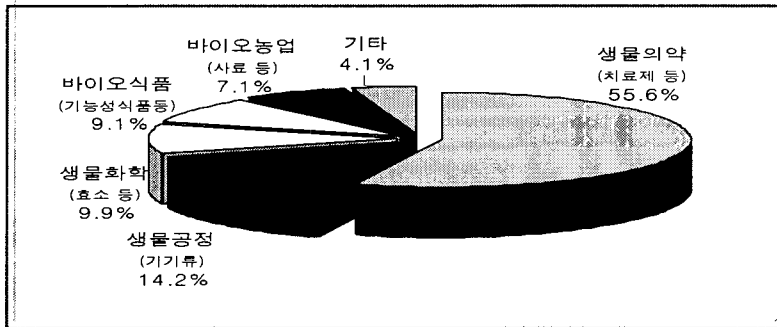
주요국의 성격별 총 연구개발비 구성 국제비교



<자료원>: 과학기술부 한국과학기술기획평가원, 「2001 과학기술연구활동조사보고」, 2001.

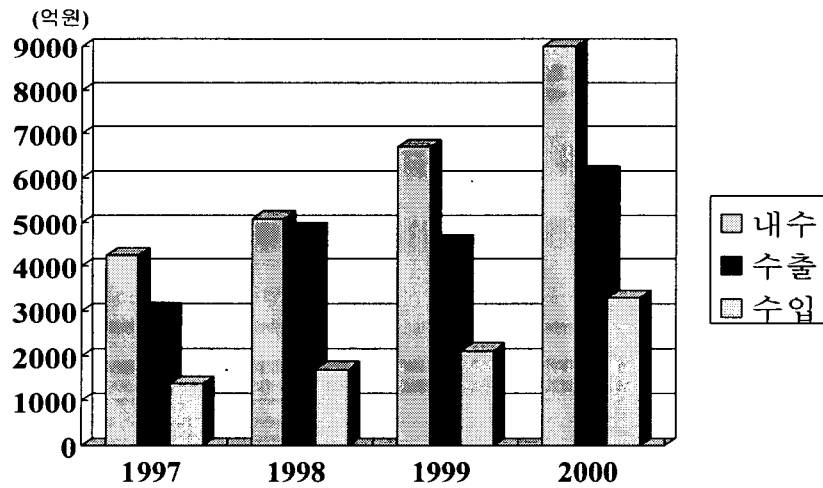
### 2000년 국내 바이오산업의 분야별 구성비

| 분야          | 구성비   | 분야         | 구성비  |
|-------------|-------|------------|------|
| 생물의약(치료제 등) | 55.6% | 바이오식품      | 9.1% |
| 생물공학(기기류)   | 14.2% | 바이오 농업(사료) | 7.1% |
| 생물화학(효소 등)  | 9.9%  | 기타         | 4.1% |

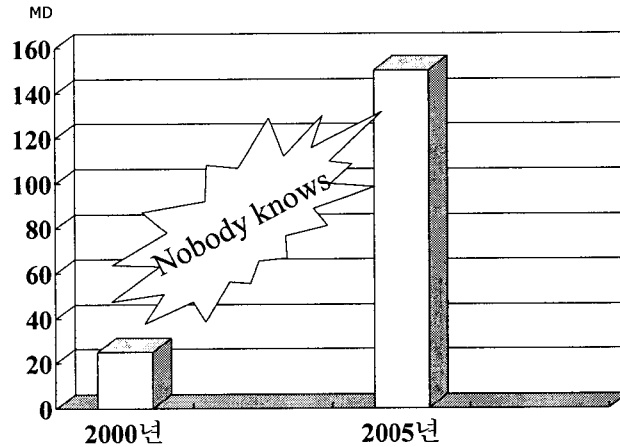


자료: 생물산업협회, 국내 생물산업 실태조사표

### 국내 바이오 산업 시장규모



자료: 생물산업협회, 국내 생물산업 실태조사표



From LG Economic Research Institute

| 년도 | 1994 | 1996 | 1998 | 1999 | 2000 | 2001 | 2002 |
|----|------|------|------|------|------|------|------|
| 억원 | 536  | 1234 | 1115 | 1608 | 2462 | 3238 | 4500 |

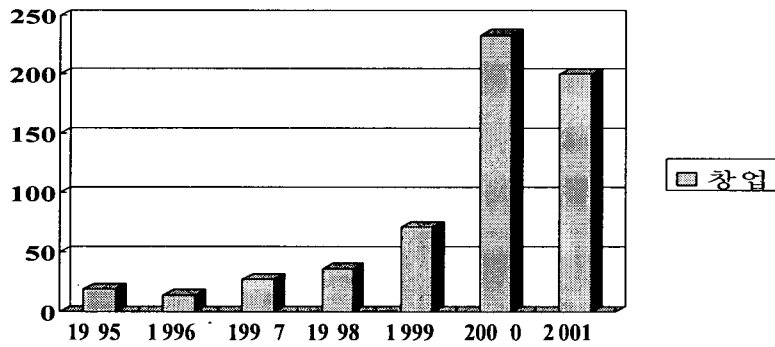
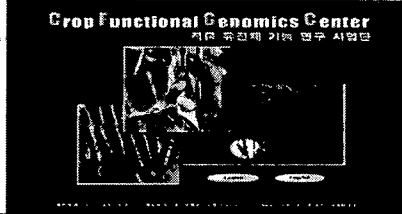
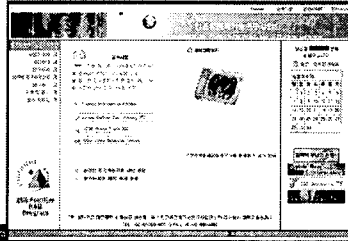
- ▶ 2000년을 기준으로 대폭 증가
- ▶ R&D 예산대비 바이오분야 비중은 8%수준
  - 94년(3.5%)→98년(5%)→2001년(8%)
  - 98년 이후 증가율: 바이오분야 46%, 정부전체 :19.4%
- ▶ 2002년 기준 정부총액 투자규모:
  - Amgen(8.5억\$/년)의 1/3수준



Genomics and Proteomics R&D Program

: Government grants of US\$10 million per year for 9 years

- >> The Center for functional Analysis of Human Genome
- >> Crop Functional Genomics Center
- >> Plant Diversity Research Center
- >> Human Proteomics Center
- >> Drug Discovery Center
- >> Medical Genomics program



- 현재 약 600여개로 추정
- 수익모델 약화로 수적양적인 구조조정이 이루어지고 있음

*Bioventure**Research fields*

|                   |   |
|-------------------|---|
| Macrogen :        | DNA chip, genome analysis, Bioinformatics solution      |
| Neodin :          | Medical diagnosis kit                                   |
| DigitalGenomics : | Medical diagnosis kit                                   |
| Bioinfomatxs:     | Genome and proteome analysis tool,                      |
| VectorKorea:      | Gene therapy  |
| IDR:              | Protein prediction and modeling, New drug development   |
| DNALink:          | New drug development                                    |
| Bioneer:          | Bio-vendor, sequencing machine and new drug development |
| Genomictree :     | DNA chip and sequencing service                         |
| Toolgen:          | New drug development                                    |
| Inbionet:         | Genome analysis and data mining,                        |
| Proteogen:        | Protein chip, Protein analysis machine                  |
| Badasoft:         | Bioinformatics solution, System development             |
| Cristalgenomics:  | New drug development, structural Proteomics             |
| Etc....,          |   |

From Korean Bioventure Association

**Industrial Research Fields-Enterprise**

- **Samsung SDS:** Genome analysis portal solution, Network solution, Analysis tools of Proteome and DNA chip
- **LG Chemicals:** New Drug discovery, Gene discovery  
Genome research.
- **SK:** New Drug discovery and Gene therapy
- **Hanhwa:** New Drug discovery in cardinal and nerves diseases

## SWOT Analysis

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**Strength(S)**

Information Technology Infra  
Need Power  
Man-Power in Biotechnology

**Opportunity(O)**

Large scale market  
Start step  
Many kinds of research fields

**Weakness(W)**

A few specialist in bioinformatics  
Government Support  
A few special Institute  
Infra in Biotechnology

**Treat(T)**

Competition with an advanced Country  
Endeavor of multinational Enterprise

## What does Government do?

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- ▶ Providing scientific, technical, organizational and policy advice
- ▶ Supplying information and information products
- ▶ Promoting new technologies, strategies and tools
- ▶ Providing training and/or information on training opportunities
- ▶ Bringing people and institutions together at national and sub-regional levels
- ▶ Assisting in writing project proposals and obtaining funds
- ▶ Implementing, coordinating and publishing research
- ▶ Allocating direct financial support to infrastructure development and activities



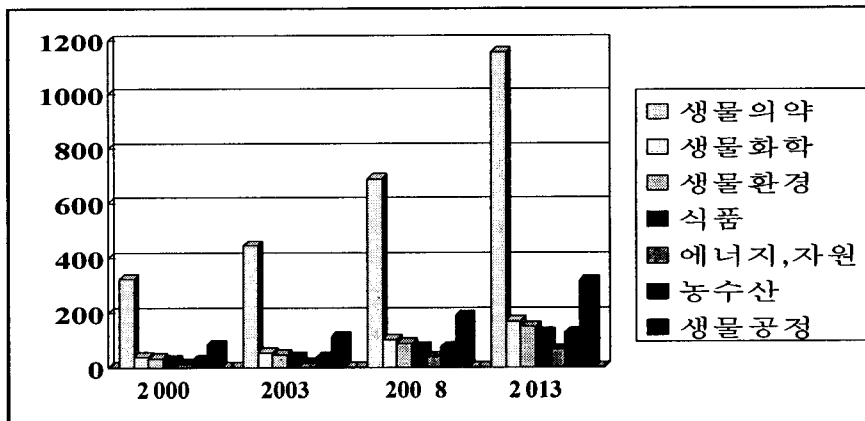
Market  
Changing Marketplace  
Pharma and Biotech  
Drug discovery  
Prospects



1. Introduction
2. Microbial Genome
3. Korean Bioinformatics
4. Bioinformatics Market
5. Education
6. BRIC's Bioinformatics



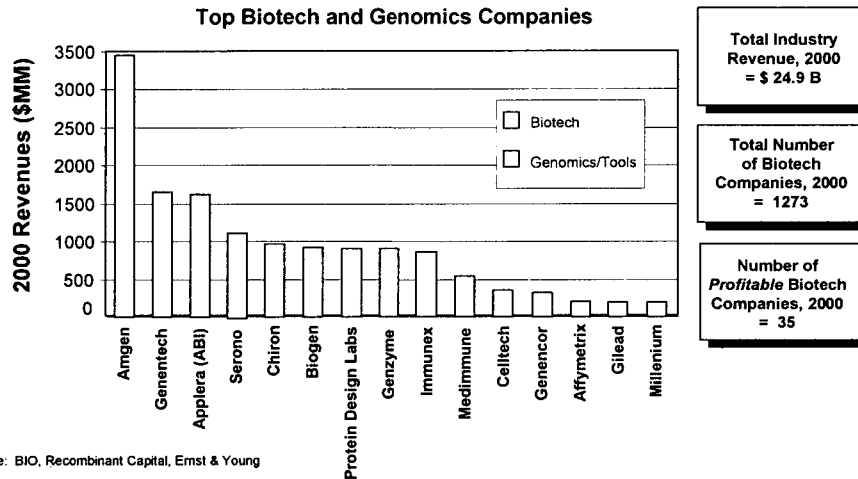
(단위: 억불)



\* 자료 : OECD, *Biotechnology & Trade*, 1997  
CRC, *U.S. Biotechnology Product Sales Forecast*, 1997

## Biotech and Genomics Companies(2000 Revenue)

The Biotech landscape is dominated by a few key players, a dozen smaller companies, and hundreds of small start-ups.



## The Changing Marketplace of Bioinformatics

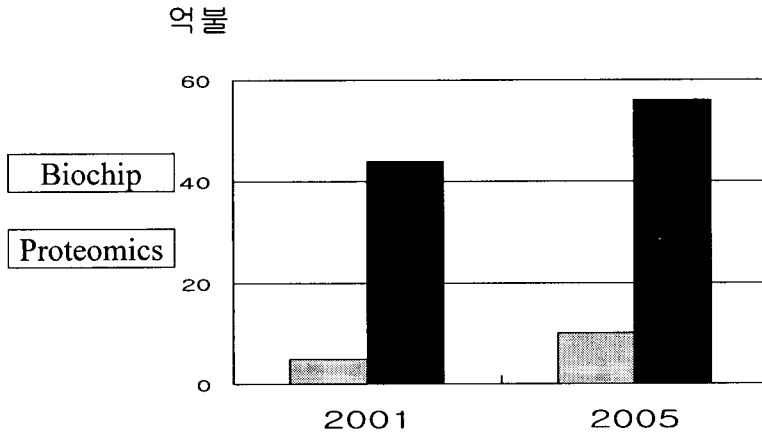
Several roles are evolving for bioinformatics in life science

- ▶ Automation of biological and chemical research for the drive toward high-throughput procedures
- ▶ Providing tools for the capture, management and analysis of disparate data
- ▶ Facilitating the exchange and dissemination of information between isolated groups in large organizations

### Emerging niches

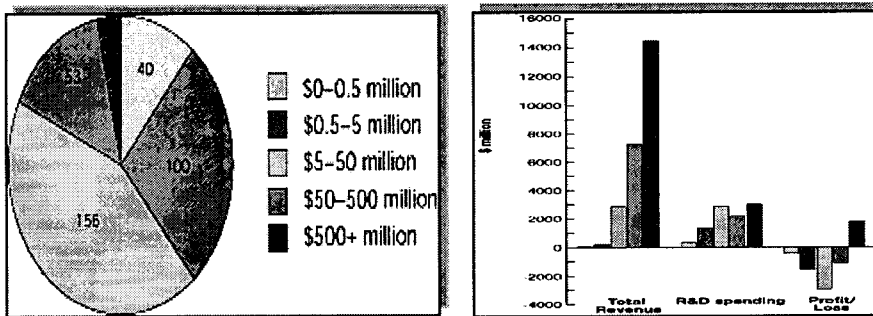
- ▶ Providing software packages for data visualization, interpretation, and Analysis  
: ex—Spotfire(Cambridge, MA), MDL(SanLeandro, CA), Silicon Genetics(SanCarlos, CA)
- ▶ Offering comprehensive bioinformatics platforms accessible to life scientists over internet  
: ex—DoubletWist(Oakland, CA), Compugen/LabonWeb(Tel Aviv, Israel), e-Bioinformatics
- ▶ To rent out Data architectures/infrastructure and bioinformatics expertise for data integration  
: ex—LionBioScience(Heidelberg, Germany), IBM, Motorola
- ▶ Biological Research-based Companies are offering tools along with their contents  
: ex—Incyte Genomics, Celera Genomics, Curagen(New Haven, CT)

Huge Market Opportunities



Citation by Genomic Solutions, 2001

Biotechnology Companies by Revenue



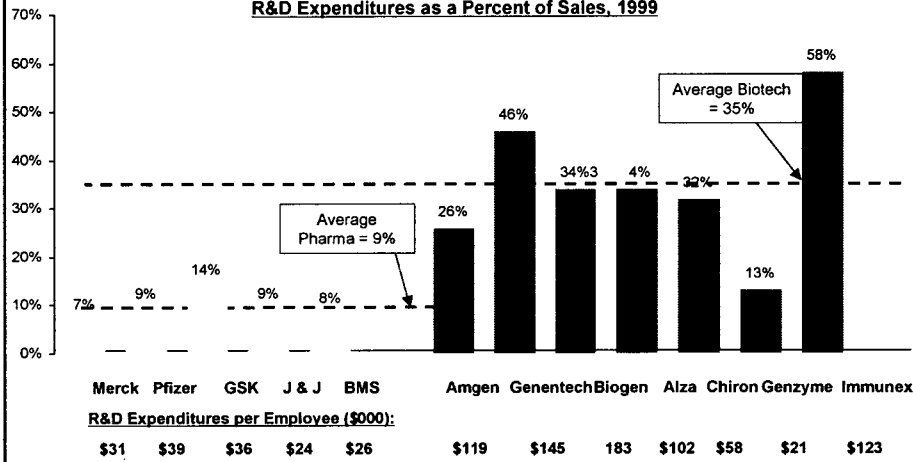
**Distribution of publicly traded biotechnology companies according to revenue generated. Total revenue, R&D spending, and profit/loss for the five categories of company described**

Nature Biotechnology 19, 407 - 412 (2001)

### Pharma and Biotech in R&D

Biotech's high % of R&D expenditures reflect a focus on innovation and a tendency to license out products as they near the commercialization phase.

**R&D Expenditures as a Percent of Sales, 1999**



**R&D Expenditures per Employee (\$000):**

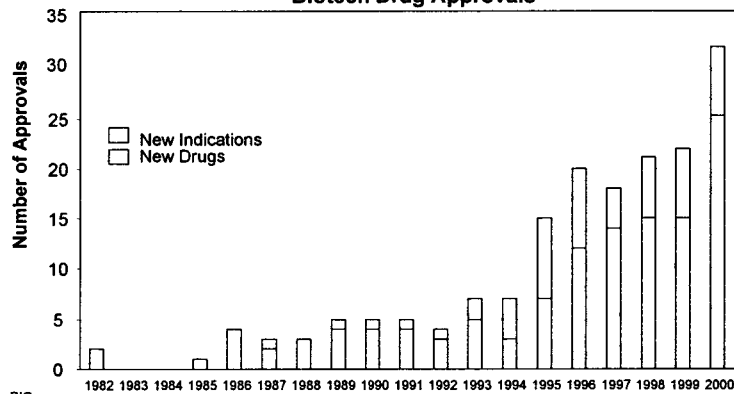
|      |      |      |      |      |       |       |     |       |      |      |       |
|------|------|------|------|------|-------|-------|-----|-------|------|------|-------|
| \$31 | \$39 | \$36 | \$24 | \$26 | \$119 | \$145 | 183 | \$102 | \$58 | \$21 | \$123 |
|------|------|------|------|------|-------|-------|-----|-------|------|------|-------|

Source: Annual reports, Ernst & Young

### Change of Drug Market

The number of Biotech drugs on the market has skyrocketed since 1994, and the trend is expected to continue.

**Biotech Drug Approvals**



Source: BIO

*An additional 350 biotech medicines targeting over 200 diseases were in late-stage development in 2001, promising to continue the upward trend in approvals.*

## Mergers and acquisitions *between Pharma and Biotech*

*The distinction between Pharma and Biotech is becoming blurred as the industry consolidates.*

Many of the top Biotechs are partially or entirely owned by traditional Pharma companies

Genentech and Roche - Chiron and Novartis  
 Immunex and AHP - Centocor and J&J

Mergers and acquisitions among Biotechs are also common

Millennium Pharmaceuticals and LeukoSite.

Genzyme and Cell Genesys, also Peptimmune.

MedImmune and U.S. Bioscience.

ALZA and Sequus Pharmaceuticals .

Few very small integrated pharmaceutical companies exist, as economies of scale in drug development, sales, and marketing are required to drive down costs and maintain growth rates

## Biotechnology company casualties during 2000

Table 3. Biotechnology company casualties during 2000

|  |  |   |
|--|--|---|
| <b>Mergers</b>   | Agrilone, by Exelixis                          | Bioscience                                    |
| Algos Pharmaceutical and Endo Pharmaceuticals  | Anesta, by Cephalon                            | NovoPharm Biotech became Viventia Biotech     |
| Atlix Biopharmaceuticals and NPS Pharmaceuticals (1999)  | Cambridge NeuroScience, by CeNeS               | Peptide Therapeutics became Acambis           |
| Aquila Biopharmaceuticals and Antigenics   | Cotrix Pharmaceuticals, by Insmed              | QLT PhotoTherapeutics became QLT              |
| Biomatrix with Genzyme Tissue Repair and Genzyme Surgical Products, to form Genzyme Biosurgery | ChRx, by Rhodia                                | UroGen changed name to GenStar (and relisted) |
| GelTex Pharmaceuticals and Genzyme Biopool and Xtrana  | CliniChem, by Biochem                          | Vanguard Medica changed name to Vernalis      |
| Celltech Chromscience and Medava, forming Celltech Group                                       | Crescendo Pharmaceuticals, by Alza             | Modi-Ject changed name to Antares Pharma      |
| Coulier Pharmaceutical and Corixa  | Trega Biosciences, by Lion Bioscience          |   |
| Creative Biomolecules with Ontogeny and Regeneration, to form Curs                             | Spiros, by Dura Pharmaceuticals                |   |
| Epitope and STC Technologies   | Oxford Molecular, by Pharmacia                 | <b>Delisted</b>                               |
| Evotec BioSystems and Oxford Asymmetry   | Pathogenesis, by Chiron                        | Amgen Biosciences                             |
| Life Technologies and Invitrogen   | L.J. Biosystems, by Molecular Devices          | BioSante Pharmaceuticals                      |
| Lunar and GE Medical Systems   | North American Vaccine, by Baxter              | Cadus Pharmaceutical                          |
|  | Quadrant Healthcare, by Elan                   | Clinico                                       |
| <b>Acquisitions</b>  | Liposome, by Elan                              | Ecogen  |
| Accumed, by Ampersand  | Dura Pharmaceuticals, by Elan                  | GalaGen                                       |
|  |  | Hybridon                                      |
|  | <b>Name changes</b>                            | Sedalia Holdings                              |
|  | CytoTherapeutics became StemCells              | Unigene                                       |
|  | Energy Biosystems became Enchira Biotechnology |   |
|  | Ethical Holdings became Amann                  | <b>Relistings</b>                             |
|  | Meridian Diagnostics became Meridian           | Antex Biologics                               |
|  |  | Epoch Pharmaceuticals                         |

Nature Biotechnology 19, 407 – 412 (2001)



## Mergers and acquisitions *between Pharma and Biotech*

Alliances and licensing have been instrumental in Biotech industry growth, but are becoming more central to Pharma's business model as well.

### Advantages for Pharma

- Efficient innovation
- Access to new technologies
- Diversification of development risk
- Enhanced product pipelines
- Downside: lack of control over pipeline

Pharma needs Biotech more now than in the past for drug discovery and new products as their scale makes sustainable growth difficult

### Advantages for Biotechs

- Access to resources
- Marketing expertise
- Established sales force
- Downside: revenue-sharing and postponement of value-chain integration

Biotechs need pharma less due to multiple sources for manufacturing and marketing, and downstream integration.

## Biotechnology Financing, 2000~2001

### Biotechnology financing, 2000–2001

| Quarter | Total<br>(billions<br>of US \$) | US    | Europe* |
|---------|---------------------------------|-------|---------|
| Q3 2001 | 2.46                            | 1.27  | 1.06    |
| Q2 2001 | 5.39                            | 4.54  | 0.59    |
| Q1 2001 | 2.34                            | 1.91  | 0.24    |
| Q4 2000 | 10.23                           | 8.69  | 1.53    |
| Q3 2000 | 8.78                            | 6.12  | 2.52    |
| Q2 2000 | 4.65                            | 3.17  | 1.30    |
| Q1 2000 | 13.09                           | 11.71 | 0.52    |

\*For all offerings in Austria, Belgium, Denmark, Finland, France, Germany, Hungary, Iceland, Ireland, Israel, Italy, Netherlands, Norway, Spain, Sweden, Switzerland, and UK  
Source: BioCentury.

## Next Waves of Innovation in Drug Discovery

Table 1. Three waves of innovation in drug discovery

| Innovation   | Tools/technologies  | Representative companies  |
|--|---|---|
| <b>First wave</b><br>Genomics<br>(target discovery,<br>anti-sense therapeutics,<br>and ultimately<br>gene therapy)                   | Gene / genome sequencing and<br>expressed sequence databases<br>Full-length cDNAs/expressions<br>Functional genomics<br>RNA expression profiling<br>Transgenics / directed mutagenesis<br>Antisense Technology<br>SNP databases<br>Gene mapping / disease genes<br>Gene diagnostics       | Incyte (Palo Alto, CA); Celera (Rockville, MD); Genset (Paris)<br>Genome Therapeutics (Walham, MA)<br>Clontech (Palo Alto, CA); Stratagene (La Jolla, CA);<br>Life Technologies (Rockville, MD);<br>Coragen (New Haven, CT); Millennium (Cambridge, MA);<br>and Pharmagene (Cambridge, UK)<br>Aflynnex (Santa Clara, CA); Gene Logic (Gaithersburg, MD);<br>Ridella Pharmaceuticals (Kirkland, WA)<br>Lexicon Genetics (Houston, TX); Genomic Systems (St. Louis, MO);<br>Ilex Pharmaceuticals (Carlsbad, CA); Hybridon (Cambridge, MA);<br>Virogenics (Cambridge, MA); Genasense (New Haven, CT); Celera<br>Oxigen (Boulder, CO); Myriad (Salt Lake City, UT); Decode (Reykjavik, Iceland);<br>Quest (Trenton, NJ); Roche (Nutley, NJ); Duocore (Palo Alto, CA); Affymetrix      |
| <b>Second wave</b><br>Proteomics<br>(target validation,<br>drug screening,<br>antibody<br>therapeutics, and<br>protein therapeutics) | Protein databases<br>and protein expression analysis<br>Protein expression technologies<br>and protein therapeutics<br>Directed evolution<br>Antibody engineering<br>High-throughput screening<br>Protein interaction databases<br>Affinity selection<br>Protein pathways / protein chips | Galathea GlycoSciences (Gatford, UK); Large Scale Biology (Granville, CA);<br>Protonix (Cambridge, MA)<br>Amgen (Thousand Oaks, CA); Genentech (S. San Francisco, CA)<br>Human Genome Sciences (Rockville, MD); Chiron (Emeryville, CA);<br>Genetica Institute (Cambridge, MA); Lanza (Saugus, UK)<br>Makymex (Redwood City, CA); Phylis (Lowington, MA);<br>Celltech/Midarex (Eastleigh, UK); Abgenix (Fremont, CA);<br>Cambridge Antibody Technology (Cambridge, UK)<br>Aurora Biosciences (La Jolla, CA); Cambridge Drug Discovery (Cambridge, UK)<br>Proteome, MDS Proteomics (Baltimore, MD); Genada; Celera<br>Norgenesus (Cambridge, MA); MDS-Proteomics<br>Zymyx (Hayward, CA); Cambiotech (Seattle, WA); CIPHERgen (Palo Alto, CA);<br>Sensio Proteomics (Cambridge, UK) |
| <b>Third wave</b><br>Molecular design<br>(protein therapeutics,<br>antibodies, and<br>small molecules)                               | Protein structure determination<br>Protein homology modeling<br>Protein engineering<br>Structure-based small molecule design<br>Molecular design tools  | Structural Genomics (San Diego, CA); Syntex (San Diego, CA); Astor (Cambridge, UK)<br>Structural Bioinformatics (San Diego, CA); Geneformix (San Diego, CA)<br>Sunesis (Redwood City, CA); Sangamo (Richmond, CA)<br>Cambridge Antibody Technology (Cambridge, UK)<br>Vertex Pharmaceuticals (Cambridge, MA); De Nova (Cambridge, UK)<br>Molecular Simulations (San Diego, CA); Topos (St. Louis, MO)   |

Nature Biotechnology 19, 207 - 209 (2001)

## Prospects

### New trends in Biology

- **New research trend in biology.**
- **Systems biology**
  - A unique approach to the study of genes and proteins which has only recently been made possible by rapid advances in computer science.
  - Unlike traditional science which examines single genes or proteins, systems biology studies the complex interaction of all levels of biological information:
    - Genomics DNA, mRNA, proteins; functional proteins, informational pathways and informational networks to understand how they work together.

## Prospects

### New needs in Biology

#### ➤ Changes of Meaning

-The role of Bioinformatics will be critical and expand.

- In the narrow view: tools for storing, processing, and searching data.
- In the broad view: discipline for creating new knowledge in Biology.

#### ➤ Into a more quantitative science.

- sophisticated mathematical, statistical, and data modeling approach.

#### ➤ More integrative Approach

- Mathematician, Statistician, Physicist, and Computer Scientist
- Engineering, Linguistics, Fluidics, Robotics, ...
- Linguistics:
  - Develop languages for Biology in the view of formalism.

## Trends in Microbial Genomics and Bioinformatics

보도자료  
전문인력  
교육과정



1. Introduction
2. Microbial Genome
3. Korean Bioinformatics
4. Bioinformatics Market
5. Education
6. BRIC's Bioinformatics

연합뉴스(01/5/29): 바이오산업 핵심 생물정보학 인력 “절대부족”

매일경제(01/6/4): 생물정보학 교육인력 부족

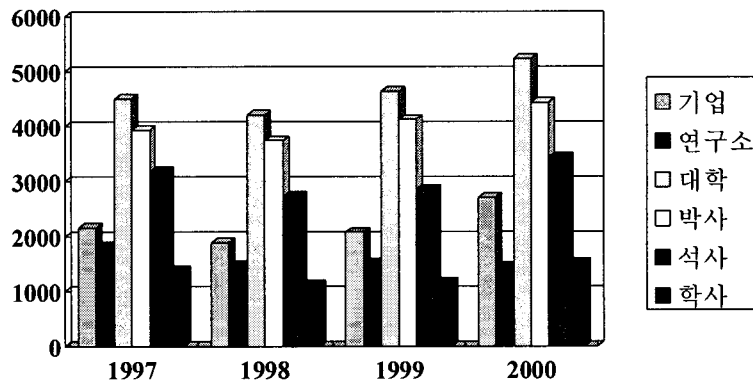
연합뉴스(01/07/19): 산자부, 서울대 BT인력 재교육 나선다

전자신문(01/8/20): 바이오업계 인력난 몸살

과학신문(01/8/29): 바이오인포매틱스 인력양성 차질

한국경제(01/10/27): 일부 바이오업체, 사업다각화 고급인력 확충나서

약업신문(01/11/15): 산자부, BT 인력 육성에 5,000억원 투입



- ▶ 대학: 5224명, 기업: 2,701명, 연구소: 1,467명
- ▶ Genomics, Proteomics, Bioinformatics분야:
- ▶ 2005년 인력 수급차질 예상: 수요—9470명, 공급—3080명(6,300명 정도 부족예상)

자료: 2002년 생명공학육성시행 계획

| 학교    | 구분   | 소속    | 학사 | 석사 | 박사 | 개강년도   | 기타       |
|-------|------|-------|----|----|----|--------|----------|
| 부산대학교 | 협동과정 | 대학원   |    | ◆  | ◆  | 2000.5 | 국내최초     |
| 숭실대학교 | 학과설립 | 자연대   | ◆  |    |    | 2001.3 | 학부최초     |
| 서울대학교 | 협동과정 | 생명과학부 |    | ◆  | ◆  | 2002.3 | 다수의 교수진  |
| KAIST | 학과설립 | 공학부   | ◆  | ◆  | ◆  | 2002.9 | 학부, 대학원  |
| 중부대학교 | 학과설립 | 생명과학부 | ◆  |    |    | 2002.3 |          |
| 국립보건원 | 단기과정 |       |    |    |    | 2001.6 | 1회 실시 종료 |

### 향후 교육과정

1. 대학원 협동과정 중심으로
2. 특화된 분야 과정
3. 재교육—수강자에 따라 다양한 프로그램개발
4. On-line 교육 활성화
5. 전문강사 확보를 위한 장기프로그램

### Some degree programs in bioinformatics/computational biology

| School  | Degree | URL   |
|---|--------|---|
| Boston University (Boston, MA)                                | MS/PhD | <a href="http://bioinformatics.bu.edu/">http://bioinformatics.bu.edu/</a>   |
| George Mason University (Fairfax, VA)                         | MS/PhD | <a href="http://www.science.gmu.edu/~michaels/bioinformatics">http://www.science.gmu.edu/~michaels/bioinformatics</a>       |
| Georgia Tech (Atlanta, GA)                                    | MS     | <a href="http://www.biology.gatech.edu/bioinformatics/">http://www.biology.gatech.edu/bioinformatics/</a>                   |
| Iowa State University (Ames, IA)                              | PhD    | <a href="http://www.bcb.iastate.edu/">http://www.bcb.iastate.edu/</a>   |
| Medical College of Wisconsin/Marquette Univ. (Milwaukee, WI)  | MS     | <a href="http://golab.mch.mcw.edu/AP/announcements/initial.html">http://golab.mch.mcw.edu/AP/announcements/initial.html</a> |
| Rensselaer Polytechnic Institute (Troy, NY)                   | BS     | <a href="http://www.rpi.edu/dept/bio/infor/bioinformatics.html">http://www.rpi.edu/dept/bio/infor/bioinformatics.html</a>   |
| Rutgers University (New Brunswick, NJ)                        | PhD    | <a href="http://cmb.rutgers.edu/">http://cmb.rutgers.edu/</a>   |
| University of the Sciences in Philadelphia (Philadelphia, PA) | MS     | <a href="http://www.usip.edu/bioinformatics/">http://www.usip.edu/bioinformatics/</a>                                       |

### Degree programs that offer specialization/emphasis in bioinformatics

| School  | Degree  | URL   |
|---|---|---|
| Baylor University (Waco, TX)                  | BS/Informatics  | <a href="http://ecswwww.baylor.edu/ecsw/computer_science/undergrad.htm">http://ecswwww.baylor.edu/ecsw/computer_science/undergrad.htm</a> |
| North Carolina State Univ. (Raleigh, NC)      | MS, PhD/Genomics  | <a href="http://genomics.ncsu.edu/">http://genomics.ncsu.edu/</a>   |
| University of California, San Diego           | PhD/Comp. Sci., Eng., Math., Chemistry, Physics, etc.   | <a href="http://www.ogsr.ucsd.edu/bioinformatics/program.htm">http://www.ogsr.ucsd.edu/bioinformatics/program.htm</a>                     |
| University of Pennsylvania (Philadelphia, PA) | BS/MS/PhD, Computer Sci., Biotech., Biology, Math.      | <a href="http://www.cbi.upenn.edu/UPCBF/">http://www.cbi.upenn.edu/UPCBF/</a>   |
| University of Pittsburgh                      | PhD/Computer Sci., Biol. Sci., Chemistry, Physics, etc. | <a href="http://www.cs.pitt.edu/teck/program.html">http://www.cs.pitt.edu/teck/program.html</a>   |
| Carnegie Mellon (Pittsburgh, PA)              |   |   |
| Washington University (St. Louis, MO)         | PhD/Medicine, D.Sc./Biomed. Eng.                        | <a href="http://www.bcb.wustl.edu/CMB/">http://www.bcb.wustl.edu/CMB/</a>   |

Nature Biotechnology 19, 285 - 286 (2001)

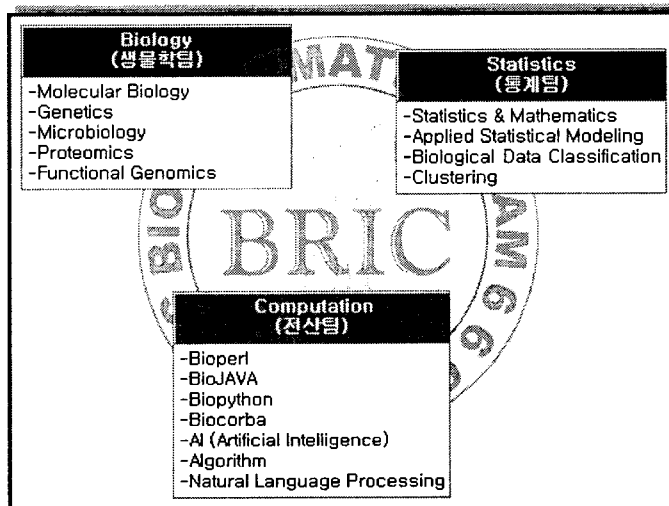
## Trends in Microbial Genomics and Bioinformatics

**Bioinformatics Team**  
대외 활동사항  
**KSDB**  
Service Items

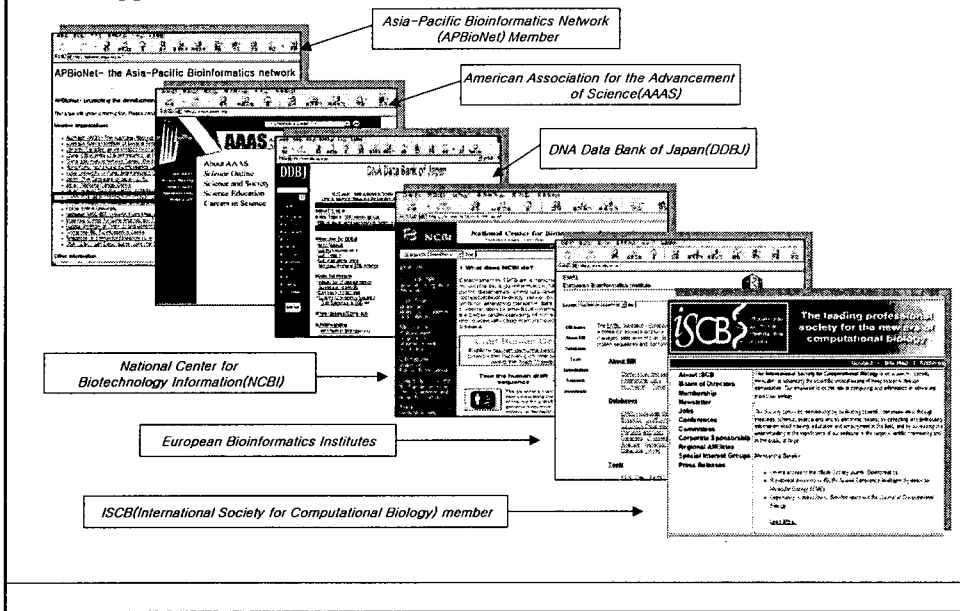


1. Introduction
2. Microbial Genome
3. Korean Bioinformatics
4. Bioinformatics Market
5. Future of Bioinformatics
6. BRIC's Bioinformatics

## BRIC's Bioinformatics Team



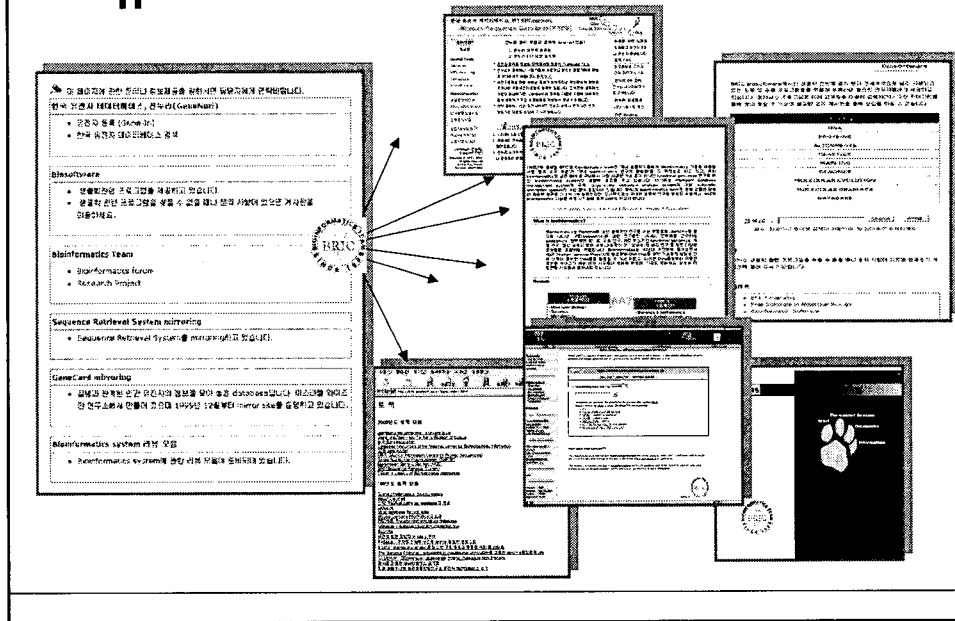
### 생물정보학관련 BRIC의 대외 활동



### Korean Sequence Database(KSDB)

#### 한국유전자 데이터베이스 (전누리: GeneNuri)

# Service Items for Bioinformatics in BRIC



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*Biological Research Information Center*