

Proposed NHGRI Reorganization



Eric Green, M.D., Ph.D. Director, NHGRI



- I. Background on NHGRI
- II. Proposed Reorganization
- III. Process for Implementation





Historical Context: 'The Genome Institute'

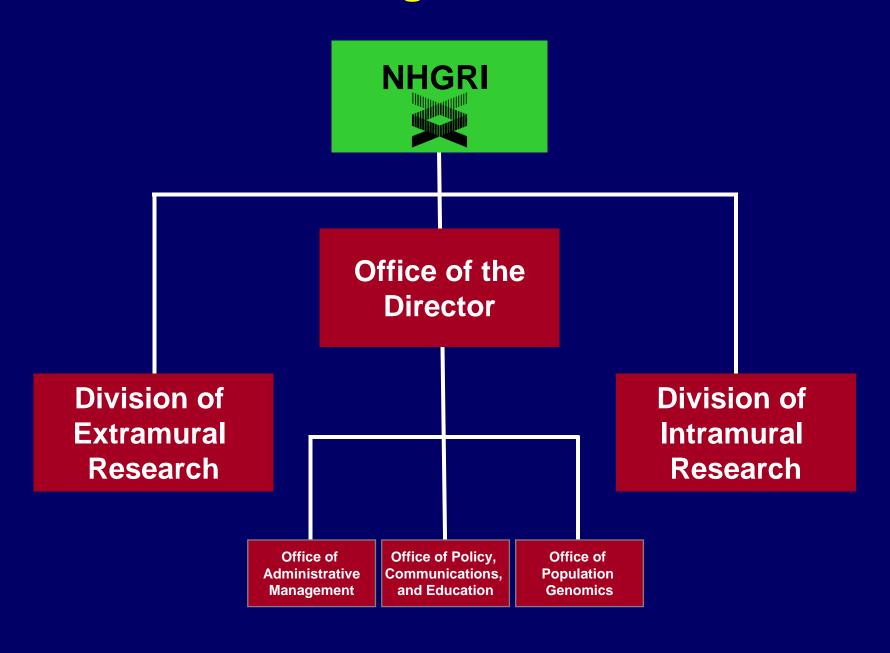


Office for Human Genome Research
1988-1989

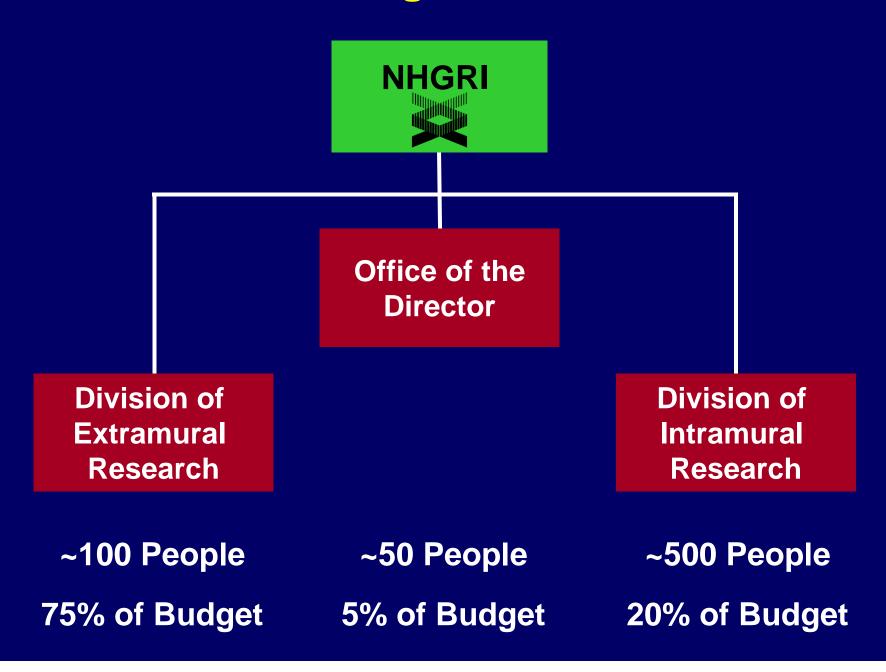
National Center for Human Genome Research 1989-1997

National Human Genome Research Institute 1997-present

Current NHGRI Organizational Structure



Current NHGRI Organizational Structure



Additional Background

- Organizational structure of the Extramural Research Program has been essentially unchanged since the Human Genome Project largely a 'flat' (non-hierarchical) structure
- The Office of the Director has grown in mission, complexity, and scale in recent years, commensurate with the Institute's expanding research portfolio

The Different Eras of NHGRI

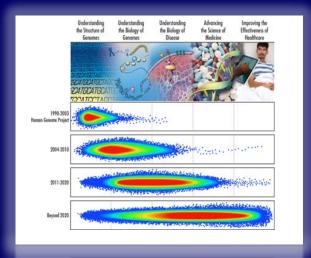
1990-2003

2003-2010

2011-??













~11 Months Ago



PERSPECTIVE

doi:10.1038/nature09764

Charting a course for genomic medicine from base pairs to bedside

Eric D. Green¹, Mark S. Guyer¹ & National Human Genome Research Institute⁴

There has been much progress in genomics in the ten years since a draft sequence of the human genome was published. Opportunities for understanding health and disease are now unprecedented, as advances in genomics are harnessed to obtain robust foundational knowledge about the structure and function of the human genome and about the genetic contributions to human health and disease. Here we articulate a 2011 vision for the future of genomics research and describe the path towards an era of genomic medicine

ince the end of the Human Genome Project (HGP) in 2003 and the quickly. Although genomics has already begun to improve diagnostics publication of a reference human genome sequence¹², genomics has a milestry of a reference human genome sequence¹², genomics has an an interaction of a reference human genome sequence¹², genomics has a milestry of himself and the attended in the second of the se become a mainstay of biomedical research. The scientific commu-(see rollfold). Optimism about the potential contributions of genomics for improving human health has been fuelled by new insights about cancer⁴⁻⁷, the molecular basis of inherited diseases (http://www.ncbi.nlm.nih.gov/ omim and http://www.genome.gov/GWAStudies) and the role of structural variation in disease⁸, some of which have already led to new therapies⁹⁻¹⁵. Other advances have already changed medical practice (for example, microarrays are now used for dinical detection of genomic imbalances and pharmacogenomic testing is routinely performed before administration of certain medications 15). Together, these achievements (see accompanying paper16) document that genomics is contributing to a better understanding nan biology and to improving human health.

As it did eight years ago¹⁷, the National Human Genome Research Institute (NHGRI) has engaged the scientific community (http://www. genome.gov/Planning) to reflect on the key attributes of genomics (Box 1) and explore future directions and challenges for the field. These discussions have led to an updated vision that focuses on understanding human biology and the diagnosis, prevention and treatment of human disease including consideration of the implications of those advances for society (but these discussions, intentionally did not address the role of genomics in agriculture, energy and other areas). Like the HGP, achieving this vision is broader than what any single organization or country can achieve realizing the full benefits of genomics will be a global effort.

This 2011 vision for genomics is organized around five domains extendng from basic research to health applications (Fig. 2). It reflects the view that, over time, the most effective way to improve human health is to understand normal biology (in this case, genome biology) as a basis for understanding disease biology, which then becomes the basis for improving health. At the same time, there are other connections among these domains. Genomics offers opportunities for improving health without a thorough understanding of disease (for example, cancer therapies can be selected based on genomic profiles that identify tumour subtypes 18,19), and clinical discoveries can lead back to understanding disease or even basic biology.

The past decade has seen genomics contribute fundamental knowledge about biology and its perturbation in disease. Further deepening this

effectiveness of healthcare cannot realistically be expected for many years nity's foresight in launching this ambitious project' is evident in the broad (Fig. 2). Achieving such progress will depend not only on research, but range of scientific advances that the HGP has enabled, as shown in Fig. 1 strated the kinds of achievements that can be anticipated with a few examples (Box 2) where a confluence of need and opportunities should lead to major accomplishments in genomic medicine in the coming decade. Similarly, we note three cross-cutting areas that are broadly relevant and fundamental across the entire spectrum of genomics and genomic medicine: bioinformatics and computational biology (Box 3), education and training (Box 4), and genomics and society (Box 5).

Understanding the biology of genomes

Substantial progress in understanding the structure of genomes has revealed much about the complexity of genome biology. Continued acquisition of basic knowledge about genome structure and function will be needed to illuminate further those complexities (Fig. 2). The contri bution of genomics will include more comprehensive sets (catalogues) of data and new research tools, which will enhance the capabilities of all researchers to reveal fundamental principles of biology

Comprehensive catalogues of genomic data

Comprehensive genomic catalogues have been uniquely valuable and widely used. There is a compelling need to improve existing catalogues and to generate new ones, such as complete collections of genetic variation functional genomic elements, RNAs, proteins, and other biological molecules, for both human and model organisms

Genomic studies of the genes and pathways associated with disease related traits require comprehensive catalogues of genetic variation, which provide both genetic markers for association studies and variants for identifying candidate genes. Developing a detailed catalogue of variation in the human genome has been an international effort that began with The SNP Consortium²⁰ and the International HapMap Project²¹ (http://hapmap. nchi.nlm.nih.gov), and is ongoing with the 1000 Genomes Project2 (http://www.1000genomes.org).

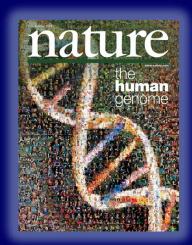
Over the past decade, these catalogues have been critical in the discovery of the specific genes for roughly 3,000 Mendelian (monogenic) disease

understanding will accelerate the transition to genomic medicine (clinical care based on genomic information). But significant change rarely comes

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February 2011 NHGRI Published New Vision for Genomics

The Path to Genomic Medicine

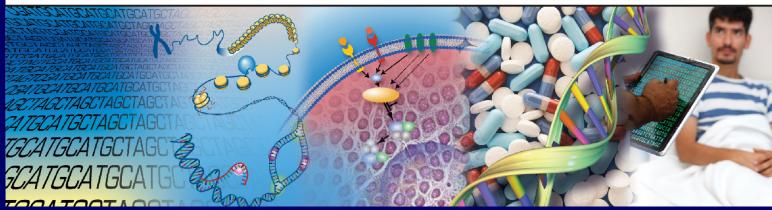


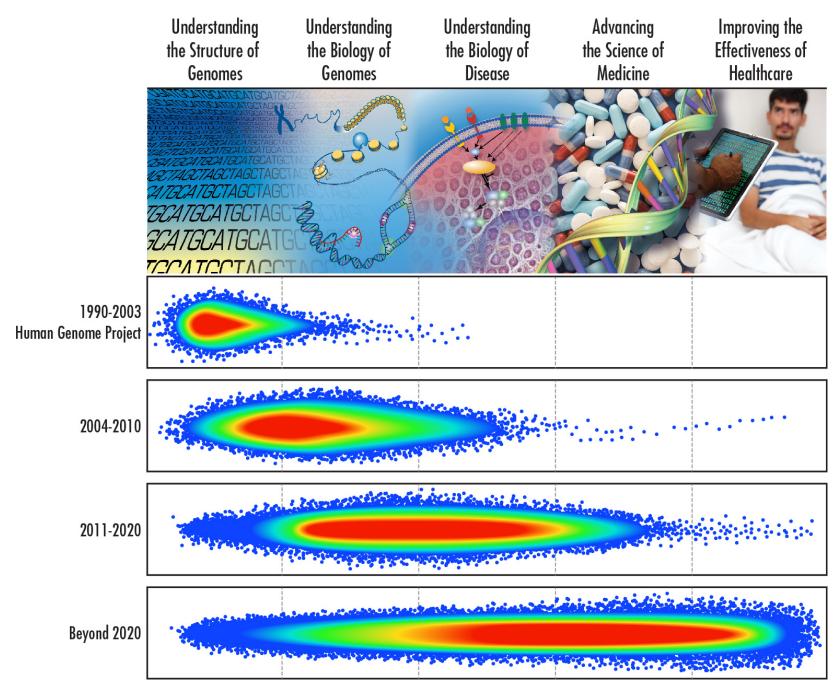


Human Genome Project

Realization of Genomic Medicine

Understanding the Structure of Genomes Understanding the Biology of Genomes Understanding the Biology of Disease Advancing the Science of Medicine Improving the Effectiveness of Healthcare





Green et al. (2011)

NHGRI Extramural Research Program: Circa 'Then'



NHGRI Extramural Research Program: Circa 'Now'

\$1000 Genome

Technology Development Program

TCGA

The Cancer Genome Atlas

KOMP

Knockout Mouse Project

PAGE

Population Architecture using Genomics and Epidemiology

1000 Genomes

ENCODE

Encyclopedia of DNA Elements Project

modENCODE

Model Organism ENCODE

Clinical Sequencing

Mendelian Disorders
Sequencing

Large-Scale Sequencing Program

eMERGE

Electronic Medical Records and Genomics

PhenX

Consensus Measures for Phenotypes and eXposures

GENEVA

Gene Environment Association Studies

CEGS

Centers of Excellence in Genomic Science

FI SI

Ethical Legal Social Implications Program

GARNET

Genomics and Randomized
Trials Network

CEER Program

Centers for Excellence in ELSI Research

KOMP2

KOMP Phenotyping

HMP

Human Microbiome **Project**

GTEx

Genotype-Tissue Expression

H3Africa

Human Heredity and Health in Africa

Protein Capture Reagents

Library of Integrated Networkbased Cellular Signatures

LINCS

MLP

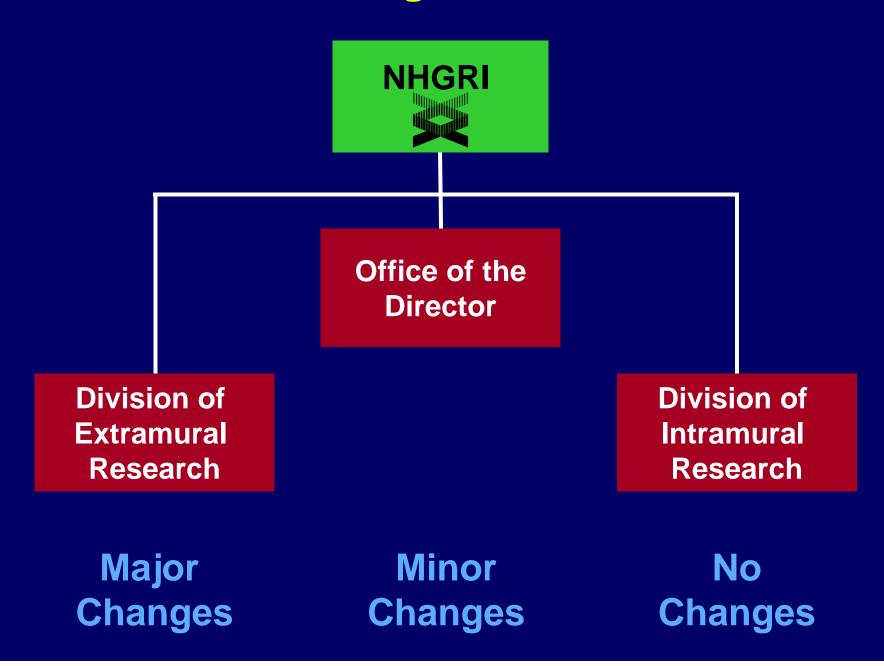
Molecular Libraries
Program

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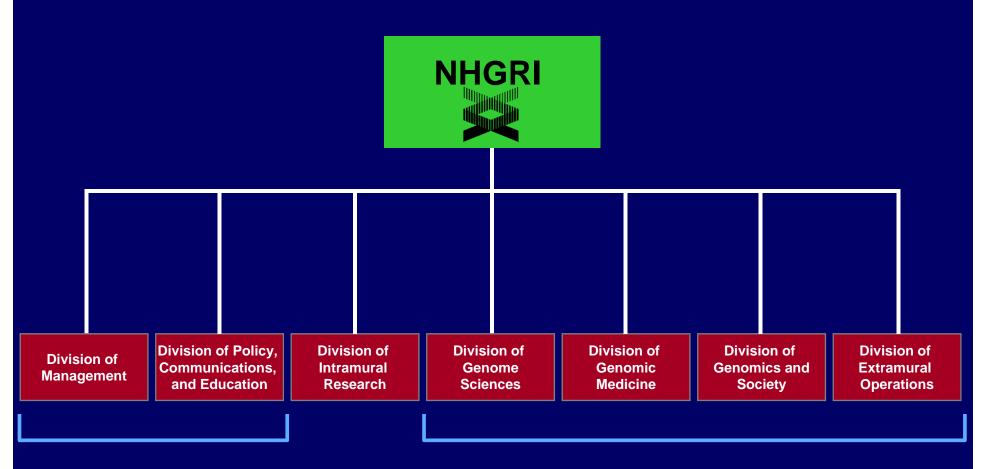




Current NHGRI Organizational Structure

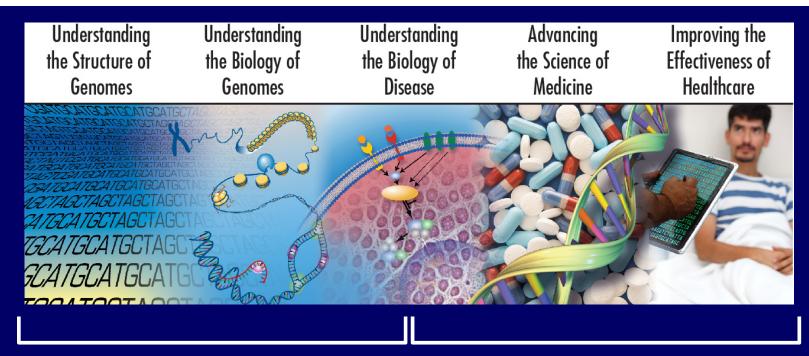


Proposed NHGRI Organizational Structure



From the Office of the Director

Extramural Research Program

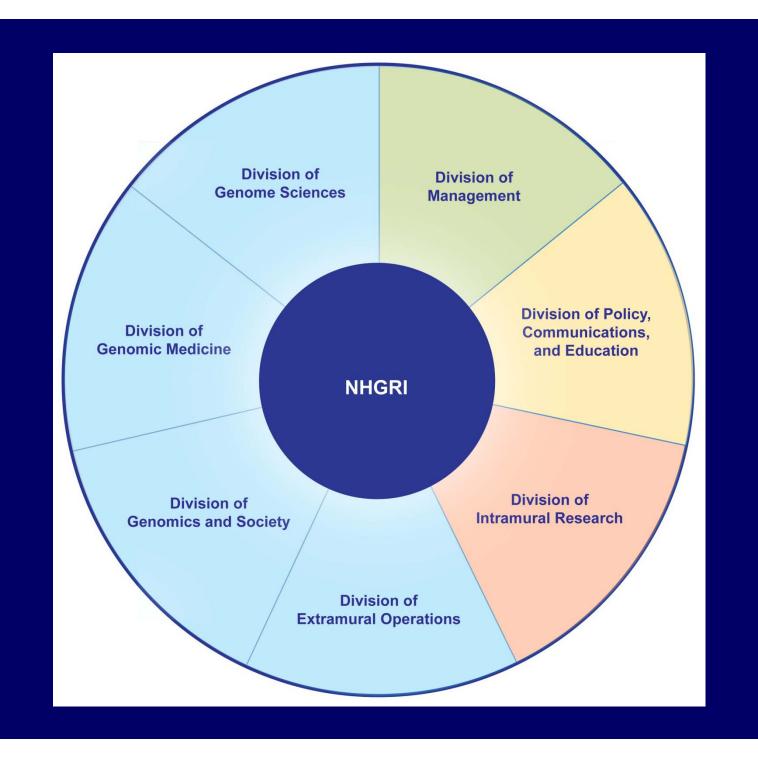


Division of Genome Sciences Division of Genomic Medicine



Genomics & Society

Division of Genomics and Society



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NIH Reform Act of 2006

H. R. 6164

One Hundred Minth Congress of the United States of America

AT THE SECOND SESSION

Begun and held at the City of Washington on Tuesday, the third day of January, two thousand and six

An Act

To amend title IV of the Public Health Service Act to revise and extend the authorities of the National Institutes of Health, and for other purposes.

Be it enacted by the Senate and House of Representatives of the United States of America in Congress assembled,

SECTION 1. SHORT TITLE.

This Act may be cited as the "National Institutes of Health Reform Act of 2006".

TITLE I-NIH REFORM

TITLE I-NIH REFORM

Reform Act of 2006".

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the United States of America in Congress assembly Representatives of



Steps to Reorganization

Public meetings:

Webinar (January 18)
NACHGR Meeting (February 13)

- Submission of reorganization package
- If approved, pursue next steps of appointing Division Directors and implementing new organizational structure

Anticipated Benefits of Reorganizing

- Organizational structure will more effectively align with the Institute's research portfolio (i.e., 'structure will match function')
- New divisions and anticipated substructures will improve succession planning of senior leadership
- New structure commensurate with Director's vision for organizational management

Additional information:

genome.gov/reorg

To provide feedback:

NHGRIcomments@nih.gov

