

Andreas Holzinger
VO 709.049 Medical Informatics
14.10.2015 11:15-12:45

Lecture 01 Introduction

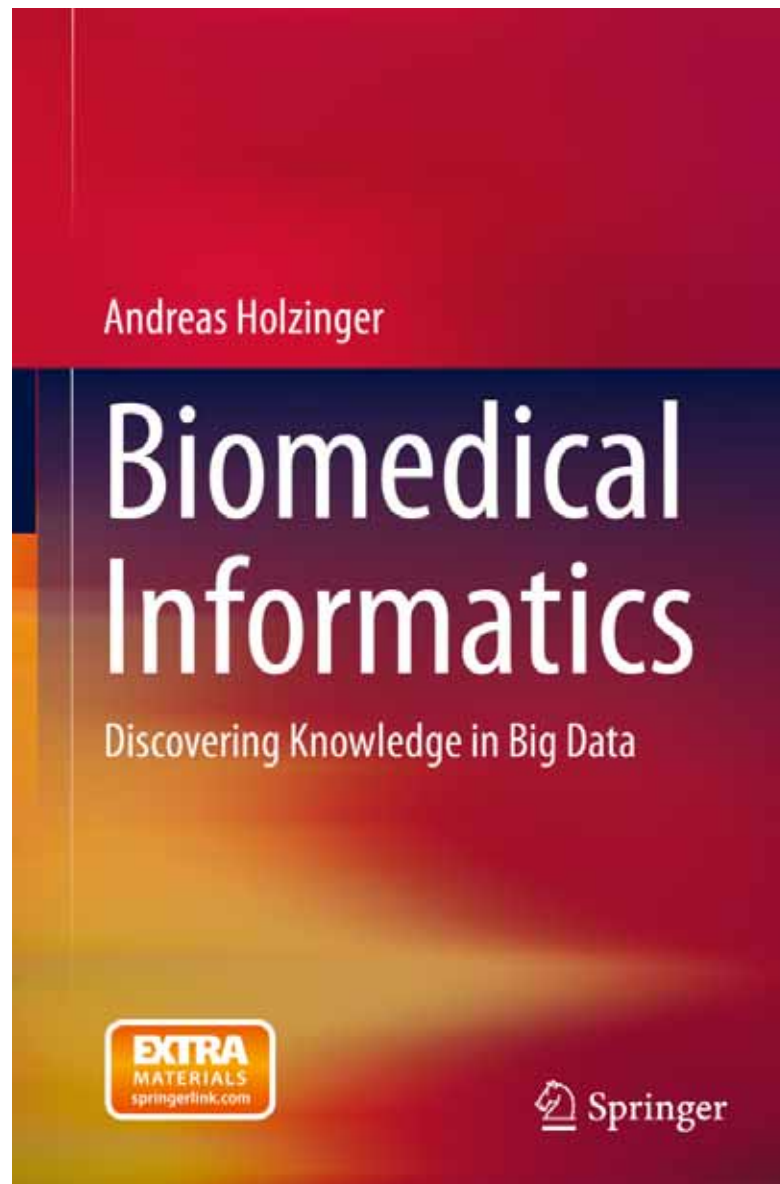
Computer Science meets Life Sciences: Challenges and Future Directions

a.holzinger@tugraz.at

Tutor: markus.plass@student.tugraz.at

<http://hci-kdd.org/biomedical-informatics-big-data>





- **01. Intro: Computer Science meets Life Sciences, challenges, future directions**
- 02. Fundamentals of Data, Information and Knowledge
- 03. Structured Data: Coding, Classification (ICD, SNOMED, MeSH, UMLS)
- 04. Biomedical Databases: Acquisition, Storage, Information Retrieval and Use
- 05. Semi structured , weakly structured data and unstructured information
- 06. Multimedia Data Mining and Knowledge Discovery
- 07. Knowledge and Decision: Cognitive Science & Human-Computer Interaction
- 08. Biomedical Decision Making: Reasoning and Decision Support
- 09. Interactive Information Visualization and Visual Analytics
- 10. Biomedical Information Systems and Medical Knowledge Management
- 11. Biomedical Data: Privacy, Safety and Security
- 12. Methodology for Info Systems: System Design, Usability & Evaluation

- Big Data
- Life
- Proteins – DNA & RNA – Cell – Tissue – Organ – Cardiovascular Systems
- Medicine – Informatics – Computer
- Personalized Medicine
- Translational Informatics – Data Integration
- Open Medical Data
- Biomarker Discovery

- At the end of this first lecture you will ...
- ... be fascinated to see our world in data;
- ... have a basic understanding of the building blocks of life;
- ... be familiar with some differences between Life Sciences and Computer Sciences;
- ... be aware of some possibilities and some limits of Biomedical Informatics;
- ... have some ideas of some future directions of Biomedical Informatics;

- **Bioinformatics** = discipline, as part of biomedical informatics, at the interface between *biology* and *information science* and *mathematics*; processing of biological data;
- **Biomarker** = a characteristic (e.g. body-temperature (fever) as a biomarker for an infection, or proteins measured in the urine) as an indicator for normal or pathogenic biological processes, or pharmacologic responses to a therapeutic intervention;
- **Biomedical data** = compared with general data, it is characterized by large volumes, complex structures, high dimensionality, evolving biological concepts, and insufficient data modeling practices;
- **Biomedical Informatics** = 2011-definition: similar to medical informatics but including the optimal use of biomedical data, e.g. from genomics, proteomics, metabolomics;
- **Classical Medicine** = is both the science and the art of healing and encompasses a variety of practices to maintain and restore health;
- **Genomics** = branch of molecular biology which is concerned with the structure, function, mapping & evolution of genomes;
- **Medical Informatics** = 1970-definition: "... scientific field that deals with the storage, retrieval, and optimal use of medical information, data, and knowledge for problem solving and decision making";
- **Metabolomics** = study of chemical processes involving metabolites (e.g. enzymes). A challenge is to integrate proteomic, transcriptomic, and metabolomic information to provide a more complete understanding of living organisms;
- **Molecular Medicine** = emphasizes cellular and molecular phenomena and interventions rather than the previous conceptual and observational focus on patients and their organs;

- **Omics data** = data from e.g. genomics, proteomics, metabolomics, etc.
- **Pervasive Computing** = similar to ubiquitous computing (Ubicomp), a post-desktop model of Human-Computer Interaction (HCI) in which information processing is integrated into every-day, miniaturized and embedded objects and activities; having some degree of “intelligence”;
- **Pervasive Health** = all unobtrusive, analytical, diagnostic, supportive etc. information functions to improve health care, e.g. remote, automated patient monitoring, diagnosis, home care, self-care, independent living, etc.;
- **Proteome** = the entire complement of proteins that is expressed by a cell, tissue, or organism;
- **Proteomics** = field of molecular biology concerned with determining the proteome;
- **P-Health Model** = Preventive, Participatory, Pre-emptive, Personalized, Predictive, Pervasive (= available to anybody, anytime, anywhere);
- **Space** = a set with some added structure;
- **Technological Performance** = machine “capabilities”, e.g. short response time, high throughput, high availability, etc.
- **Time** = a dimension in which events can be ordered along a time line from the past through the present into the future;
- **Translational Medicine** = based on interventional epidemiology; progress of Evidence-Based Medicine (EBM), integrates research from basic science for patient care and prevention;
- **Von-Neumann-Computer** = a 1945 architecture, which still is the predominant machine architecture of today (opp.: Non-Vons, incl. analogue, optical, quantum computers, cell processors, DNA and neural nets (in silico));

- AI = Artificial Intelligence
- AL = Artificial Life
- CPG = Clinical Practice Guideline
- CPOE = Computerized physician order entry
- CMV = Controlled Medical Vocabulary
- DEC = Digital Equipment Corporation (1957-1998)
- DNA = Deoxyribonucleic Acid
- EBM = Evidence Based Medicine
- EPR = Electronic Patient Record
- GBM = Genome Based Medicine
- GC = Gas Chromatography
- GPM = Genetic Polymorphism
- HCI = Human–Computer Interaction
- LC = Liquid Chromatography
- LNCS = Lecture Notes in Computer Science
- MS = Mass Spectrometry
- mRNA = Messenger RNA
- NGC = New General Catalogue of Nebulae and Star clusters in Astronomy
- NGS = Next Generation Sequencing
- NMR = Nuclear Magnetic Resonance
- PDB = Protein Data Base
- PDP = Programmable Data Processor (mainframe)
- PPI = Protein-Protein Interaction
- RFID = Radio-frequency identification device
- RNA = Ribonucleic Acid
- SNP = Single Nucleotide Polymorphism
- TNF = Tumor Necrosis Factor
- TQM = Total Quality Management

- **Zillions** of different biological species (humans, animals, bacteria, virus, plants, ...);
- Enormous **complexity** of the medical domain [1];
- **Complex**, heterogeneous, high-dimensional, big data in the life sciences [2];
- Limited **time**, e.g. a medical doctor in a public hospital has only 5 min. to make a decision [3];
- Limited **computational power** in comparison to the complexity of life (and the natural limitations of the Von-Neumann architecture, ...);

1. Patel VL, Kahol K, & Buchman T (2011) Biomedical Complexity and Error. *J. Biomed. Inform.* 44(3):387-389.
2. Holzinger A, Dehmer M, & Jurisica I (2014) Knowledge Discovery and interactive Data Mining in Bioinformatics - State-of-the-Art, future challenges and research directions. *BMC Bioinformatics* 15(S6):I1.
3. Gigerenzer G (2008) *Gut Feelings: Short Cuts to Better Decision Making* (Penguin, London).

What is the challenge ?

ESO, Atacama, Chile (2011)

Time

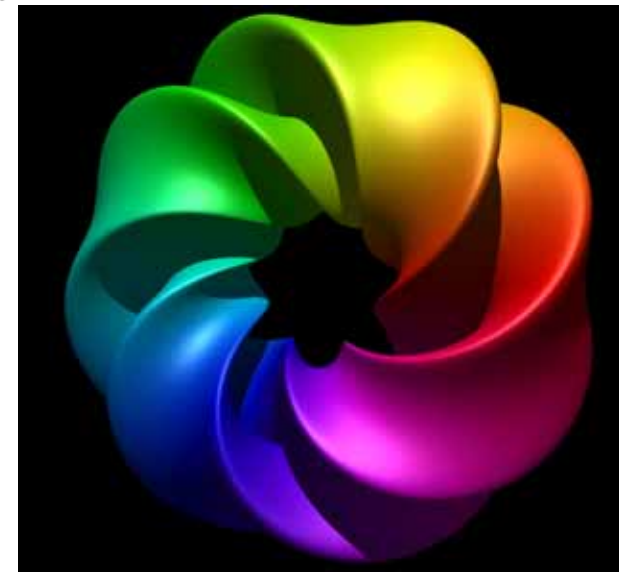
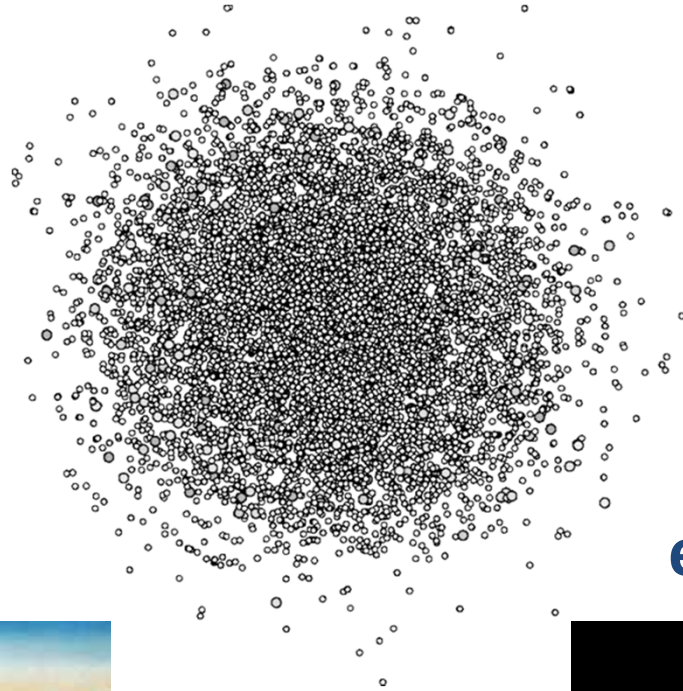
e.g. Entropy



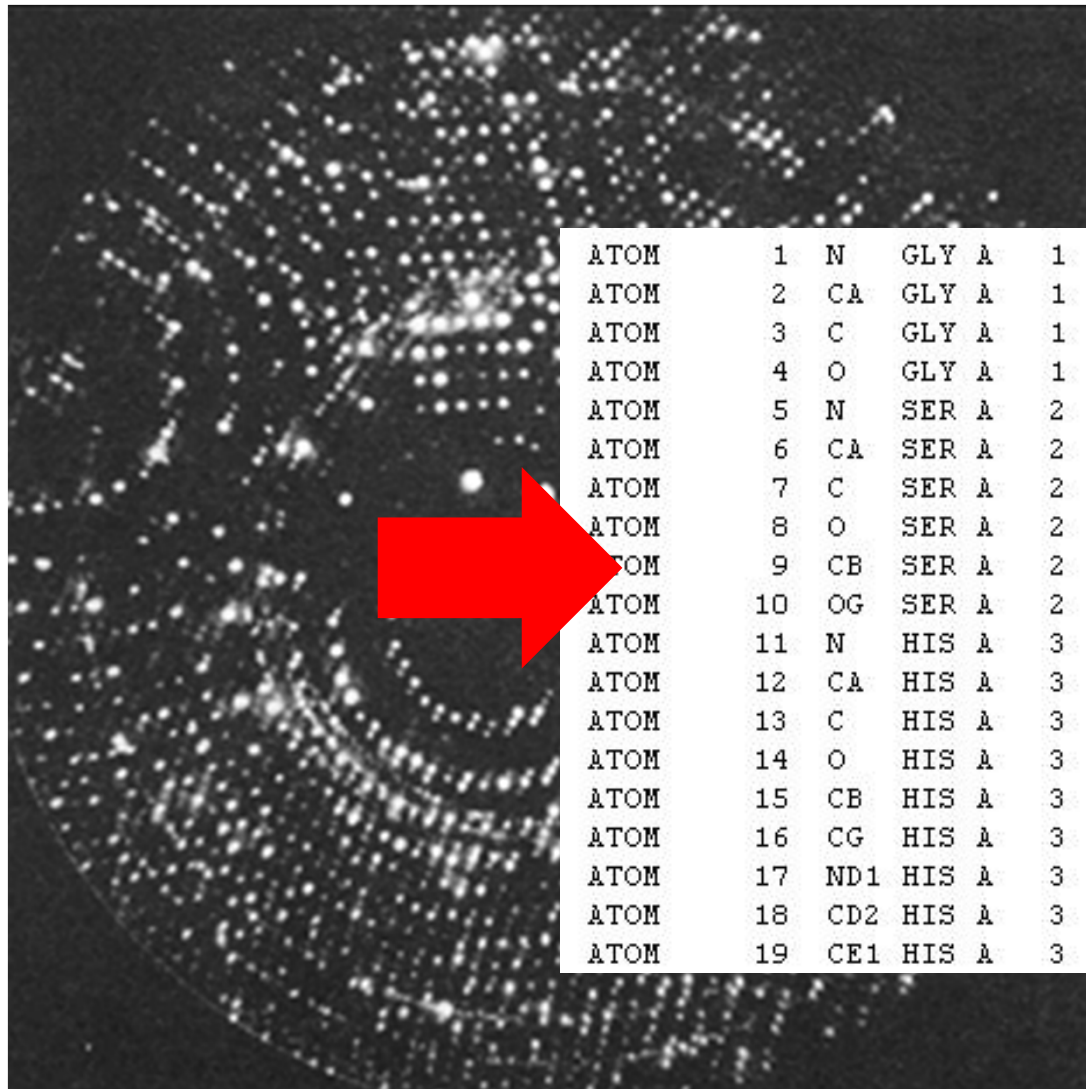
Dali, S. (1931) The persistence of memory

Space

e.g. Topology

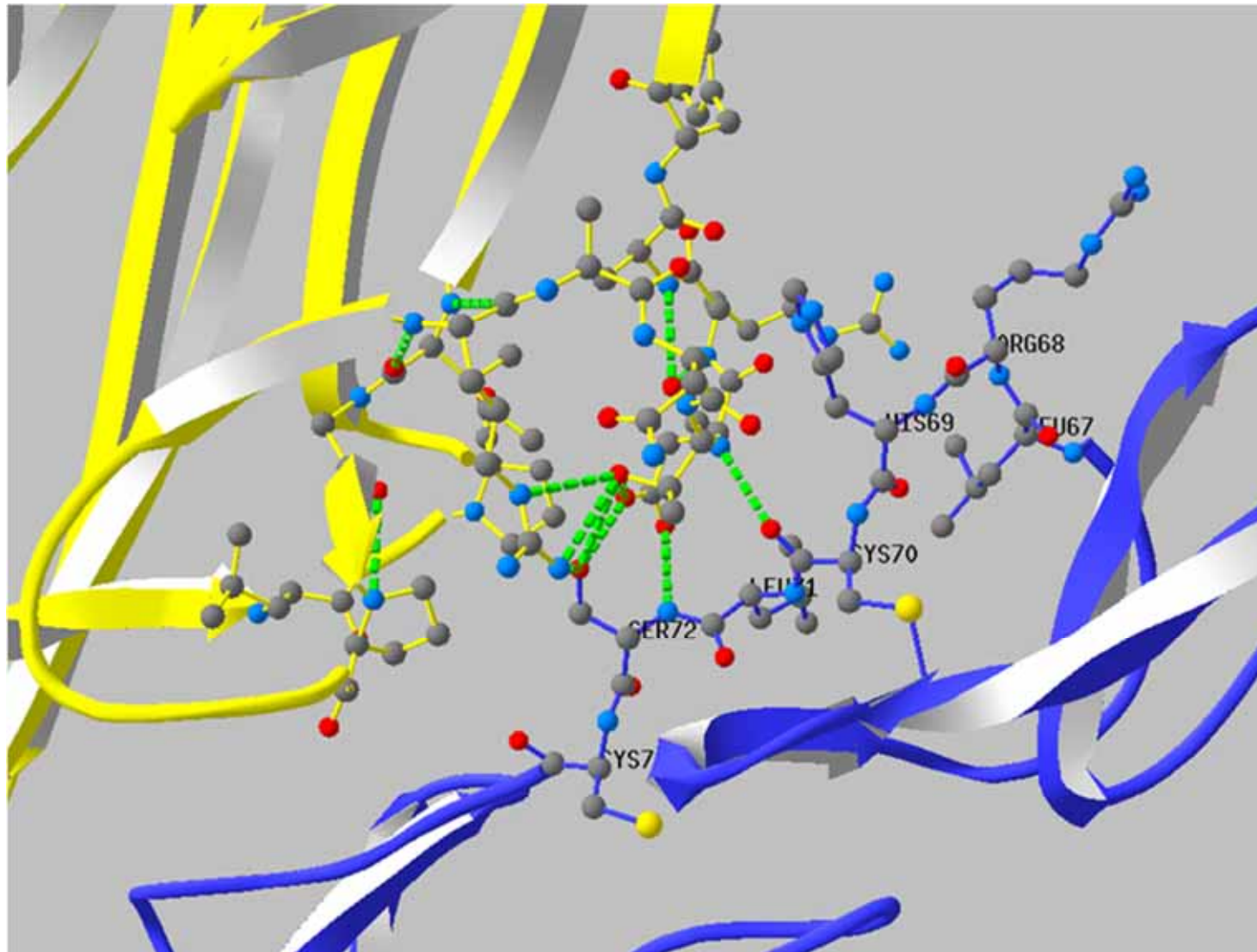


Bagula & Bourke (2012) Klein-Bottle



ATOM	1	N	GLY	A	1	44.842	51.034	101.284	0.01	27.20
ATOM	2	CA	GLY	A	1	45.640	50.230	100.389	0.01	26.99
ATOM	3	C	GLY	A	1	46.692	49.648	101.308	0.01	26.80
ATOM	4	O	GLY	A	1	46.895	50.222	102.381	0.01	26.91
ATOM	5	N	SER	A	2	47.283	48.516	100.951	1.00	26.26
ATOM	6	CA	SER	A	2	48.277	47.866	101.761	1.00	26.17
ATOM	7	C	SER	A	2	49.212	47.031	100.845	1.00	24.21
ATOM	8	O	SER	A	2	49.060	47.195	99.630	1.00	19.77
ATOM	9	CB	SER	A	2	47.438	47.091	102.800	1.00	26.31
ATOM	10	OG	SER	A	2	46.276	46.356	102.404	1.00	27.99
ATOM	11	N	HIS	A	3	50.147	46.186	101.370	1.00	23.93
ATOM	12	CA	HIS	A	3	51.129	45.389	100.609	1.00	21.44
ATOM	13	C	HIS	A	3	50.953	43.905	100.849	1.00	20.32
ATOM	14	O	HIS	A	3	50.530	43.595	101.950	1.00	22.00
ATOM	15	CB	HIS	A	3	52.555	45.674	100.990	1.00	19.69
ATOM	16	CG	HIS	A	3	52.940	47.090	100.611	1.00	21.44
ATOM	17	ND1	HIS	A	3	53.371	47.470	99.422	1.00	20.87
ATOM	18	CD2	HIS	A	3	52.956	48.175	101.433	1.00	21.69
ATOM	19	CE1	HIS	A	3	53.676	48.730	99.476	1.00	20.57

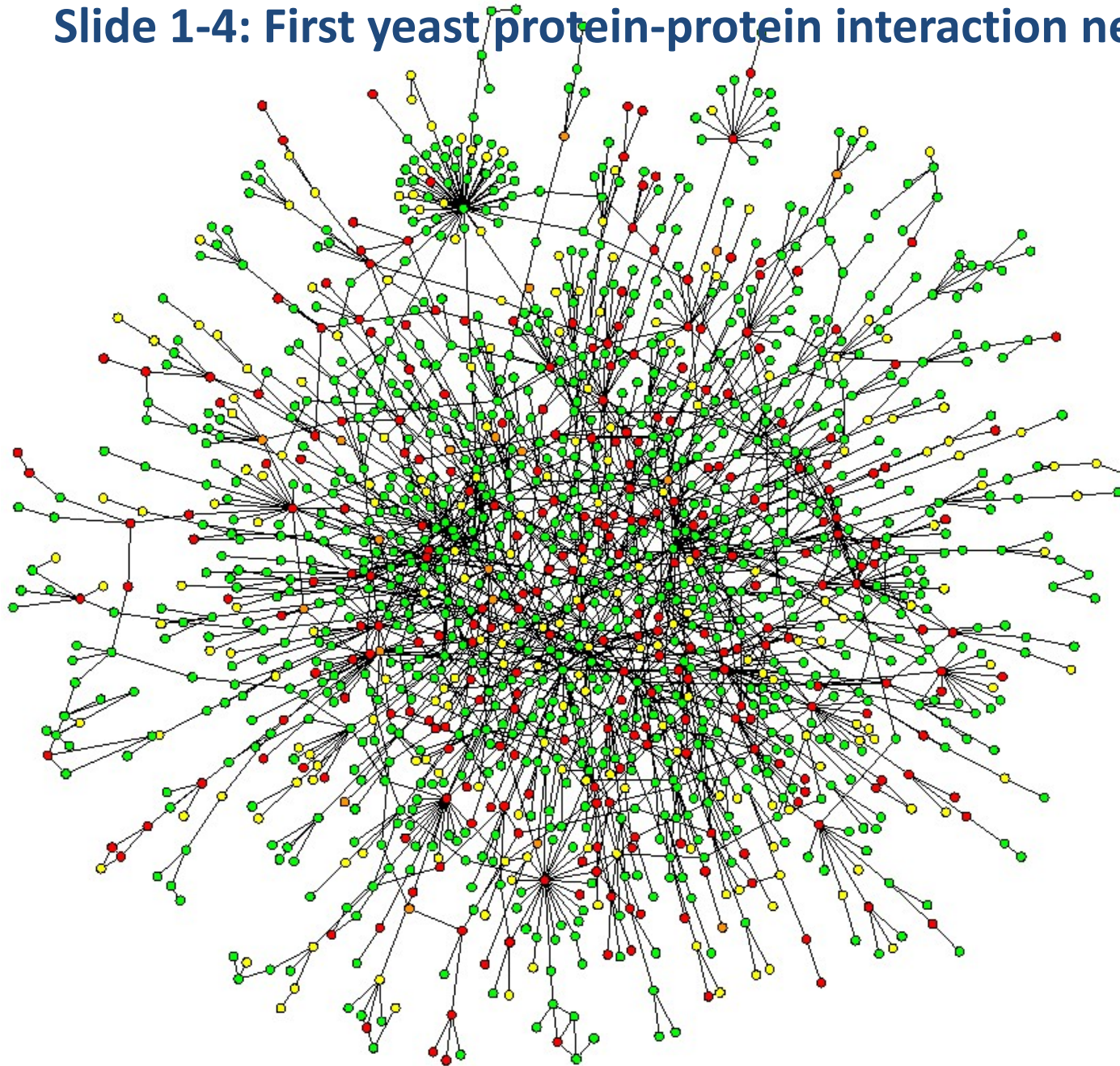
Wiltgen, M. & Holzinger, A. (2005) Visualization in Bioinformatics: Protein Structures with Physicochemical and Biological Annotations. In: *Central European Multimedia and Virtual Reality Conference. Prague, Czech Technical University (CTU)*, 69-74



Wiltgen, M., Holzinger, A. & Tilz, G. P. (2007) Interactive Analysis and Visualization of Macromolecular Interfaces Between Proteins. In: *Lecture Notes in Computer Science (LNCS 4799)*. Berlin, Heidelberg, New York, Springer, 199-212.

Slide 1-4: First yeast protein-protein interaction network

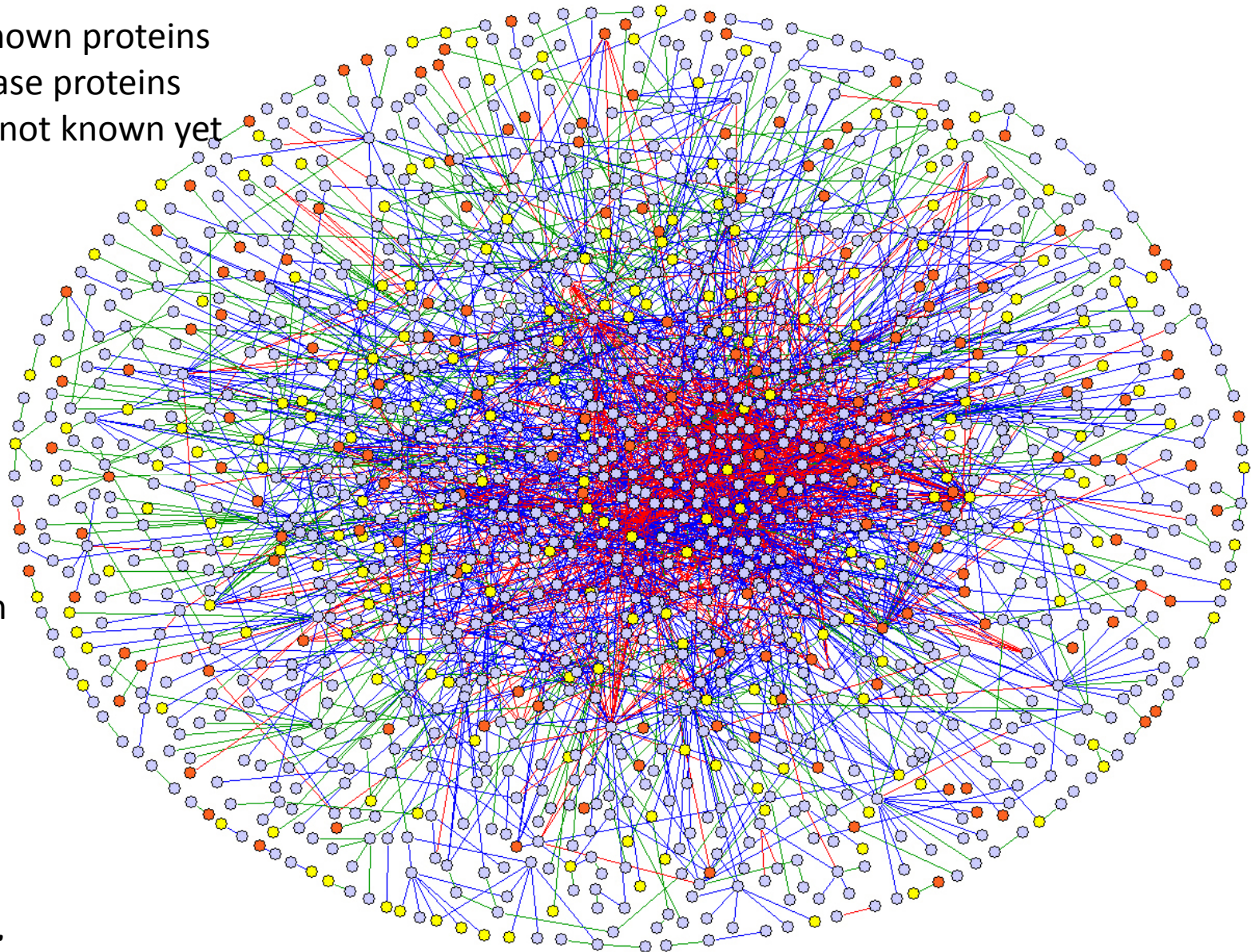
Nodes = proteins
Links = physical interactions
(bindings)
Red Nodes = lethal
Green Nodes = non-lethal
Orange = slow growth
Yellow = not known



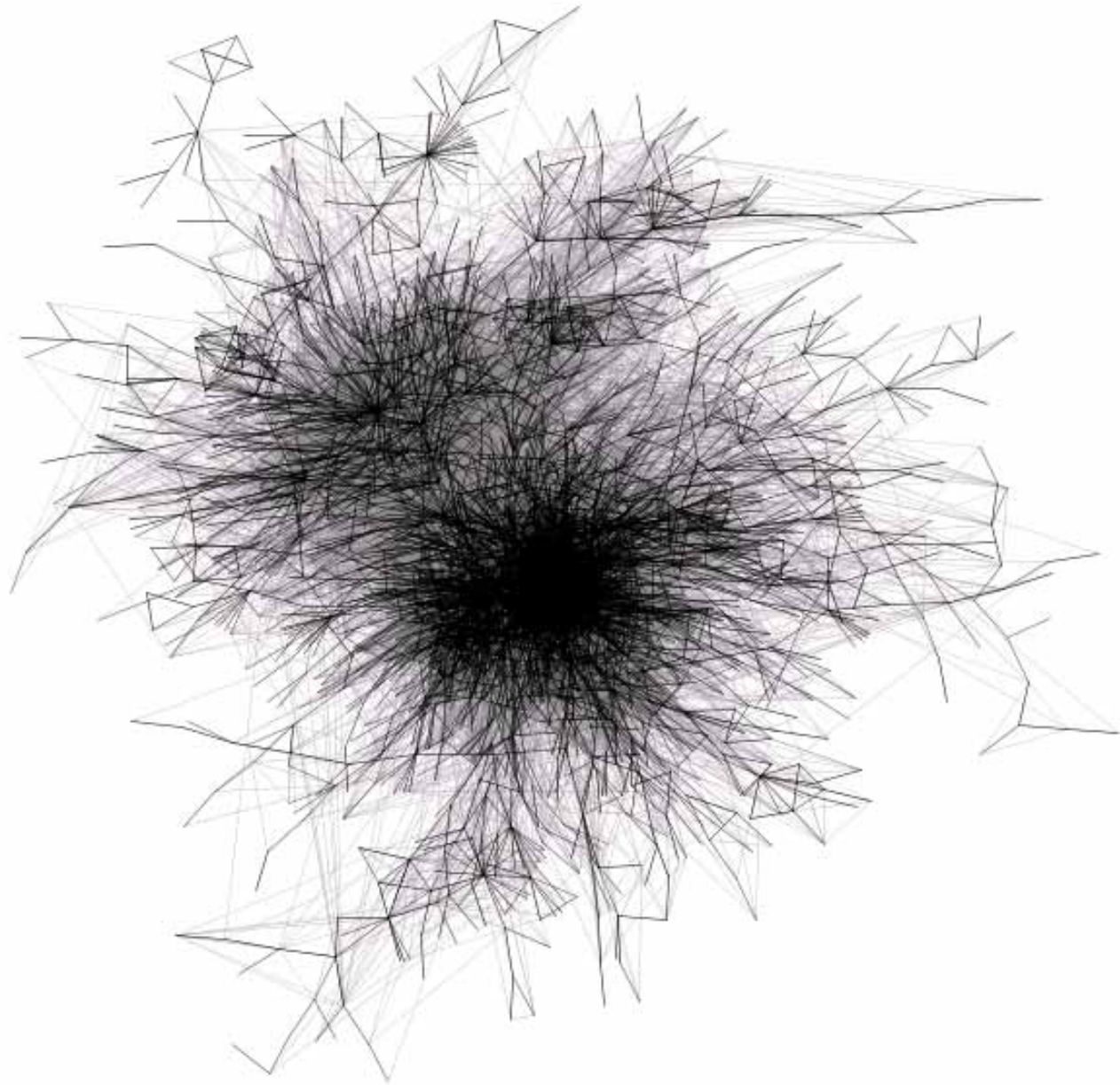
Jeong, H., Mason, S. P., Barabasi, A. L. & Oltvai, Z. N. (2001) Lethality and centrality in protein networks. *Nature*, 411, 6833, 41-42.

Slide 1-5: First human protein-protein interaction network

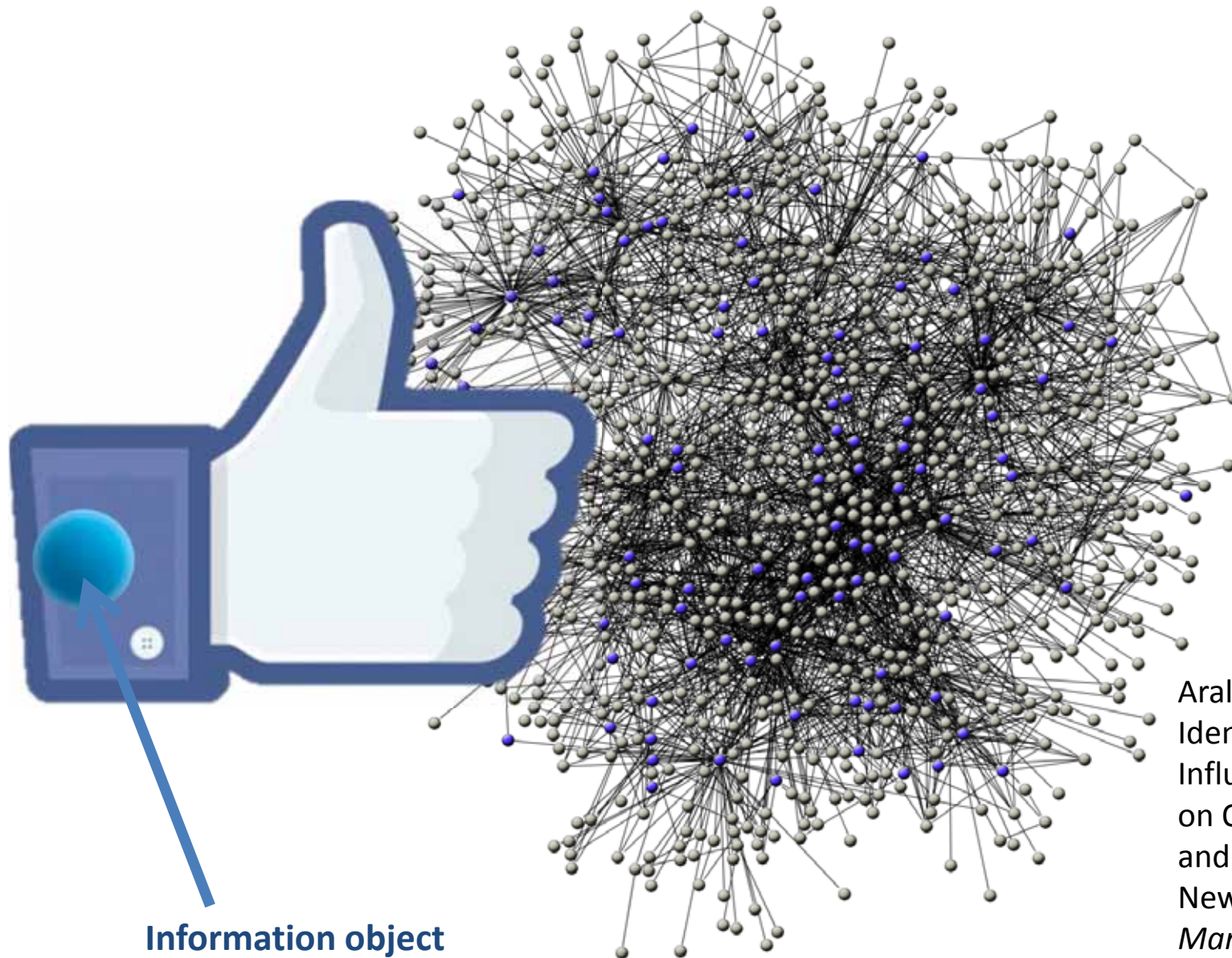
Light blue = known proteins
Orange = disease proteins
Yellow ones = not known yet



Stelzl, U. et al.
(2005) A Human
Protein-Protein
Interaction
Network: A
Resource for
Annotating the
Proteome. *Cell*,
122, 6, 957-968.

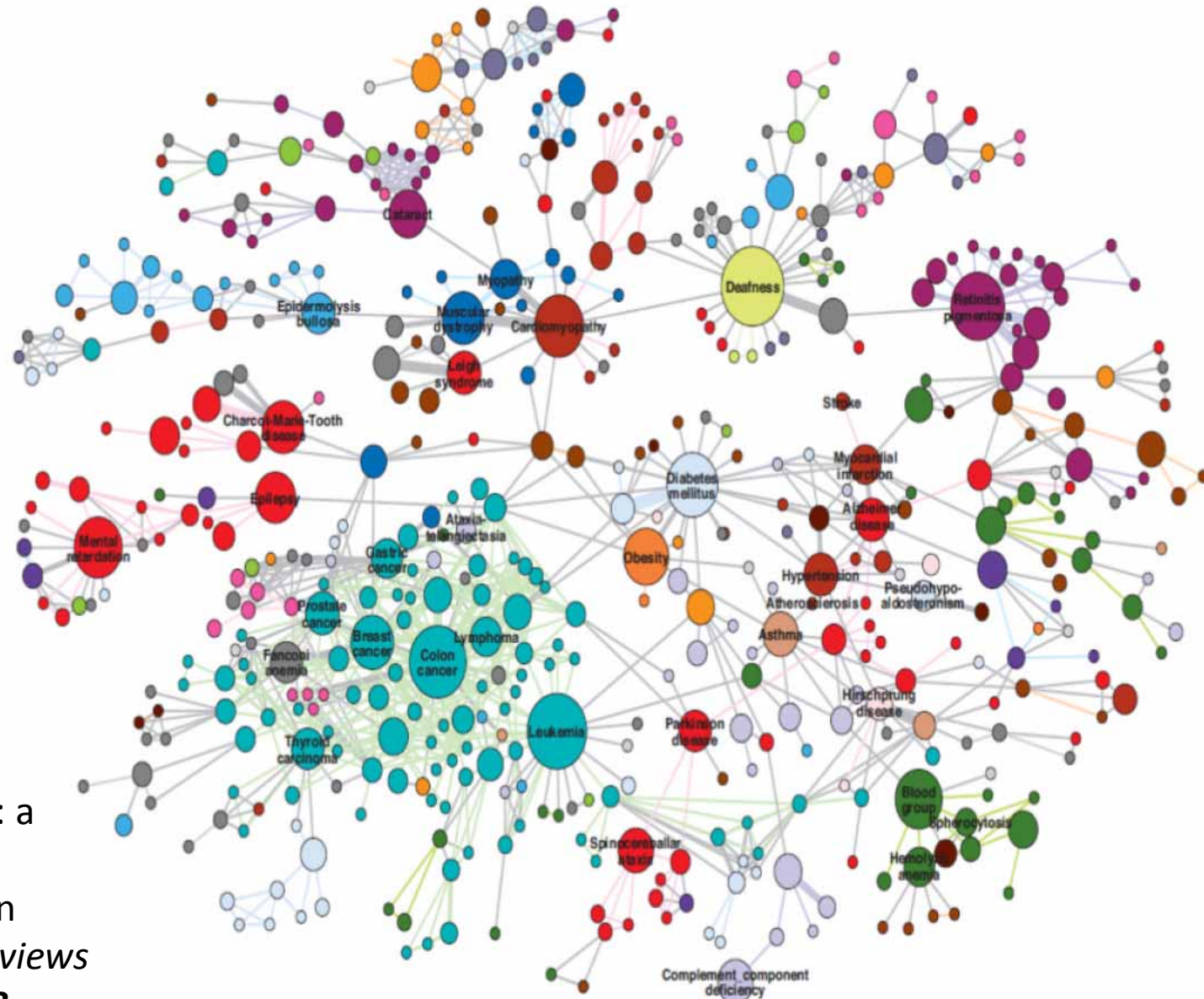


Hurst, M. (2007), Data Mining: Text Mining, Visualization and Social Media. Online available: http://datamining.typepad.com/data_mining/2007/01/the_blogosphere.html, last access: 2011-09-24



Information object

Aral, S. (2011)
Identifying Social
Influence: A Comment
on Opinion Leadership
and Social Contagion in
New Product Diffusion.
Marketing Science, 30,
2, 217-223.



Barabási, A. L.,
Gulbahce, N. &
Loscalzo, J. 2011.
Network medicine: a
network-based
approach to human
disease. *Nature Reviews
Genetics*, 12, **56-68**.



$$\left(-\frac{\hbar^2}{2m} \Delta + U(\vec{r}, t) \right) \psi(\vec{r}, t) = i\hbar \frac{\partial}{\partial t} \psi(\vec{r}, t)$$

to reproduce ...

to grow ...

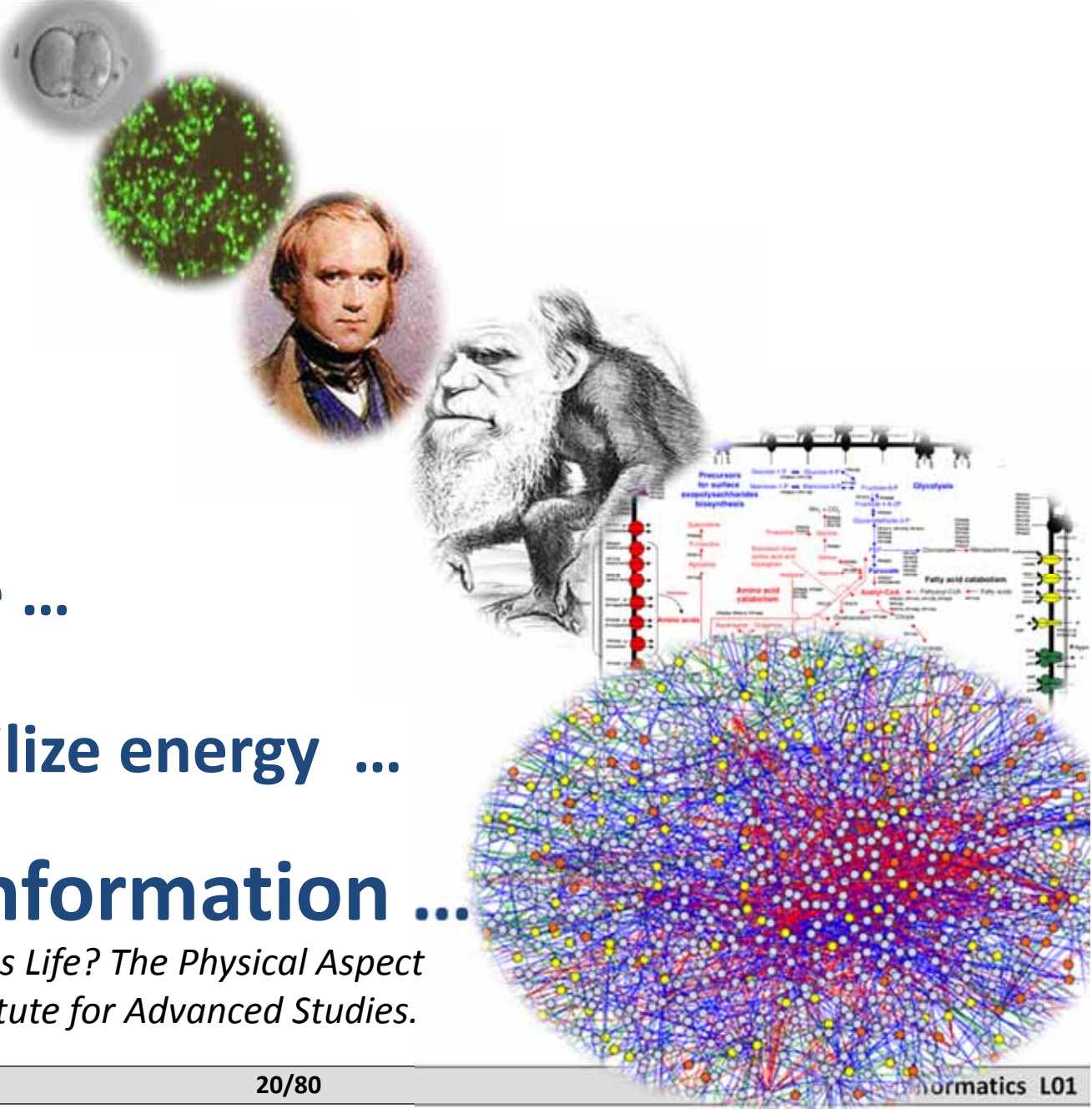
to evolve ...

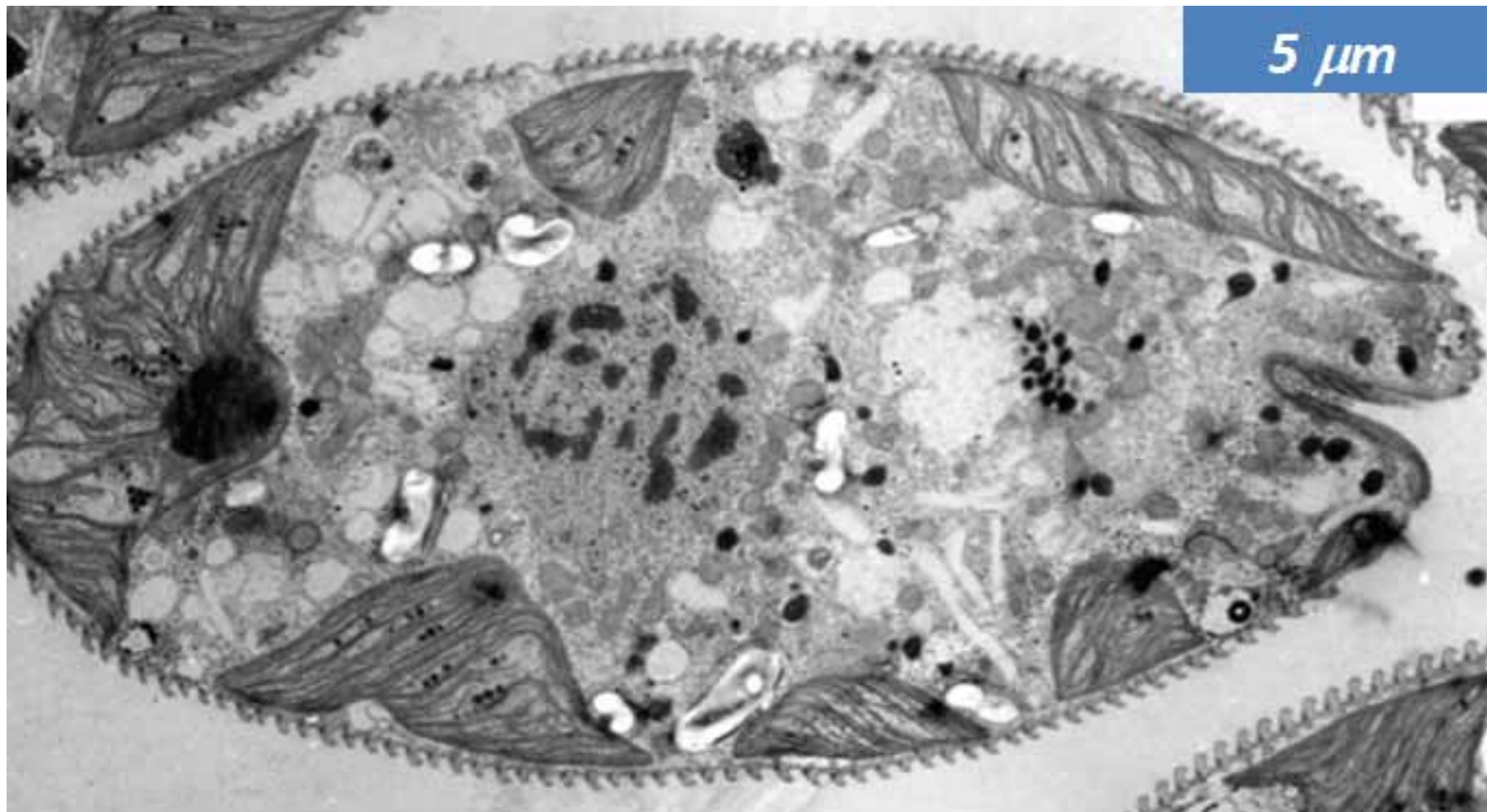
to self-replicate ...

to generate/utilize energy ...

to process information ...

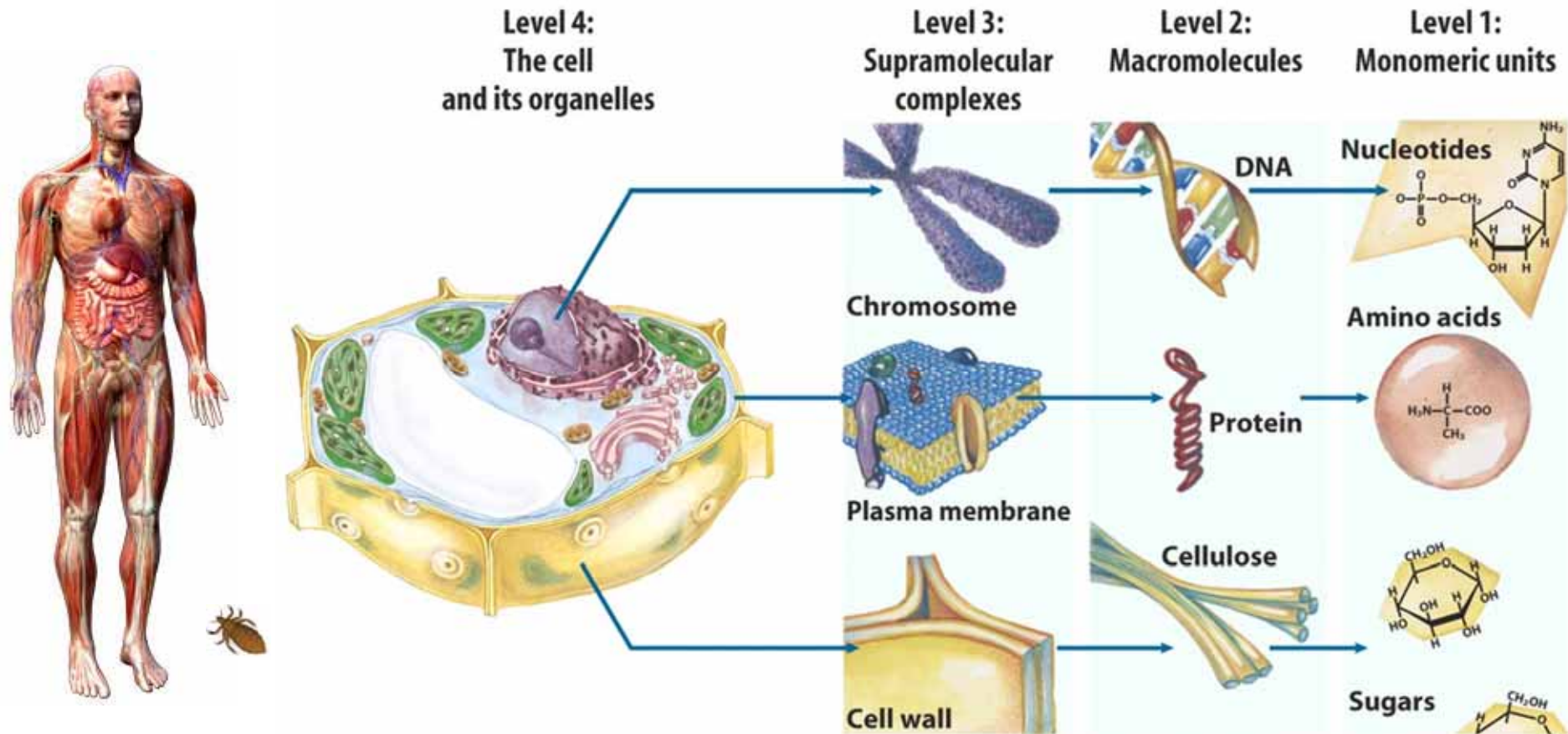
Schrödinger, E. (1944) *What Is Life? The Physical Aspect of the Living Cell*. Dublin Institute for Advanced Studies.



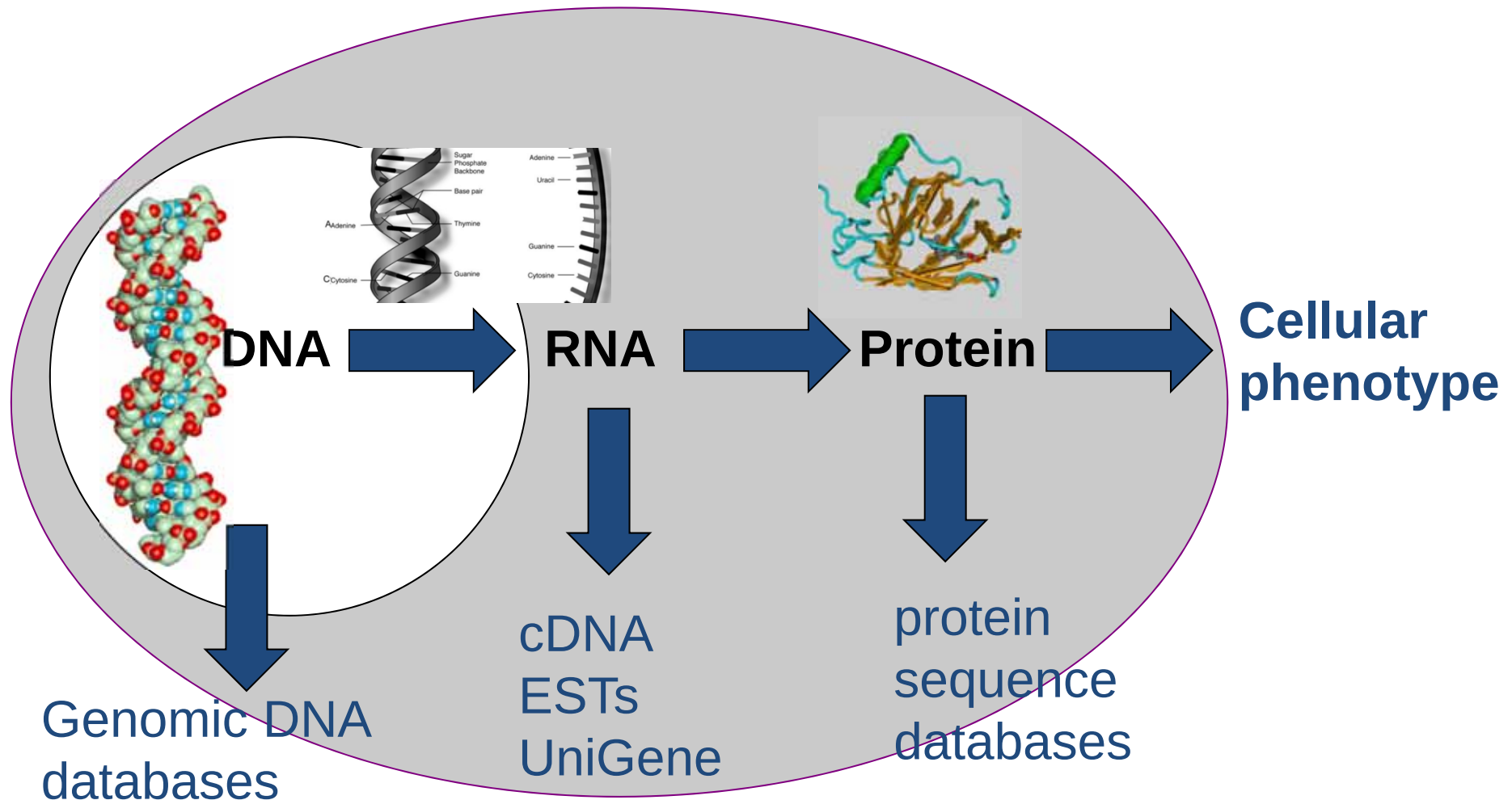


Lane, N. & Martin, W. (2010) The energetics of genome complexity.
Nature, 467, 7318, 929-934.

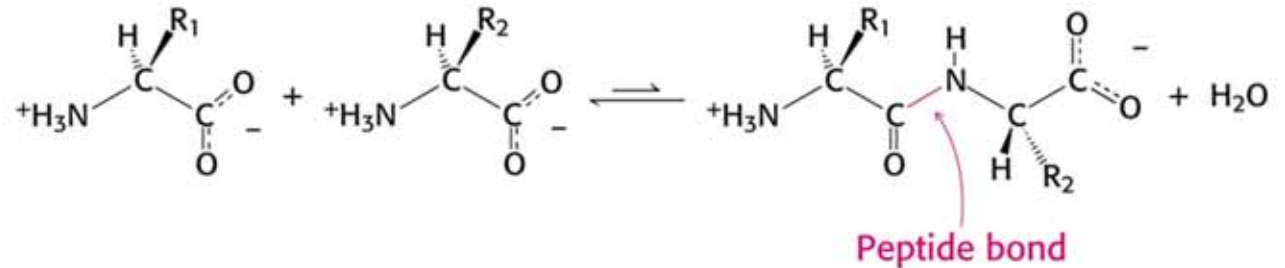
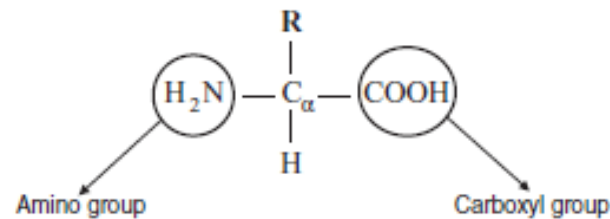
Slide 1-11 Building Blocks of Life - Overview



Human eye		Light microscope	Electron microscope	Special
1m	1mm	1 μm	1nm	100 pm



Crick, F. 1970. Central Dogma of Molecular Biology. *Nature*, 227, (5258), 561-563.

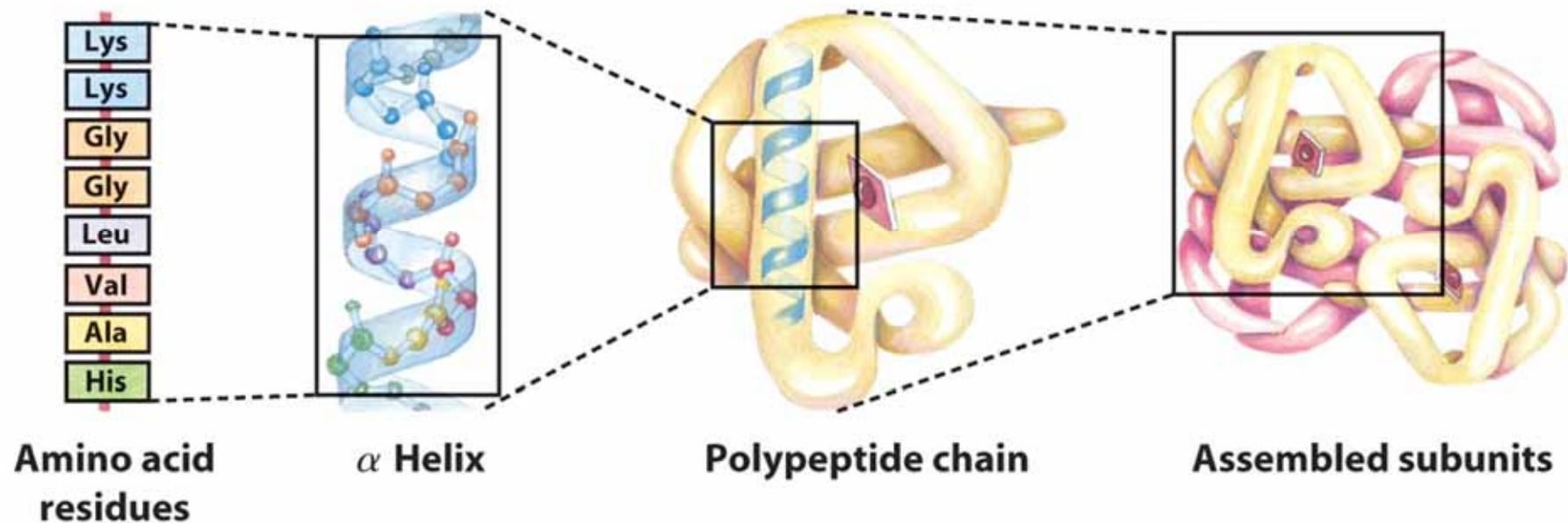


**Primary
structure**

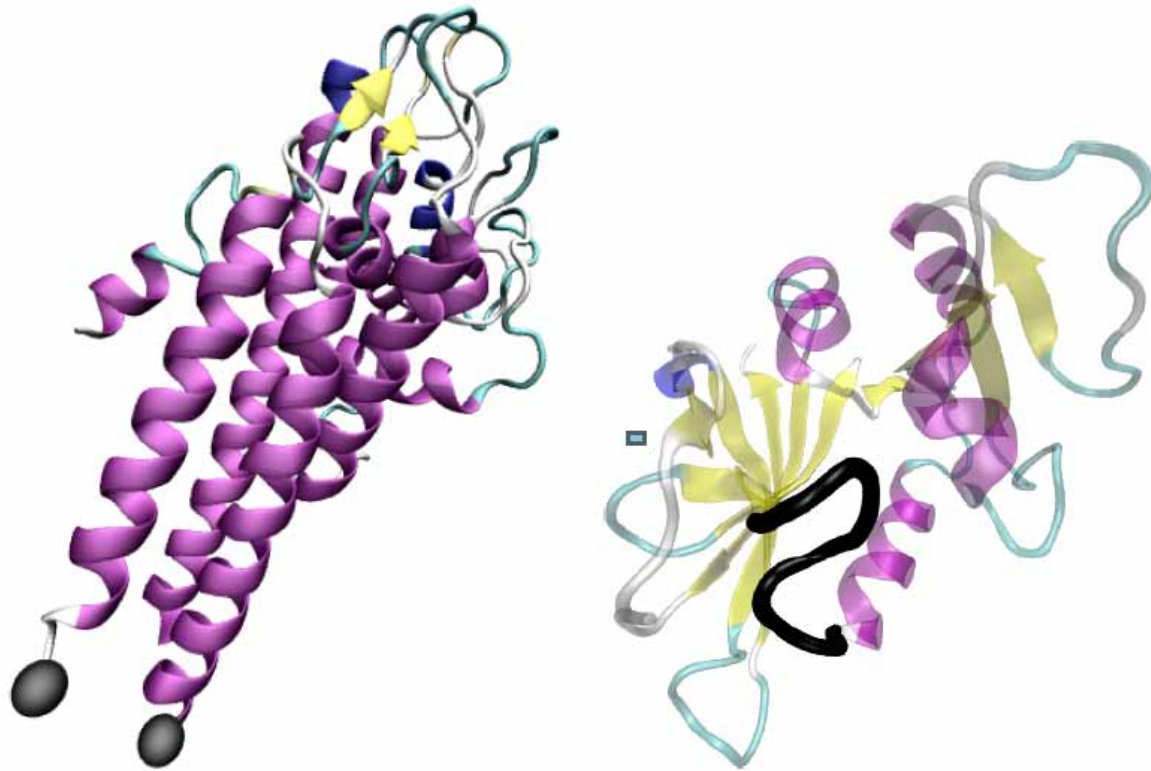
**Secondary
structure**

**Tertiary
structure**

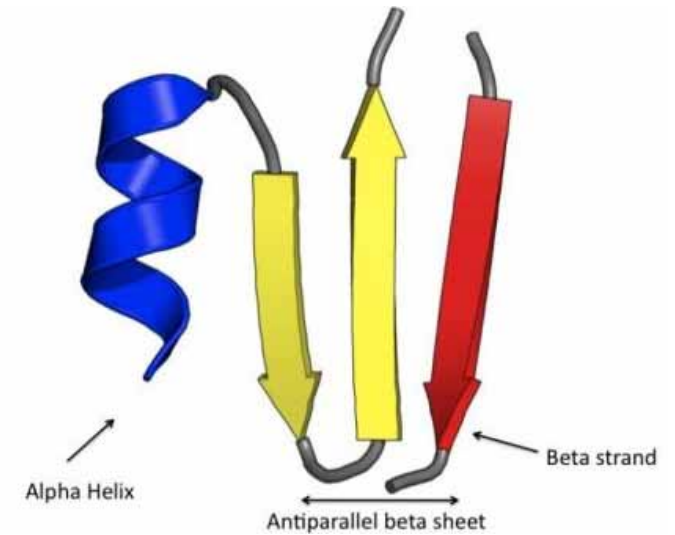
**Quaternary
structure**



Gromiha, M. 2010. *Protein Bioinformatics*, Amsterdam, Elsevier.



Shehu, A. & Kavraki, L. E. 2012. Modeling structures and motions of loops in protein molecules. *Entropy*, 14, (2), 252-290.



Tertiary structure

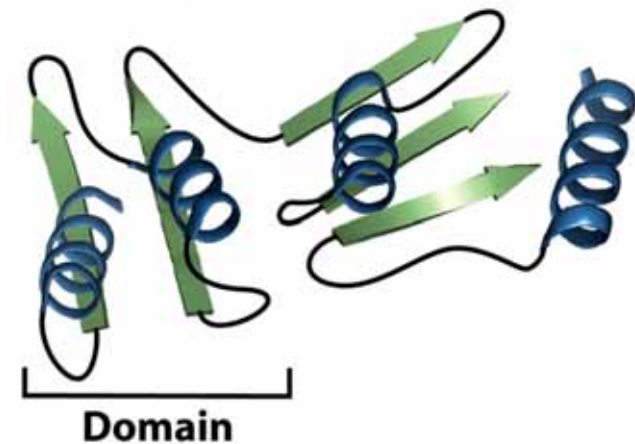
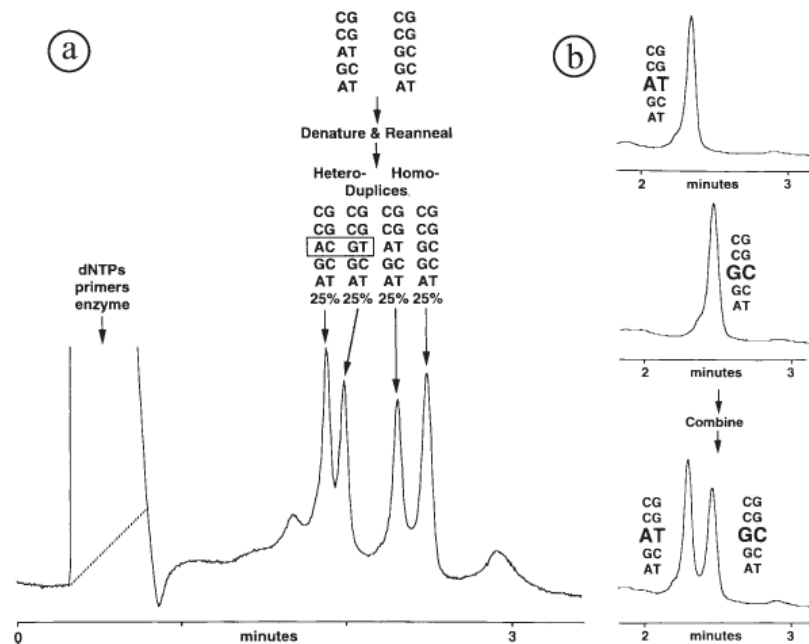
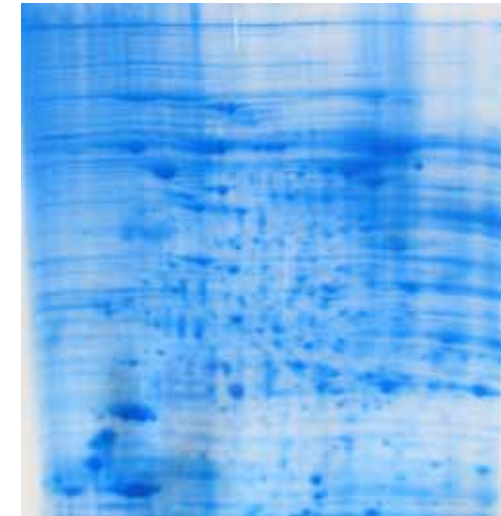
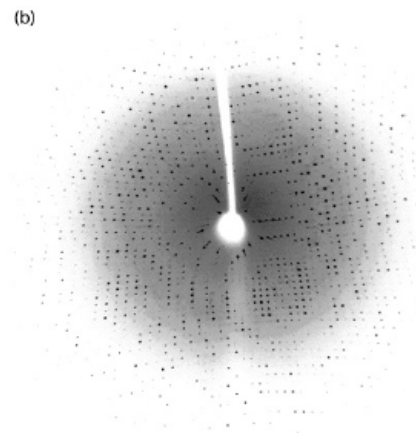
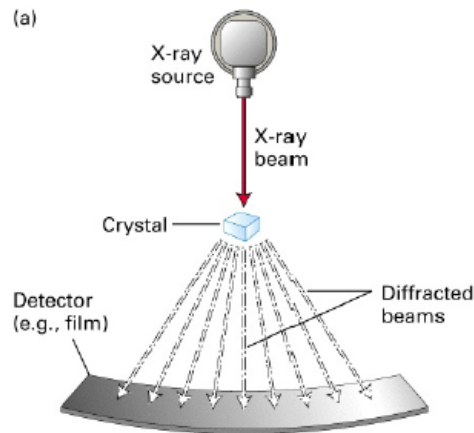


Figure 3-3a
Molecular Cell Biology, Sixth Edition
© 2008 W. H. Freeman and Company



Rabilloud, et al. 2010. Two-dimensional gel electrophoresis in proteomics: past, present and future. *Journal of proteomics*, 73, (11), 2064-2077.

Xiao, W. Z. & Oefner, P. J. 2001. Denaturing high-performance liquid chromatography: A review. *Human Mutation*, 17, (6), 439-474.

Slide 1-16: Comparison of some current Methods

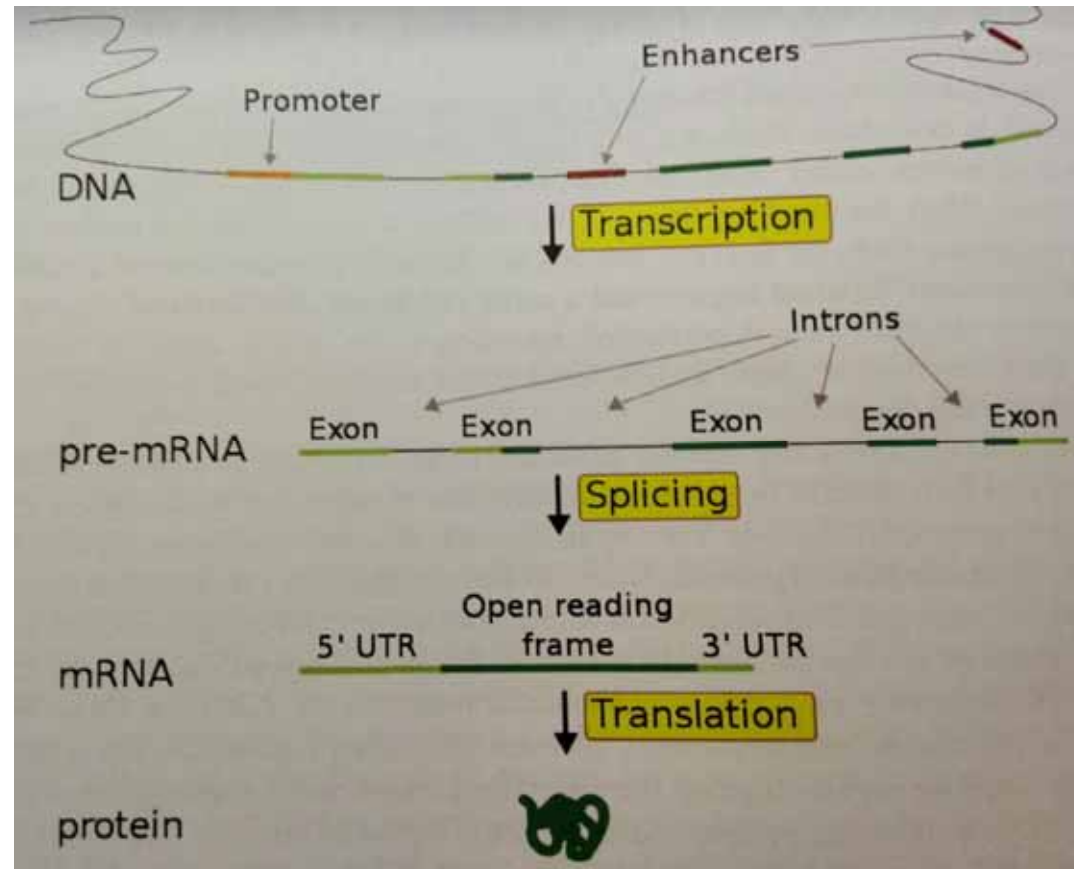
Technology	Sensitivity	Subcellular resolution	Cellular resolution	Minimally invasive?	Live cells?	Real time?
Genetically encoded nanosensors	Nanomolar to millimolar	Nanometer to millimeter	Yes	Yes	Yes	Yes
MRI	Mid-micromolar to millimolar (213)	No	Yes	Yes	Yes	Yes
PET	1–40 Bq mm ⁻² (18)	No	No	No	Yes	Yes
X-ray synchrotron	<1 mg kg ⁻¹ tissue (transit metals) (204)	No	Yes	No	No	No
SIMS	<1 fmol (67)	Yes	Yes	No	No	No
MALDI or TOF imaging	<1 ppm	Yes	50–300 µm (MALDI) 1–2 µm (TOF)	No	No	No
NIMS imaging	Yoctomolar (85)	No	50–300 µm	No	No	No
Mass spectrometry	Yoctomolar	No?	Yes	No	No	No
Raman	50 µM (70)	Yes	Yes	Yes	Yes	Yes

Okumoto, S., Jones, A. & Frommer, W. B. 2012. Quantitative imaging with fluorescent biosensors. *Annual review of plant biology*, 63, 663-706.

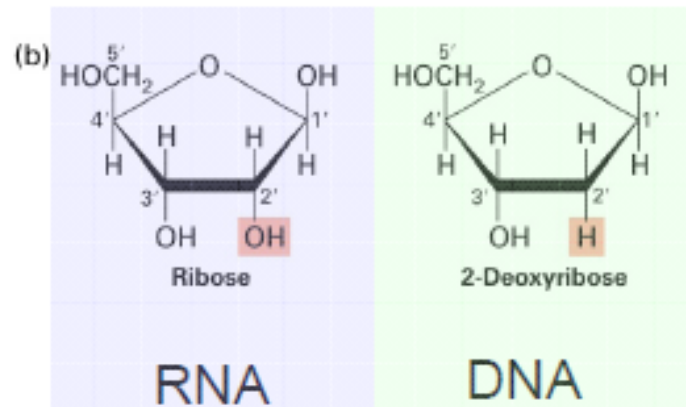
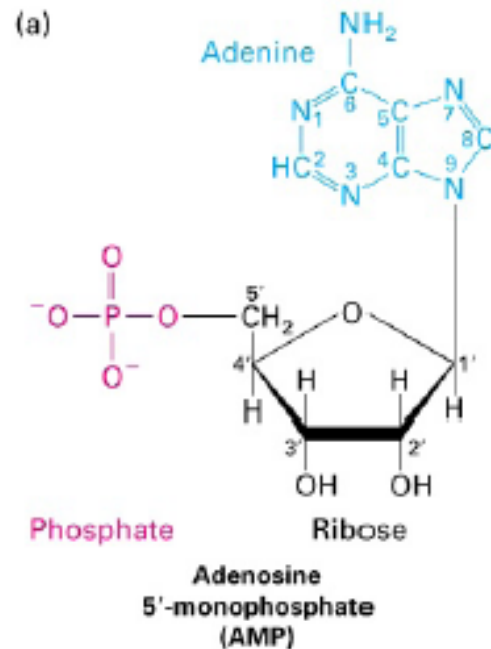
Klibanov, A. M. 2001.
Improving enzymes by
using them in organic
solvents. *Nature*, 409,
(6817), 241-246.



The DNA, the RNA and the proteins are the three major macromolecules essential for all known forms of life.

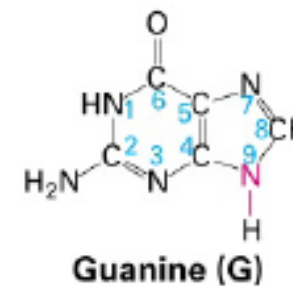
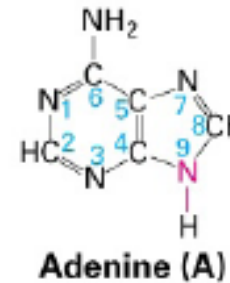


Manca, V. (2013). Infobiotics. Springer.

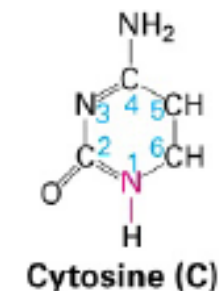
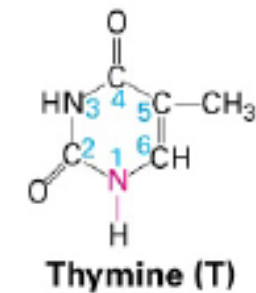
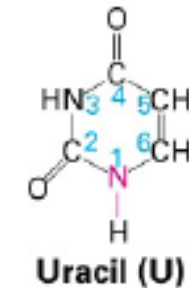


The five principal bases

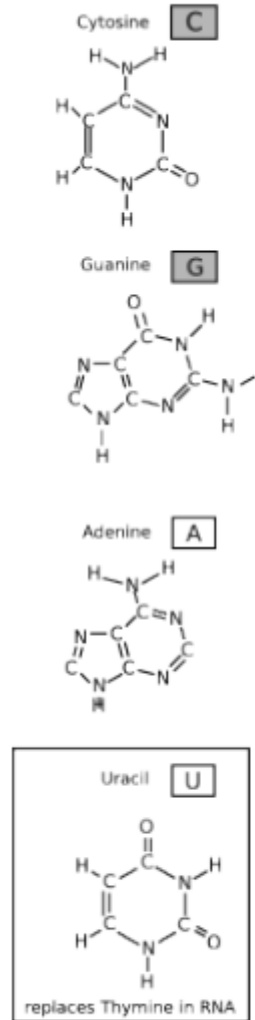
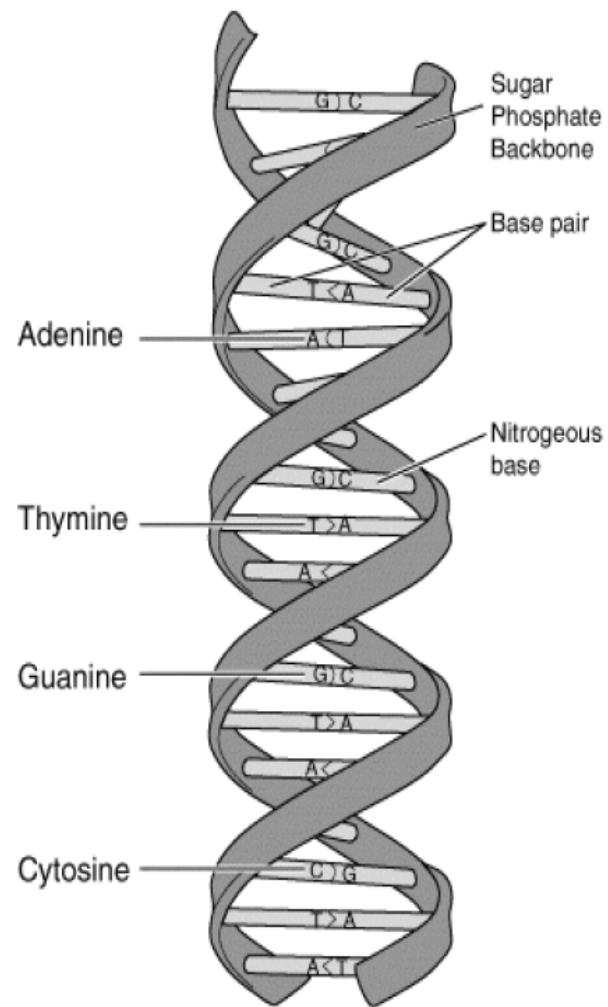
PURINES



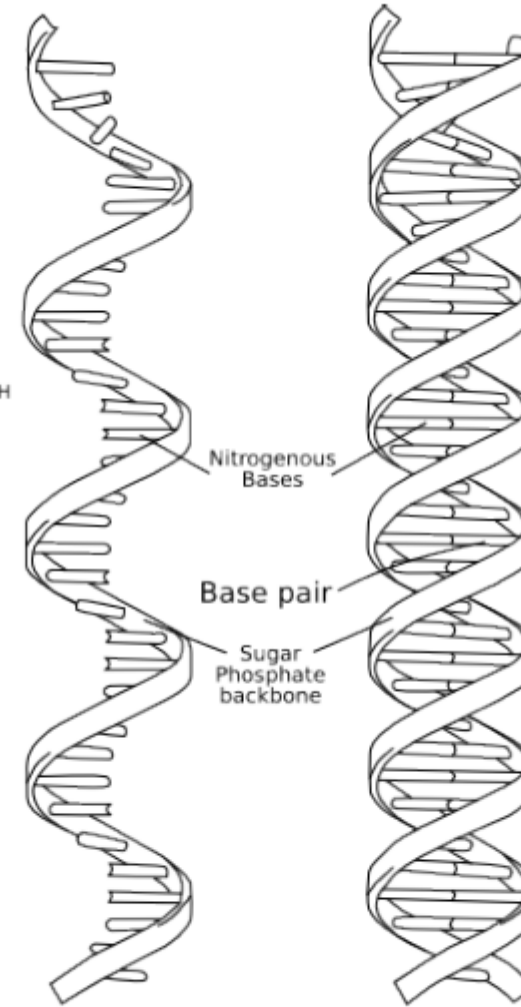
PYRIMIDINES



A, G, T, C are present in DNA (DeoxyriboNucleic Acid)
A, G, U, C are present in RNA (RiboNucleic Acid)



Nitrogenous Bases



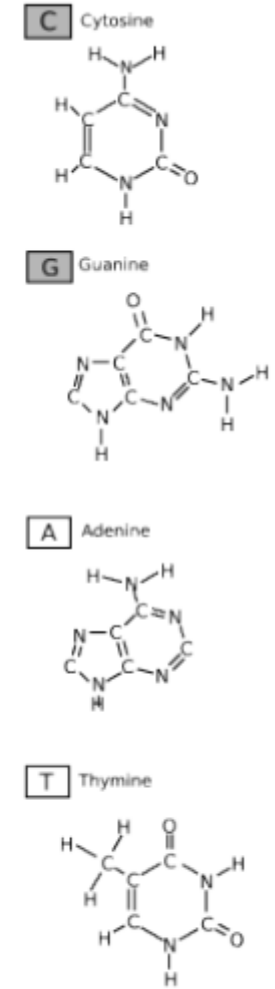
RNA

Ribonucleic acid

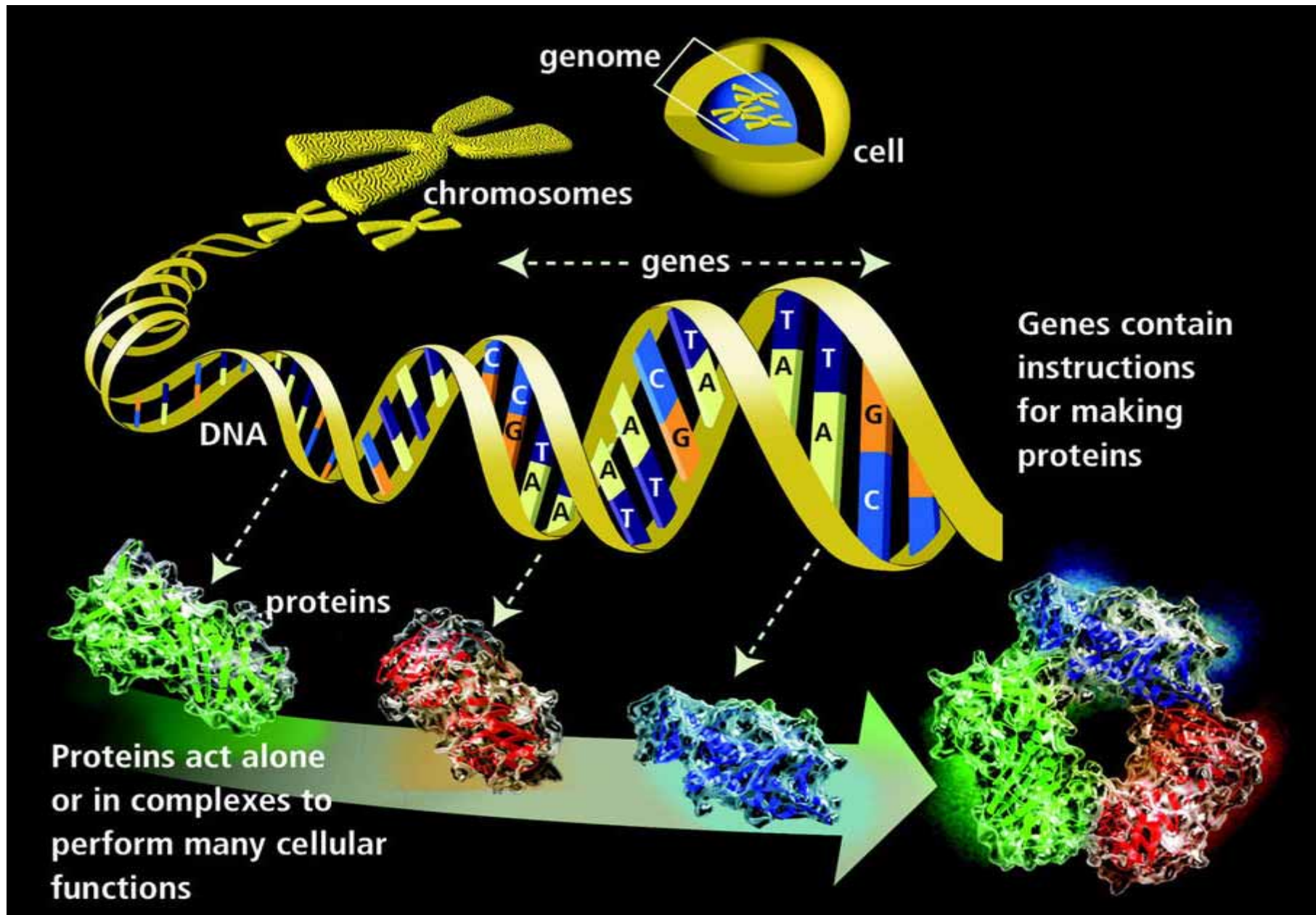


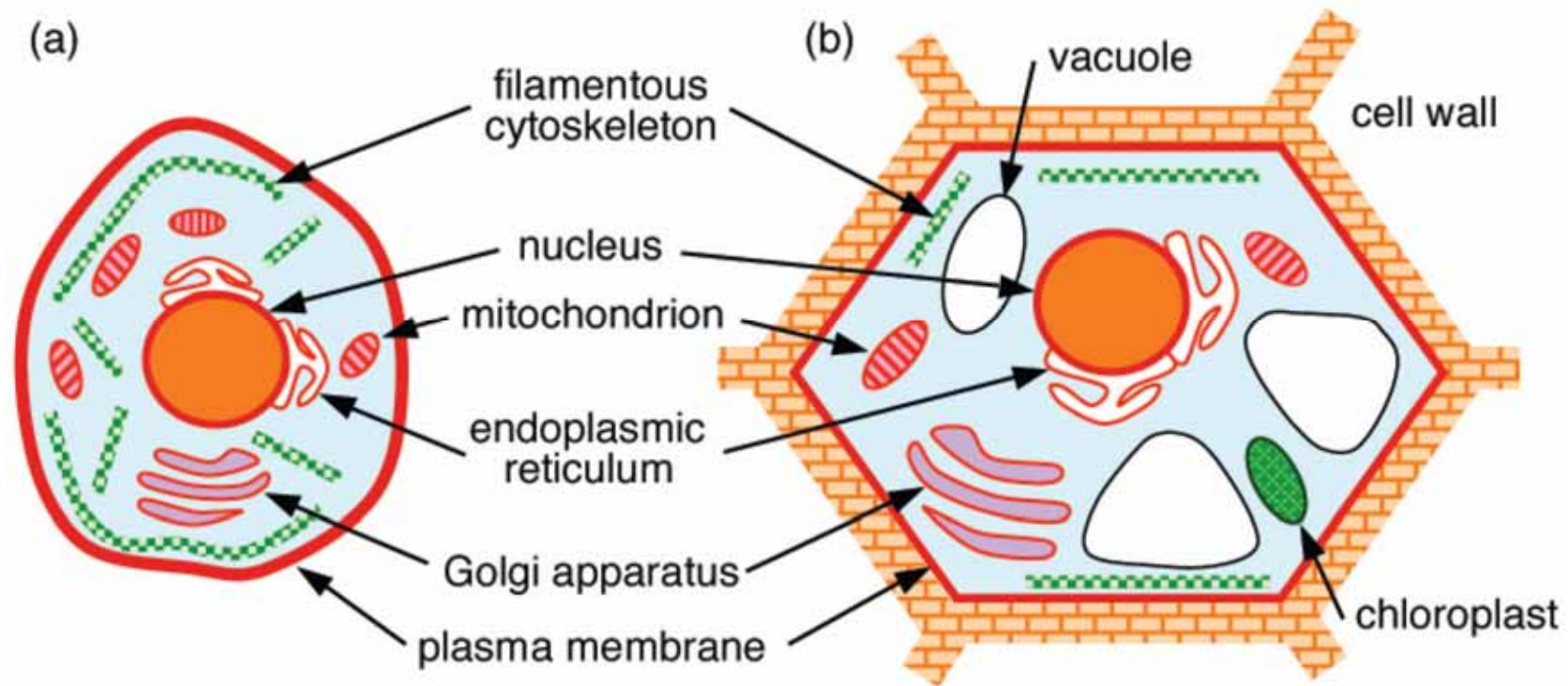
DNA

Deoxyribonucleic acid

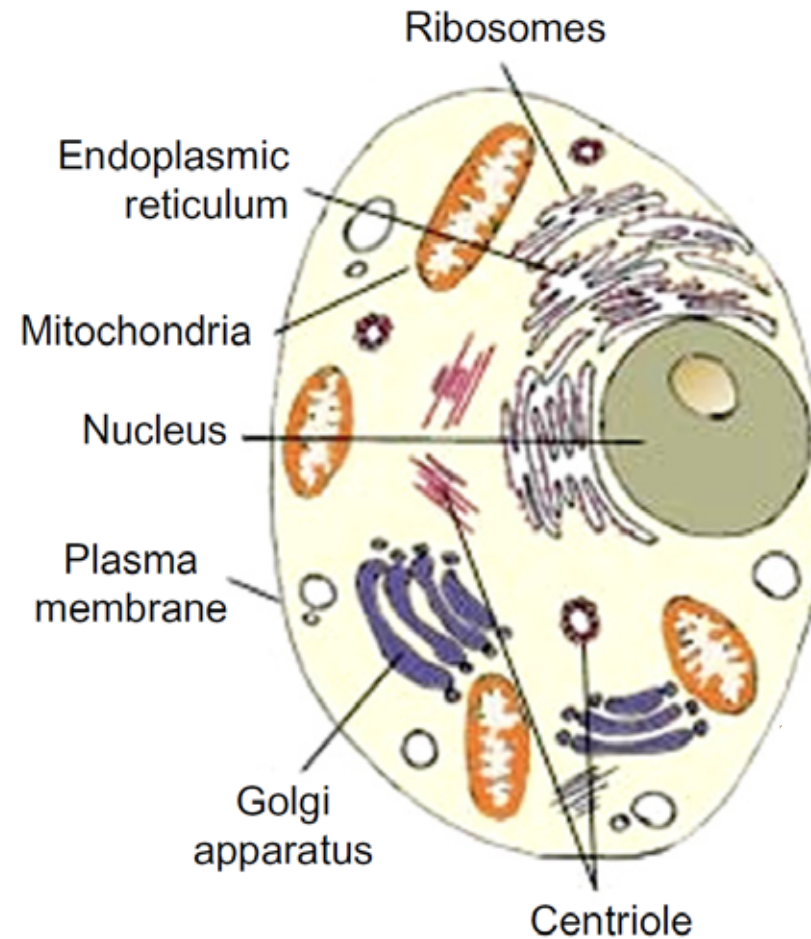
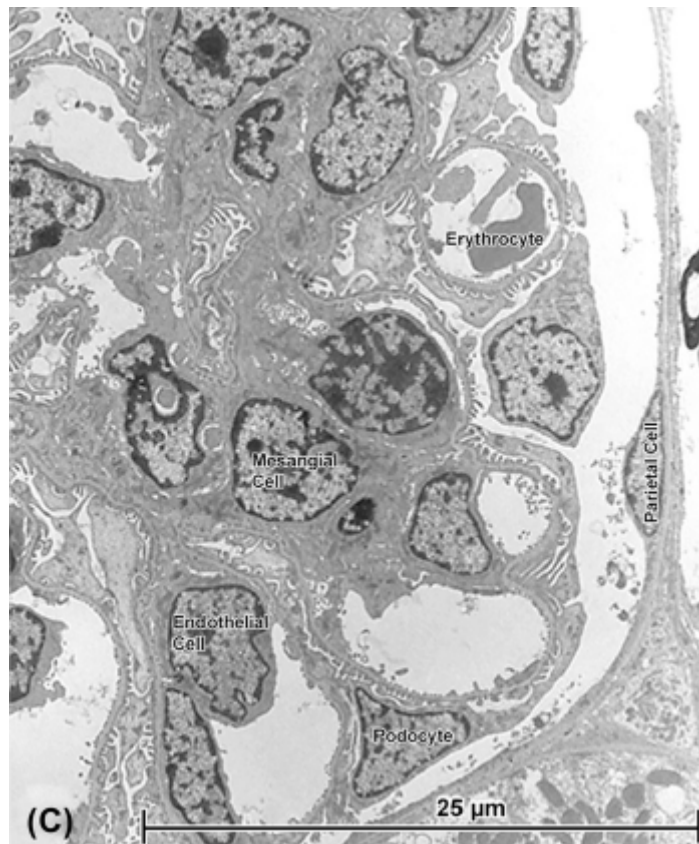


Nitrogenous Bases

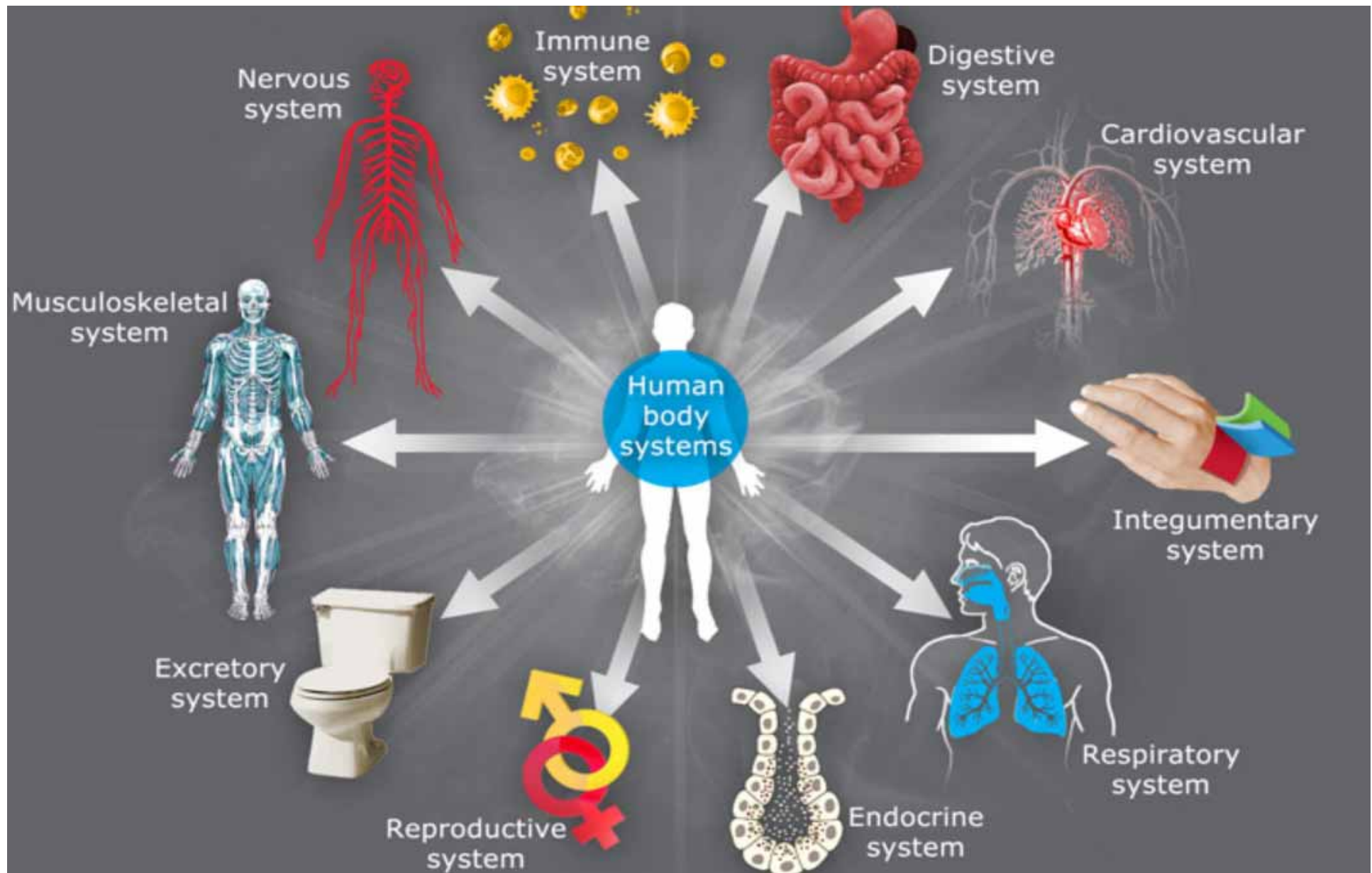


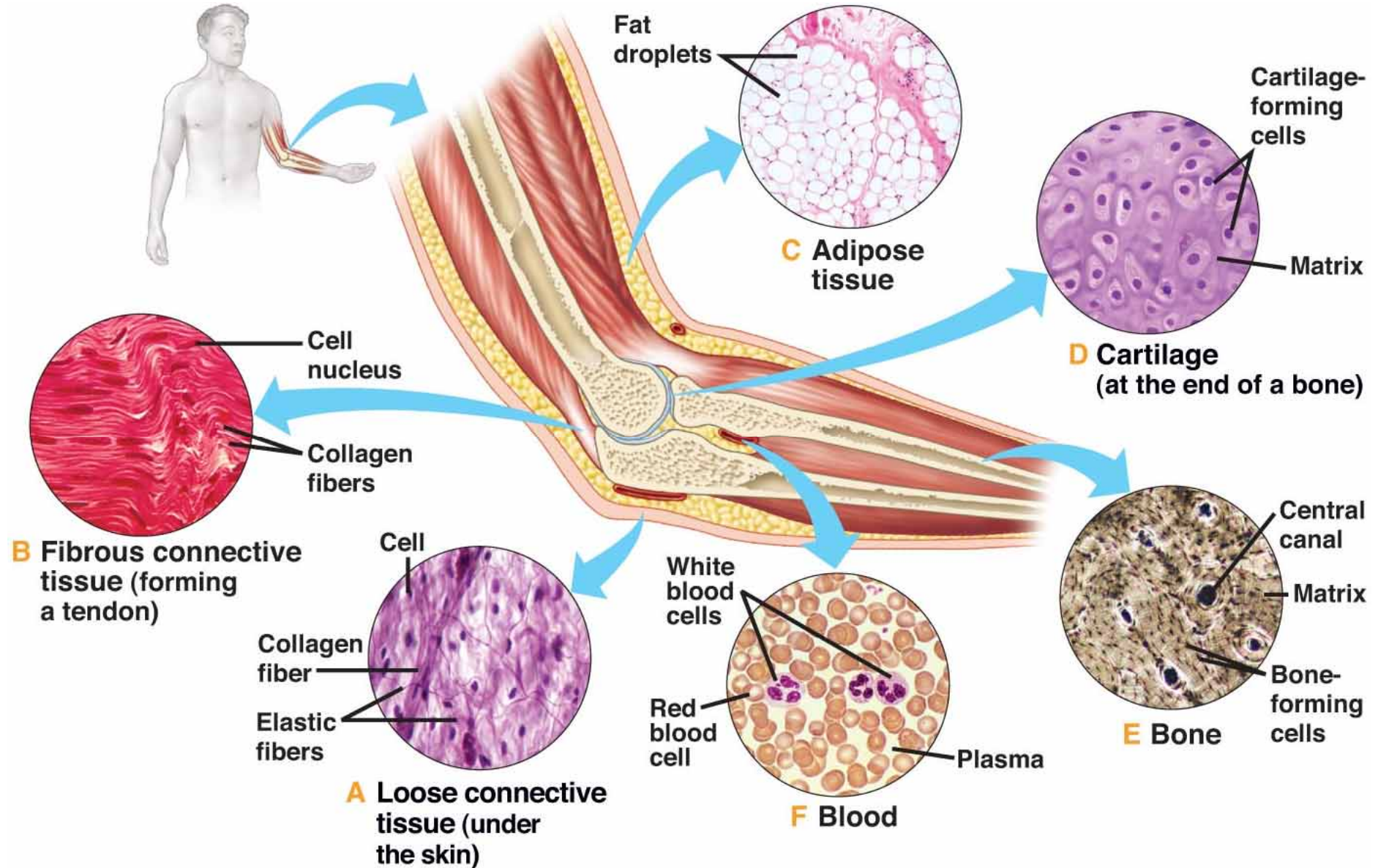


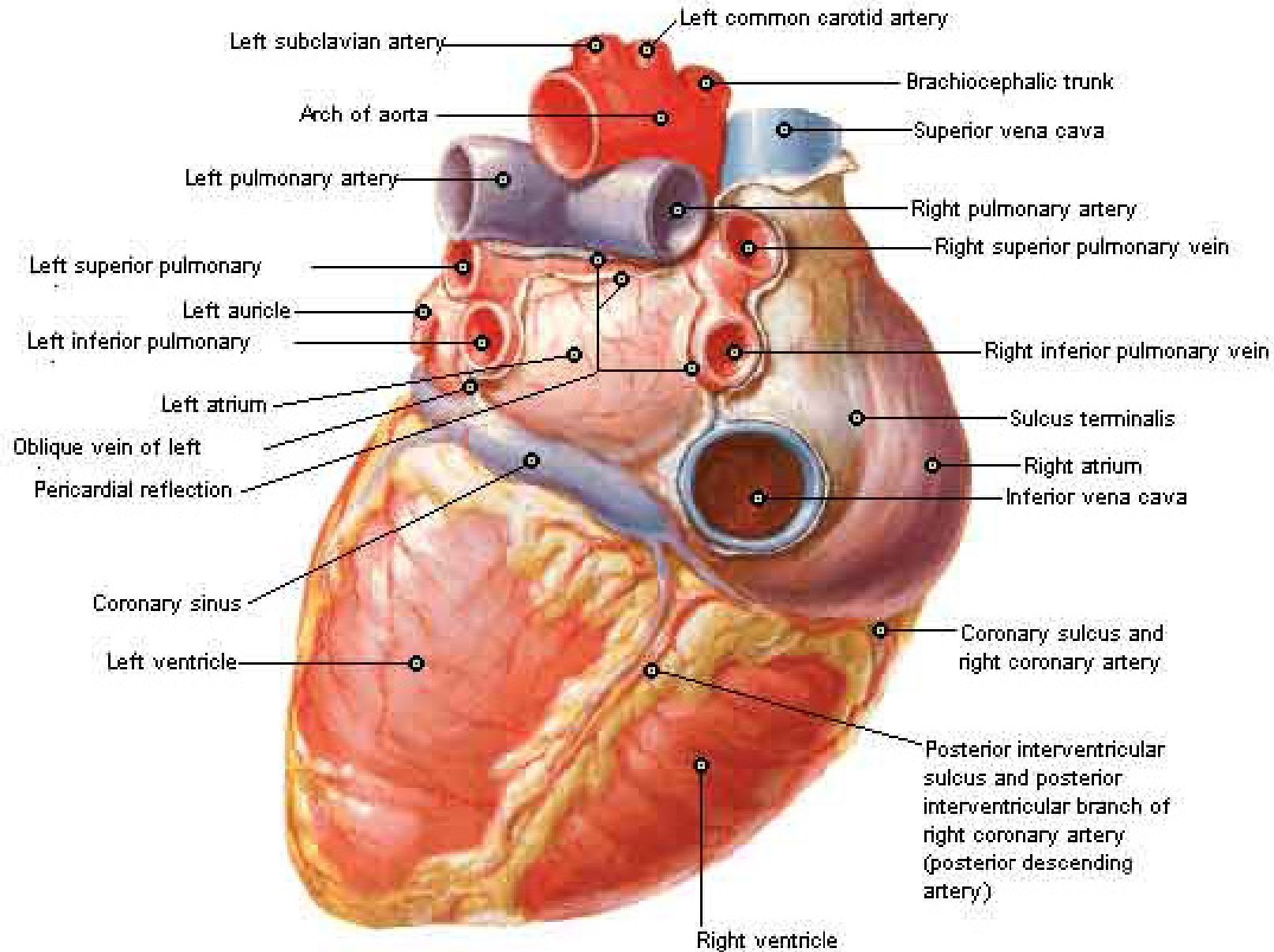
Boal, D. 2012. *Mechanics of the Cell*, Cambridge University Press.

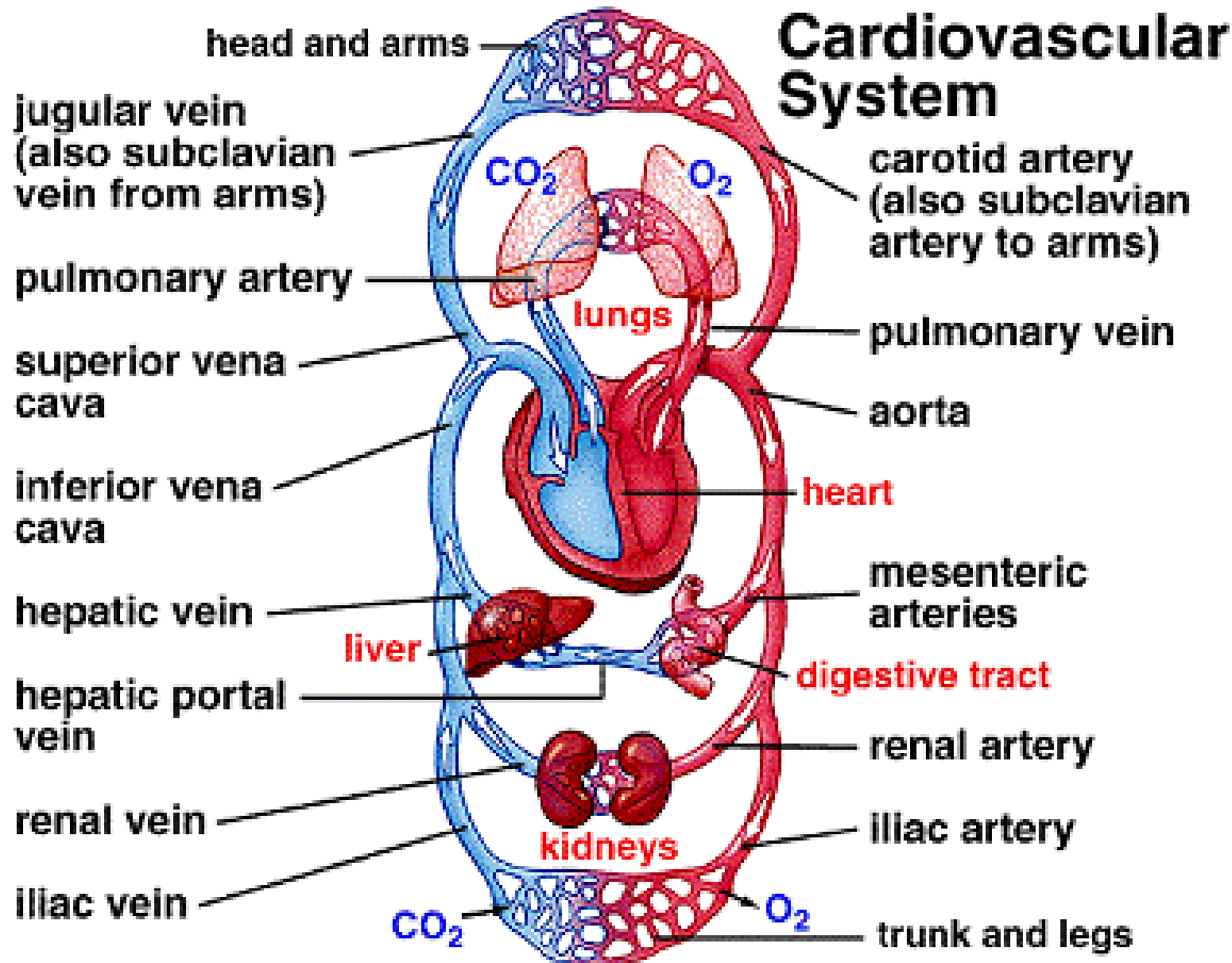


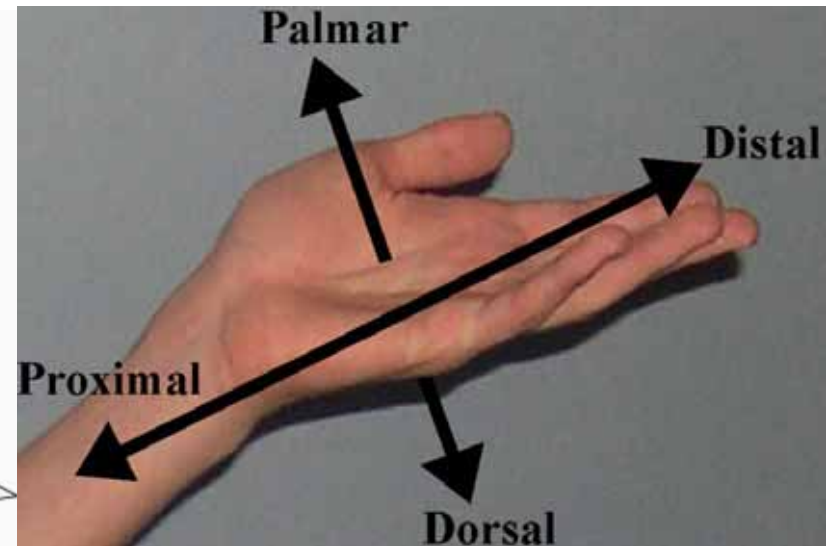
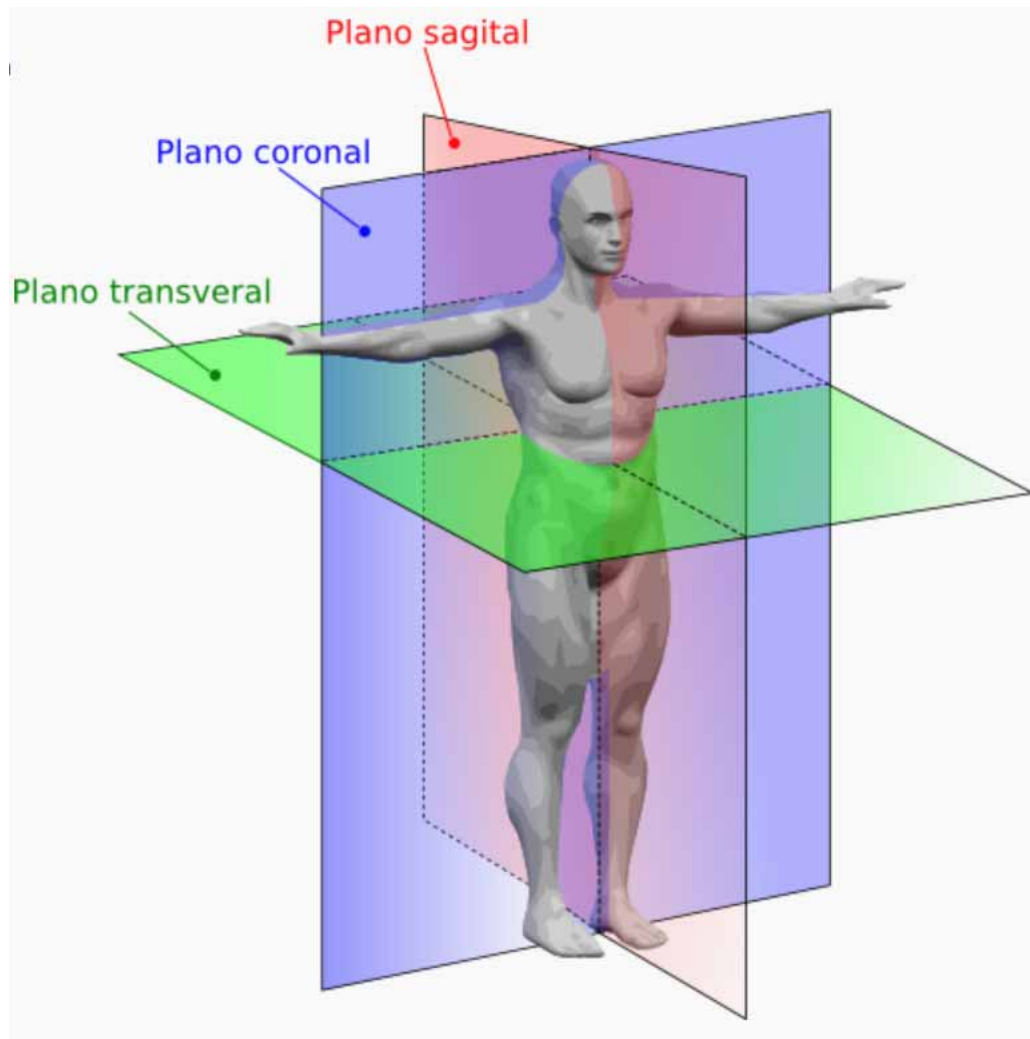
Sperelakis, N. 2012. *Cell Physiology Sourcebook: Essentials of Membrane Biophysics. Fourth Edition*, Amsterdam, Elsevier.

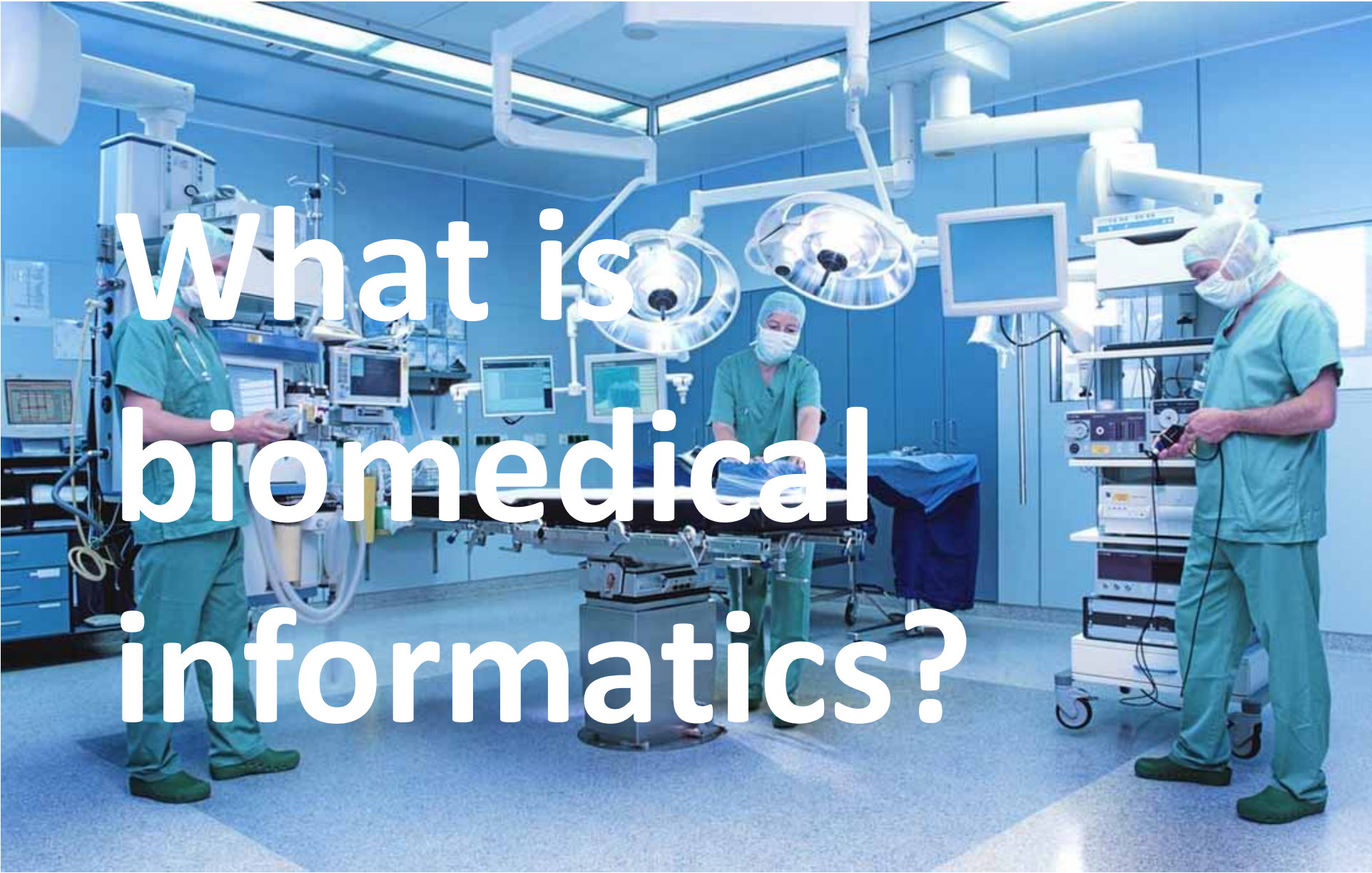










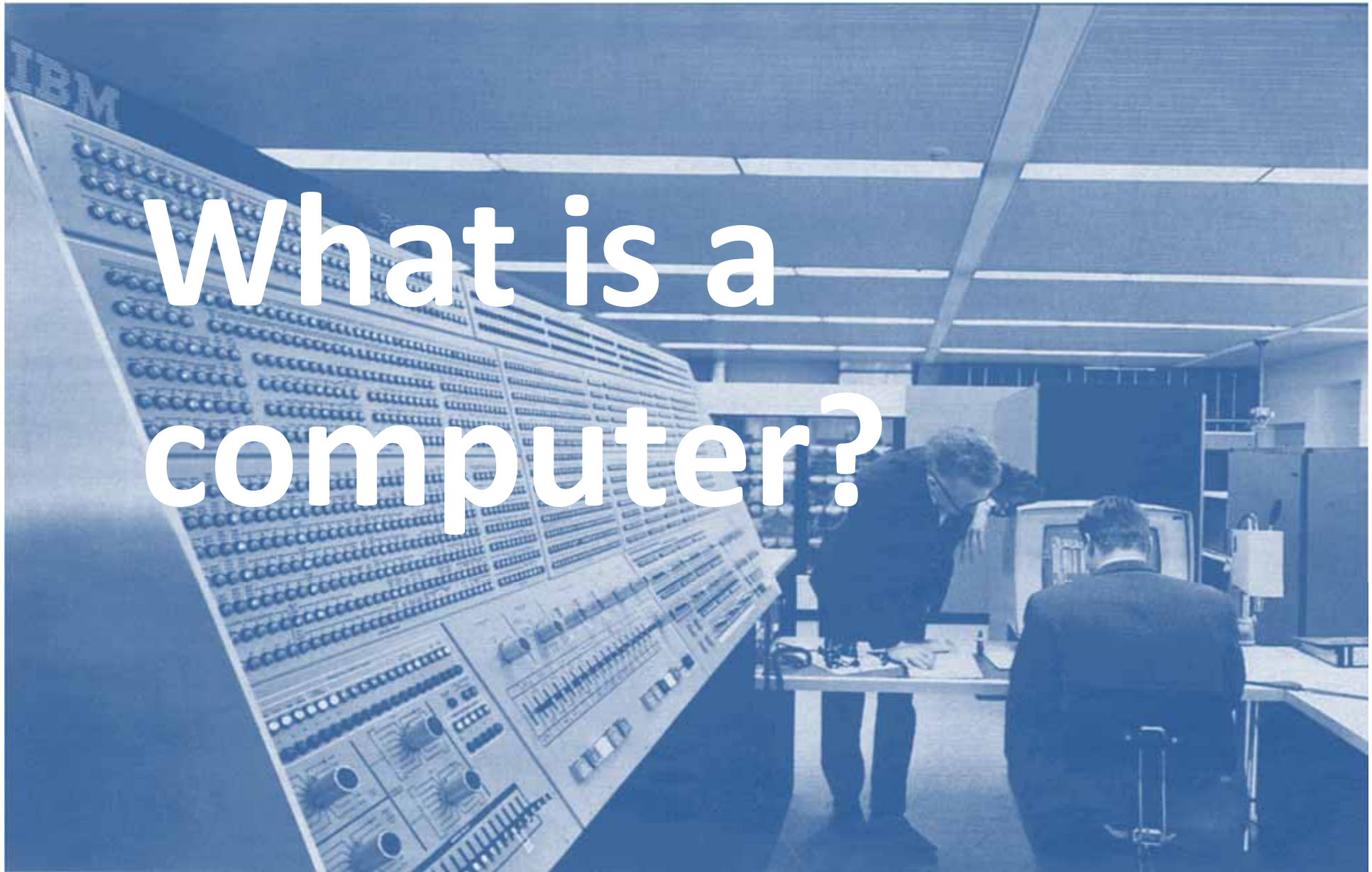
A photograph of an operating room with three surgeons in green scrubs and masks. They are surrounded by medical equipment, including monitors and surgical lights. The text "What is biomedical informatics?" is overlaid in large white letters.

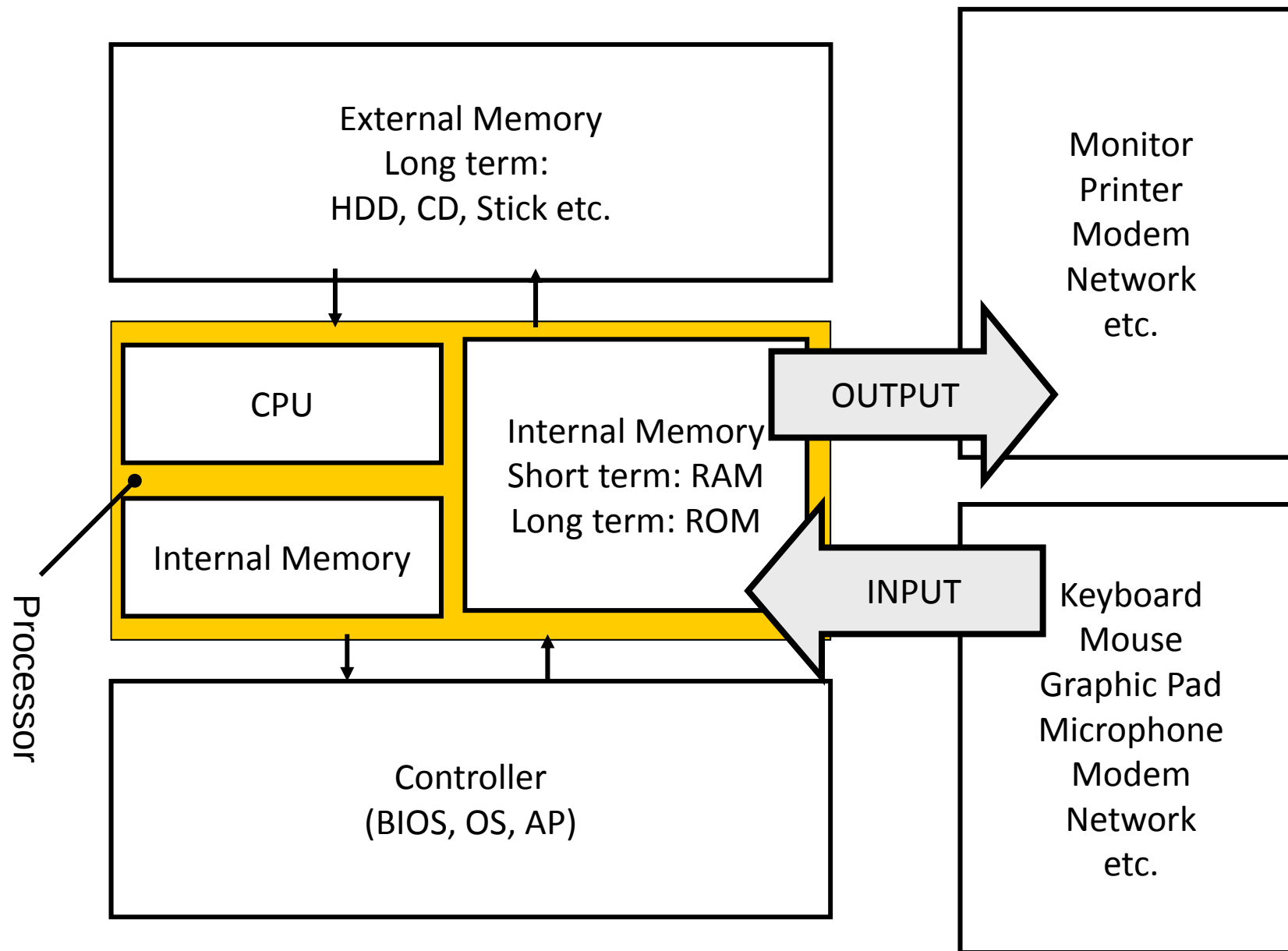
What is biomedical informatics?



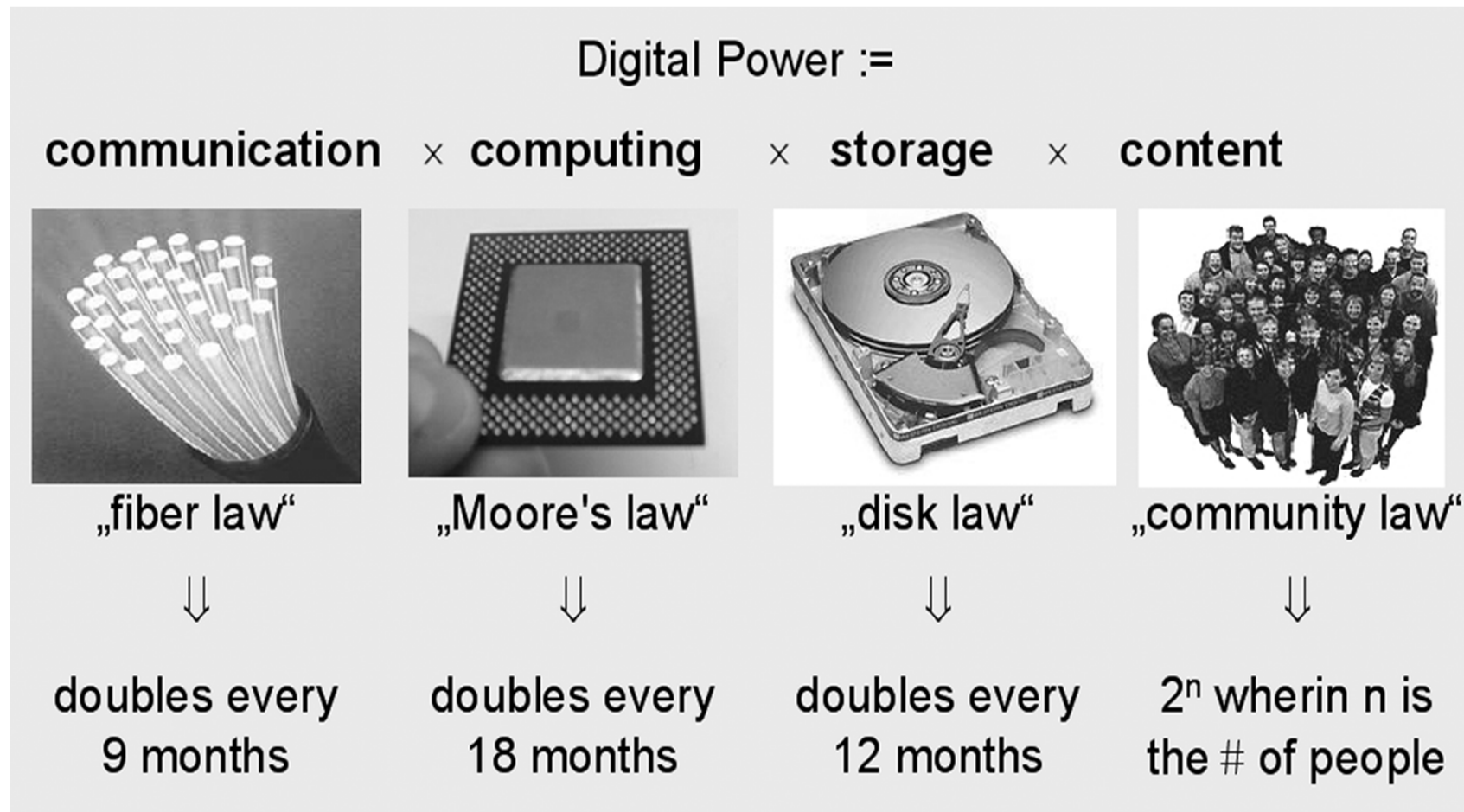
Yapijakis, C. (2009) Hippocrates of Kos, the Father of Clinical Medicine, and Asclepiades of Bithynia, the Father of Molecular Medicine. *In Vivo*, 23, 4, 507-514.

What is a computer?



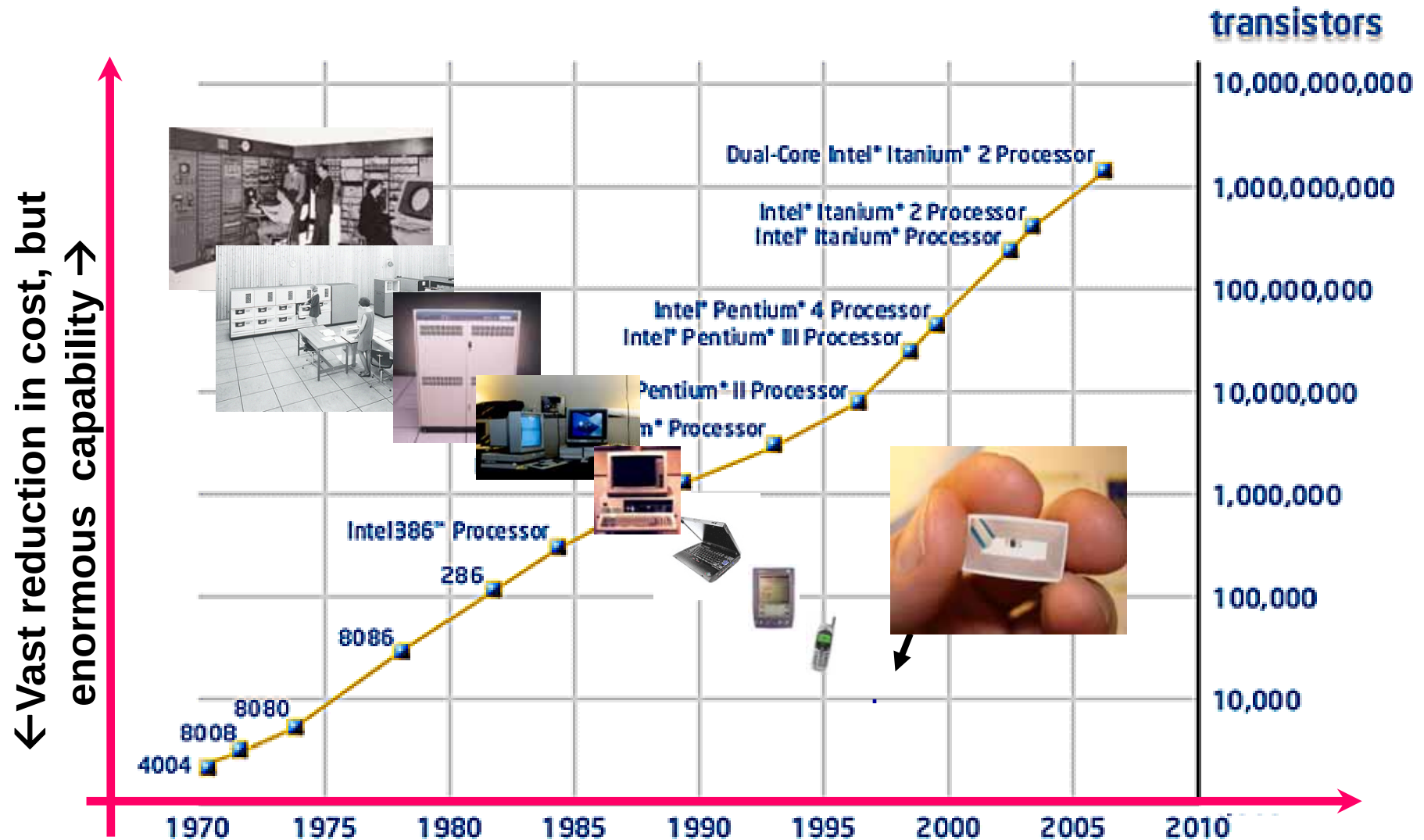


Gordon E. Moore (1965, 1989, 1997)

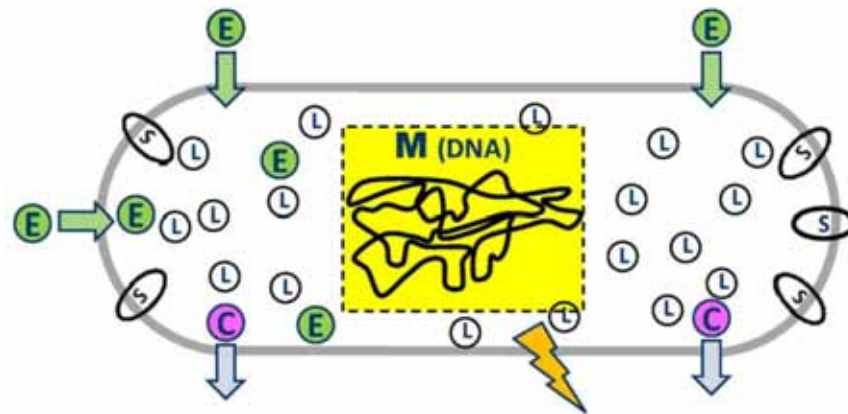


Holzinger, A. 2002. *Basiswissen IT/Informatik Band 1: Informationstechnik. Das Basiswissen für die Informationsgesellschaft des 21. Jahrhunderts*, Wuerzburg, Vogel Buchverlag.

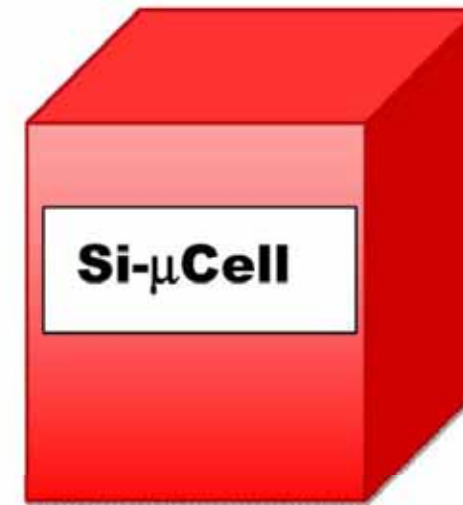
Slide 1-31: Computer cost/size versus Performance



Cf. with Moore (1965), Holzinger (2002), Scholtz & Consolvo (2004), Intel (2007)



Memory:	10^7 bit
Logic:	$>10^6$ bit
Power:	10^{-13} W
Heat:	10^{-6} W/cm ²
Energy/task*:	10^{-10} J
Task time*:	2400s=40min

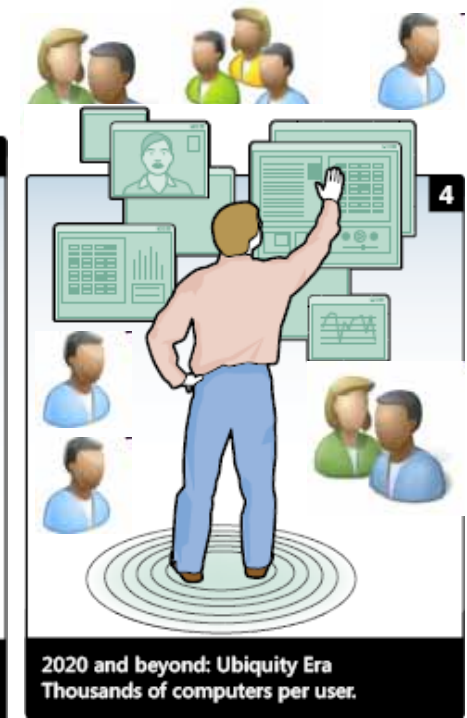
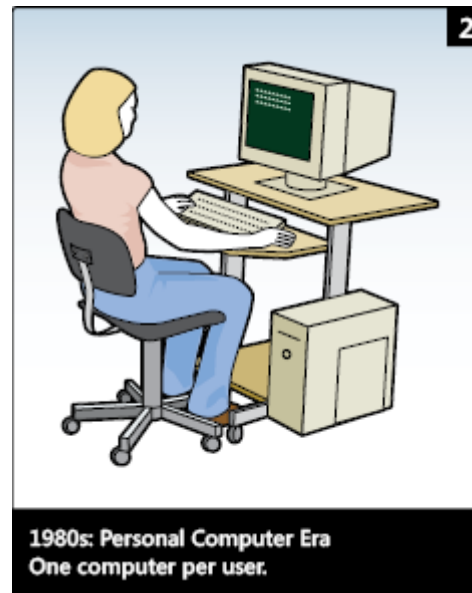


Memory:	$\sim 10^4$ bit
Logic:	$\sim 300-150,000$ bit
Power:	$\sim 10^{-7}$ W
Heat:	~ 1 W/cm ²
Energy/task*:	$\sim 10^{-2}$ J
Task time*:	510,000 s $\sim$ 6 days

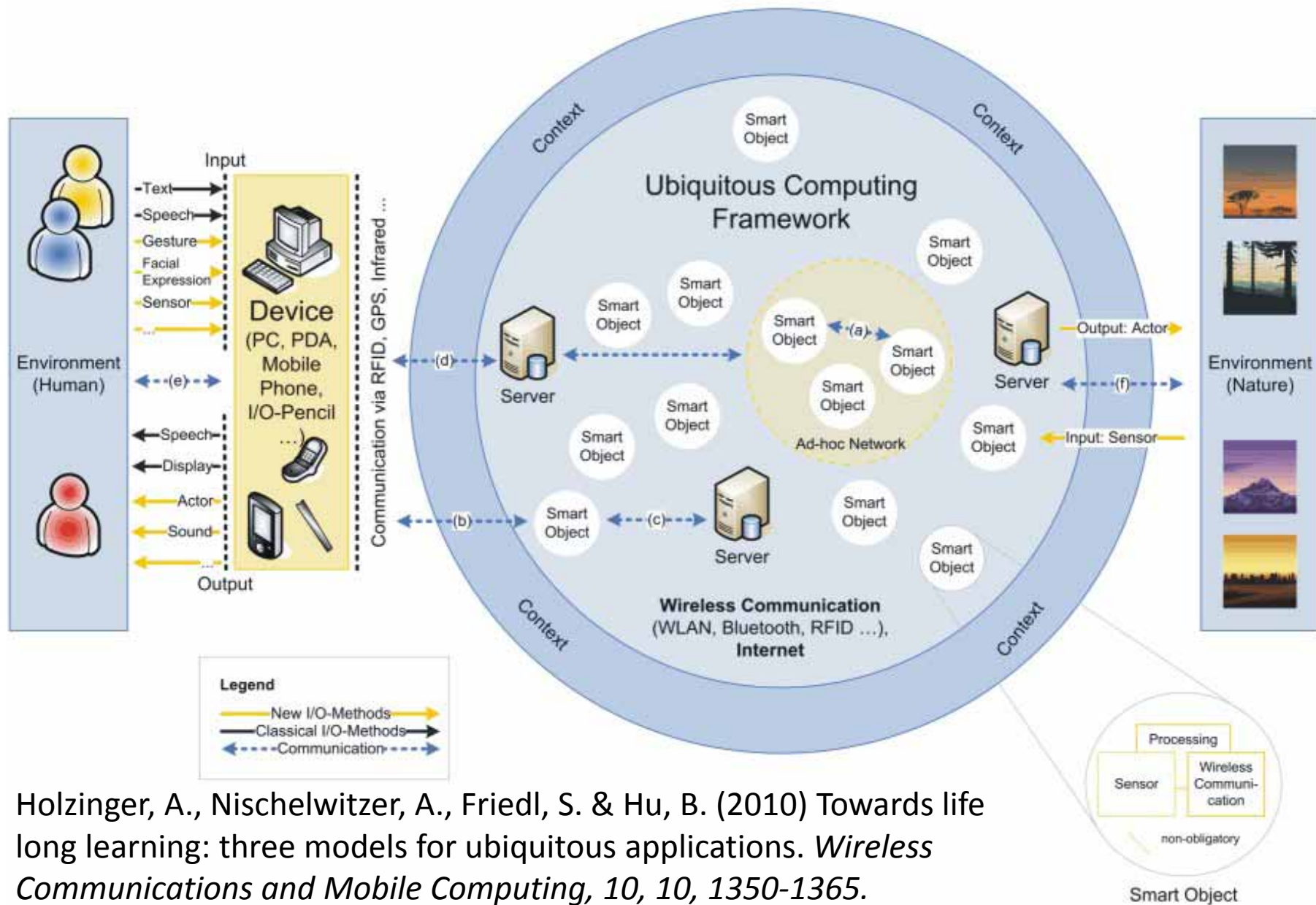
*Equivalent to 10^{11} output bits

Cavin, R., Lugli, P. & Zhirnov, V. 2012. Science and Engineering Beyond Moore's Law. *Proc. of the IEEE*, 100, 1720-49 (L=Logic-Protein; S=Sensor-Protein; C=Signaling-Molecule, E=Glucose-Energy)

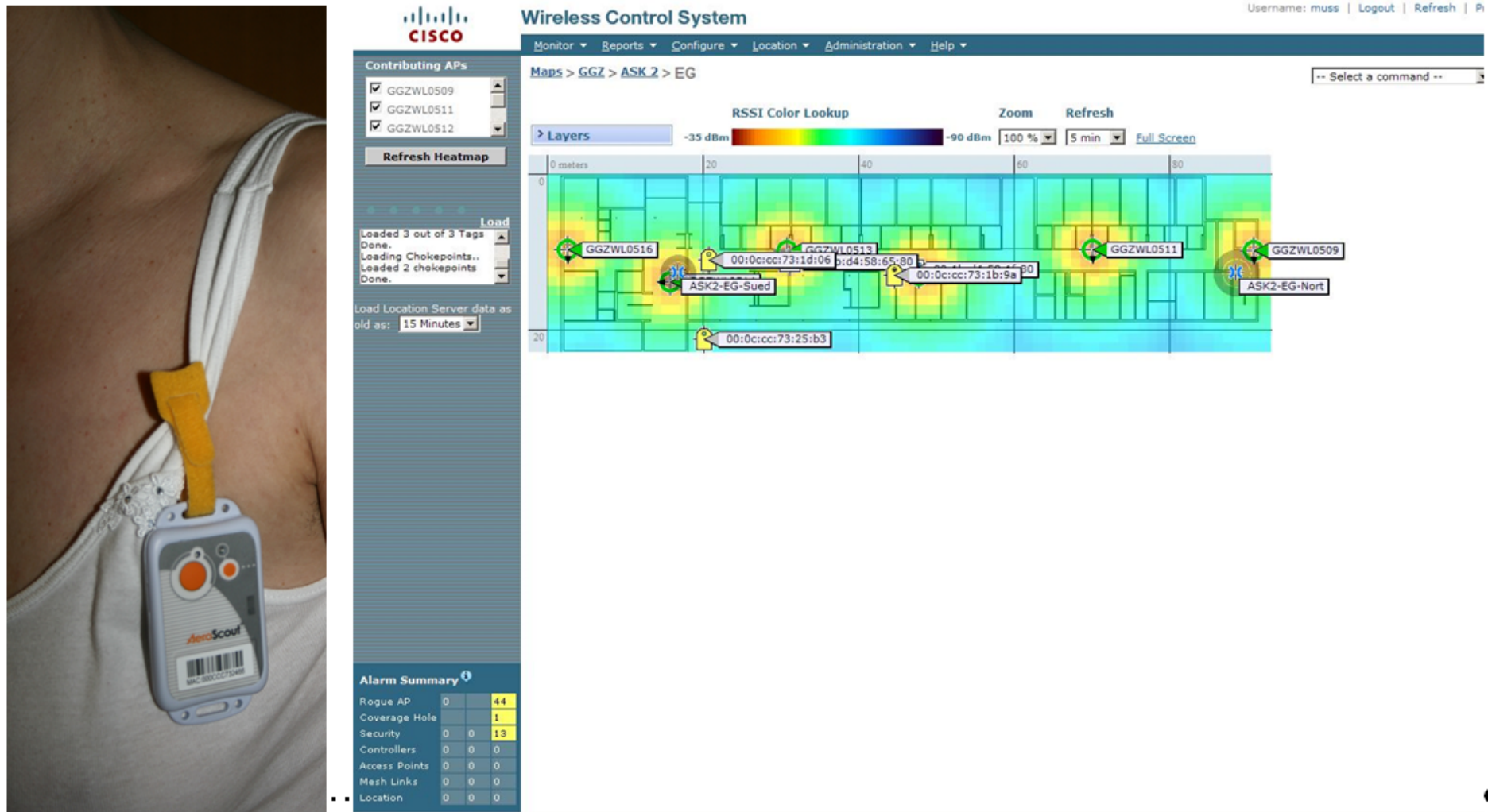
- ... using technology to augment human capabilities for structuring, retrieving and managing information



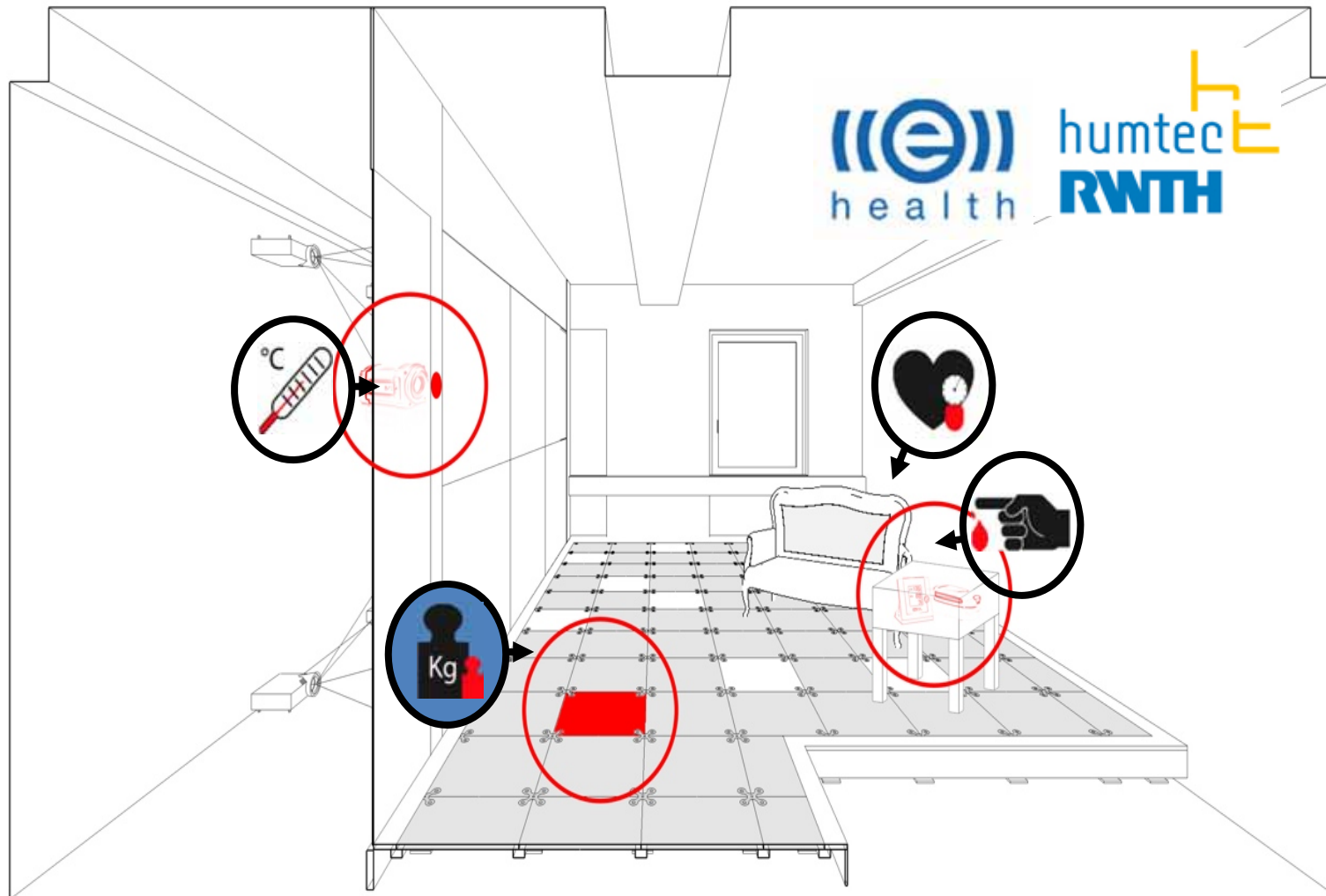
Harper, R., Rodden, T., Rogers, Y. & Sellen, A. (2008) *Being Human: Human-Computer Interaction in the Year 2020*. Cambridge, Microsoft Research.



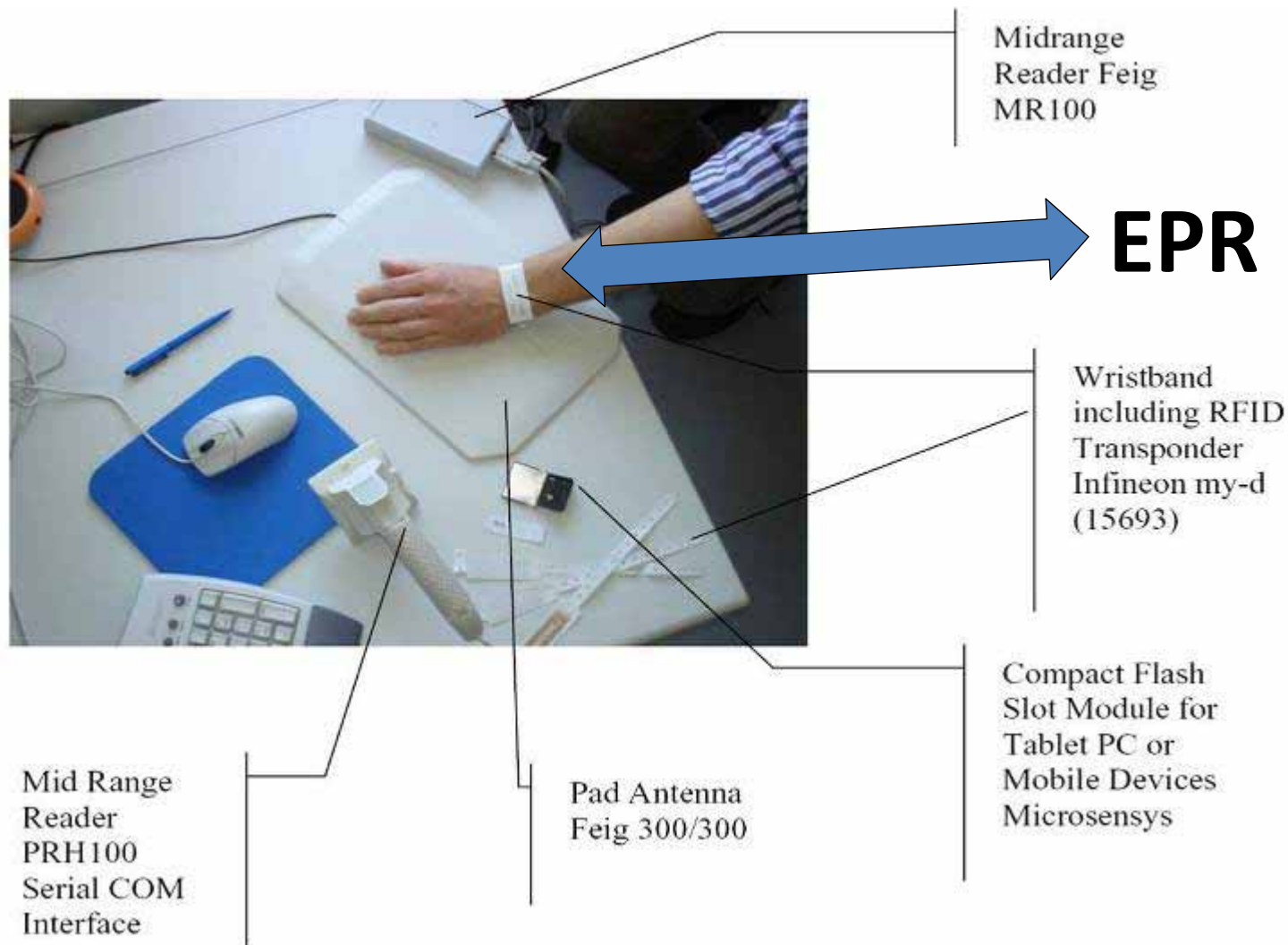
Holzinger, A., Nischelwitz, A., Friedl, S. & Hu, B. (2010) Towards life long learning: three models for ubiquitous applications. *Wireless Communications and Mobile Computing*, 10, 10, 1350-1365.



Holzinger, A., Schaupp, K. & Eder-Halbedl, W. (2008) An Investigation on Acceptance of Ubiquitous Devices for the Elderly in an Geriatric Hospital Environment: using the Example of Person Tracking In: *Lecture Notes in Computer Science (LNCS 5105)*. Heidelberg, Springer, 22-29.

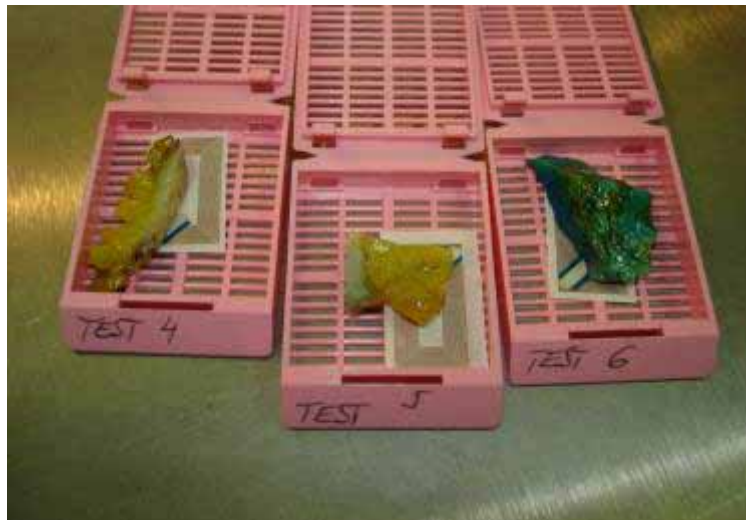


Alagoez, F., Valdez, A. C., Wilkowska, W., Ziefle, M., Dorner, S. & Holzinger, A. (2010) From cloud computing to mobile Internet, from user focus to culture and hedonism: The crucible of mobile health care and Wellness applications. *5th International Conference on Pervasive Computing and Applications (ICPCA)*. IEEE, 38-45.

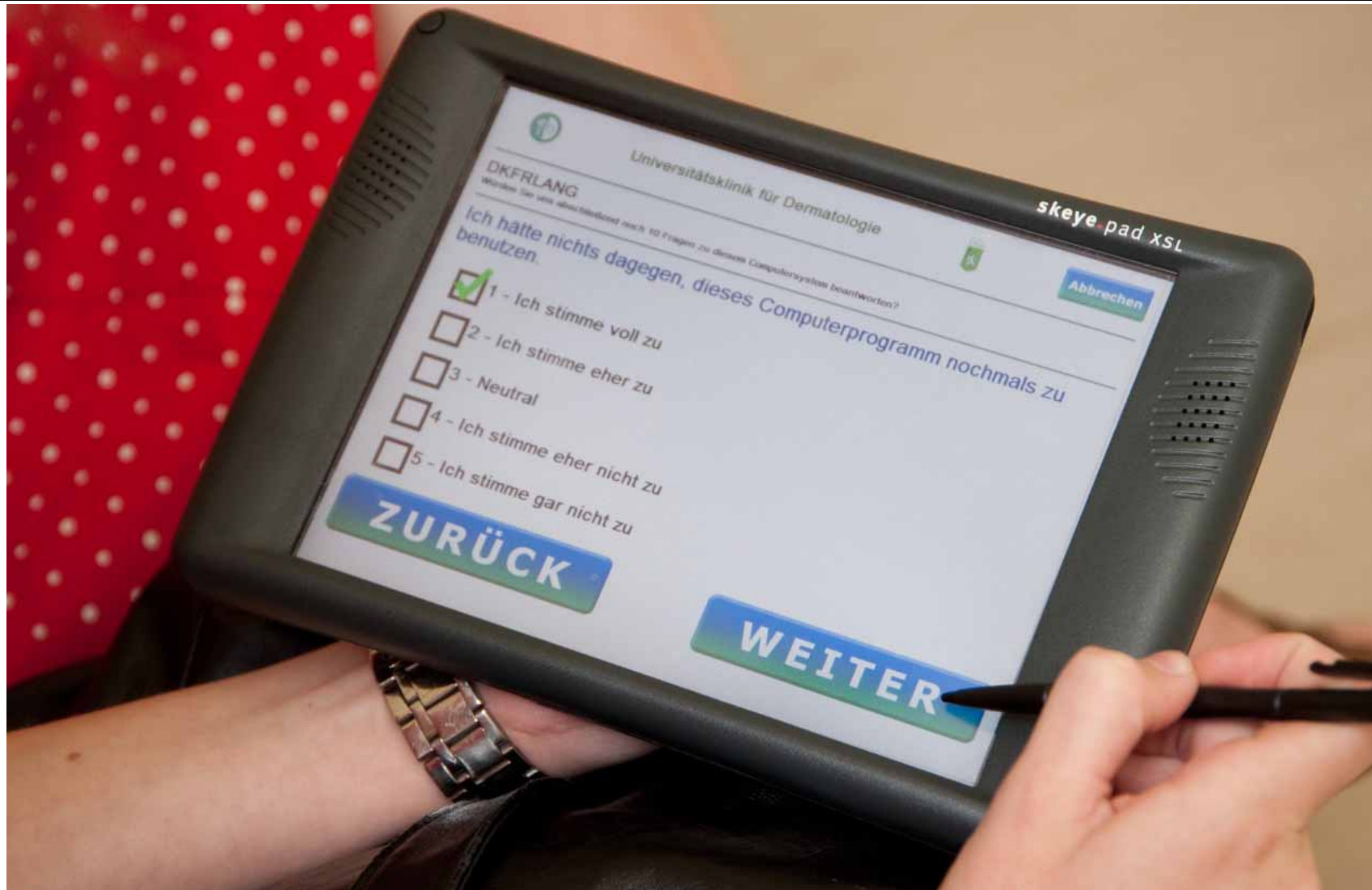


Holzinger, A., Schwabberger, K. & Weitlaner, M. (2005) Ubiquitous Computing for Hospital Applications: RFID-Applications to enable research in Real-Life environments *29th Annual IEEE International Computer Software & Applications Conference (IEEE COMPSAC)*, 19-20.

Slide 1-38: Smart Objects in the pathology



Holzinger et al. (2005)



Holzinger, A., Kosec, P., Schwantzer, G., Debevc, M., Hofmann-Wellenhof, R. & Frühauf, J. 2011. Design and Development of a Mobile Computer Application to Reengineer Workflows in the Hospital and the Methodology to evaluate its Effectiveness. *Journal of Biomedical Informatics*, 44, 968-977.

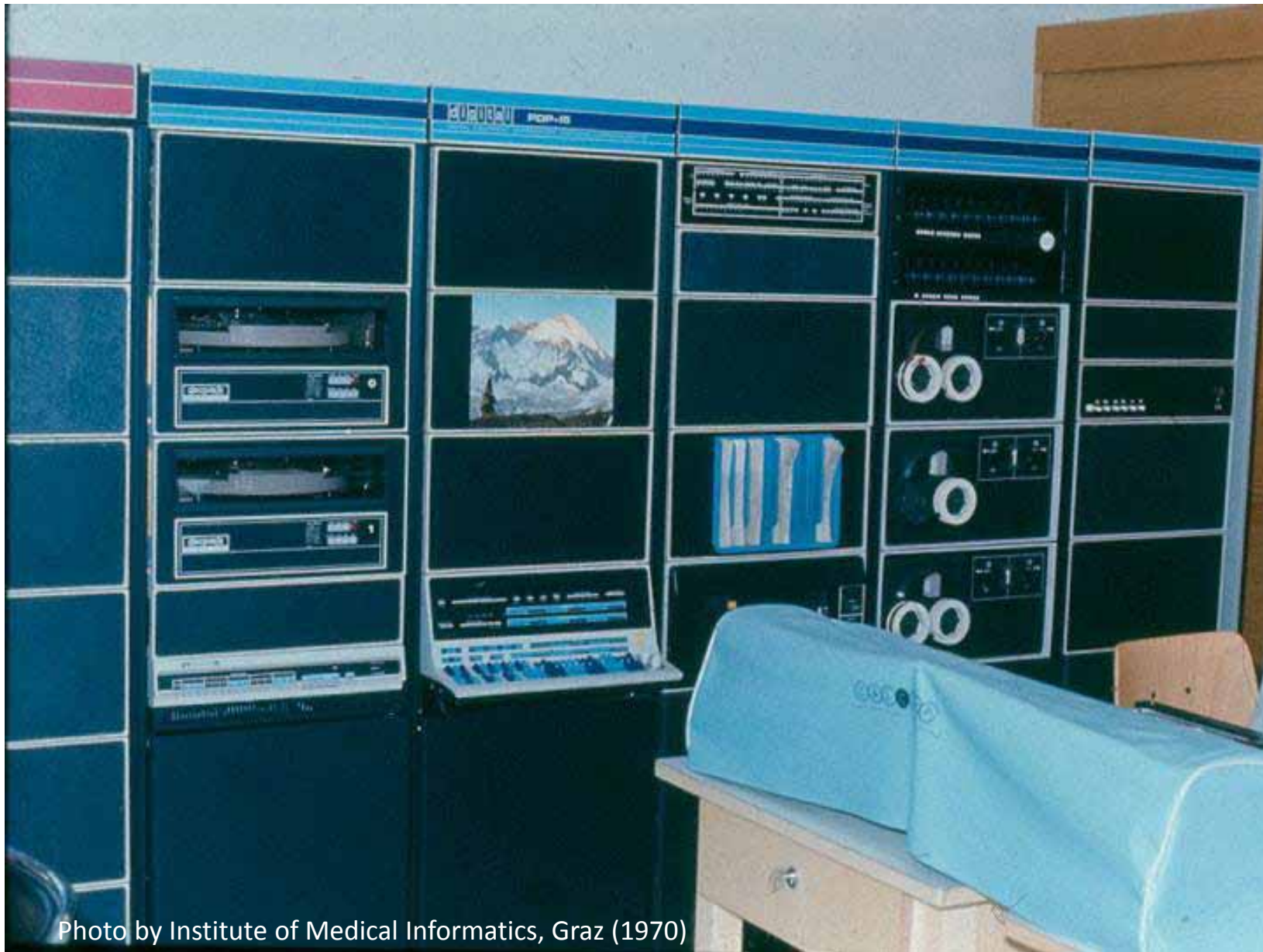


Photo by Institute of Medical Informatics, Graz (1970)

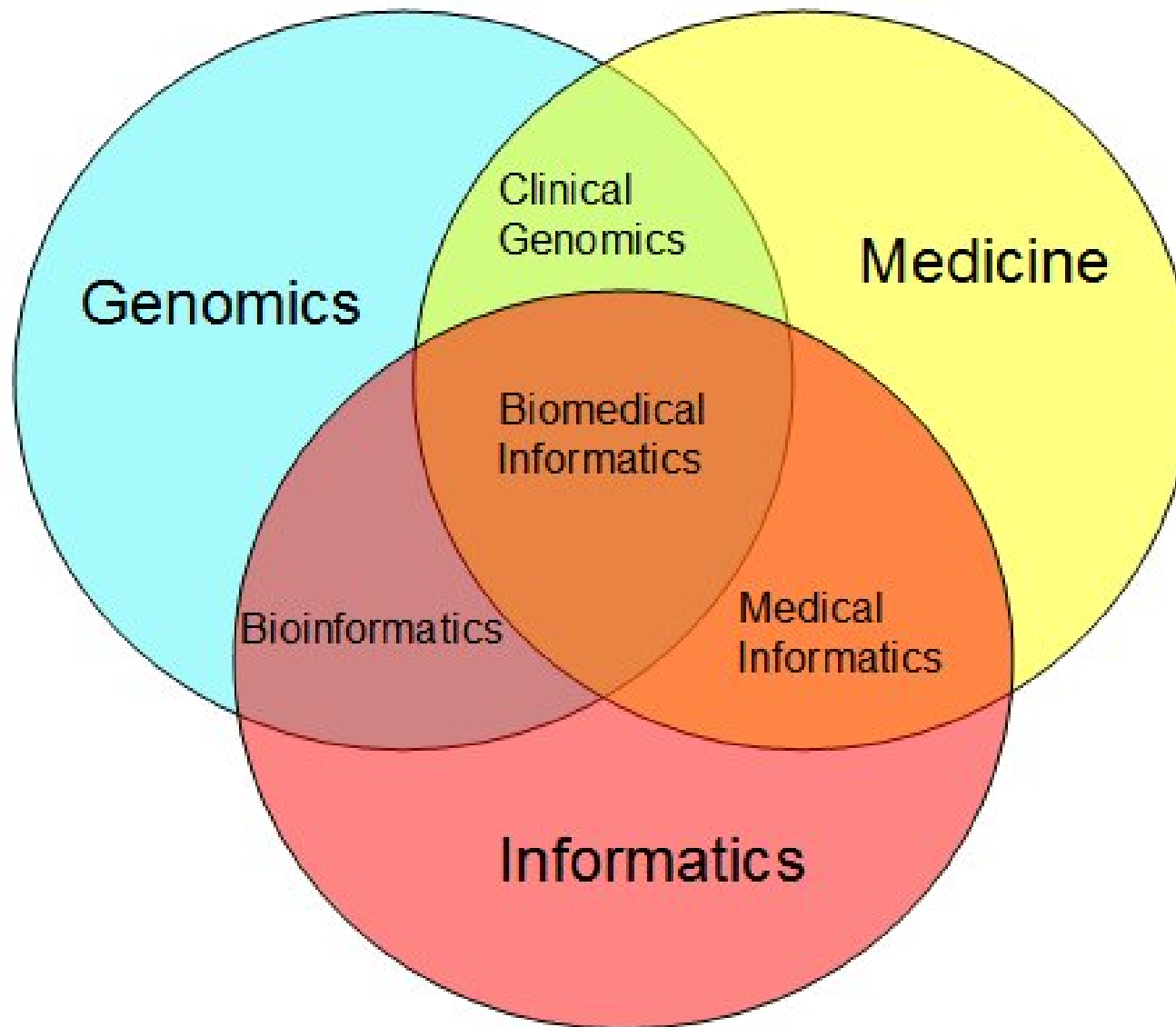
- 1970+ Begin of **Medical Informatics**
 - Focus on data acquisition, storage, accounting (typ. “EDV”)
 - The term was first used in 1968 and the first course was set up 1978
- 1985+ Health Telematics
 - Health care networks, Telemedicine, CPOE-Systems etc.
- 1995+ Web Era
 - Web based applications, Services, EPR, etc.
- 2005+ Ambient Era
 - Pervasive & Ubiquitous Computing
- 2010+ Quality Era – **Biomedical Informatics**
 - Information Quality, Patient empowerment, individual molecular medicine, End-User Programmable Mashups





- ***Biomedical informatics (BMI)*** is the interdisciplinary field that studies and pursues the effective use of biomedical data, information, and knowledge for scientific problem solving, and decision making, motivated by efforts to improve human health

Shortliffe, E. H. (2011). Biomedical Informatics: Defining the Science and its Role in Health Professional Education. In A. Holzinger & K.-M. Simoncic (Eds.), *Information Quality in e-Health. Lecture Notes in Computer Science LNCS 7058* (pp. 711-714). Heidelberg, New York: Springer.



<http://www.bioinformaticslaboratory.nl/twiki/bin/view/BioLab/EducationMIK1-2>



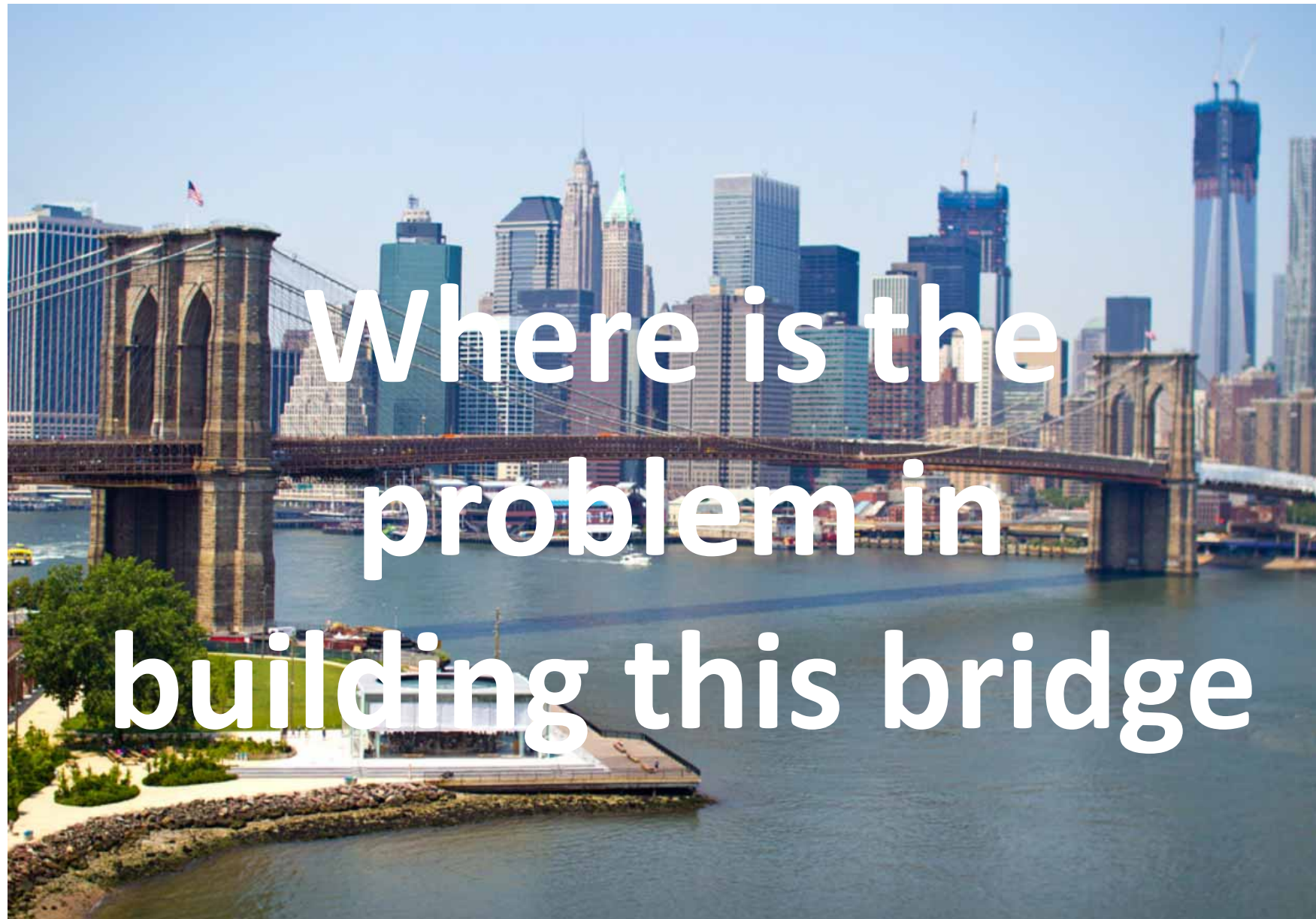
Our central hypothesis: Information bridges this gap

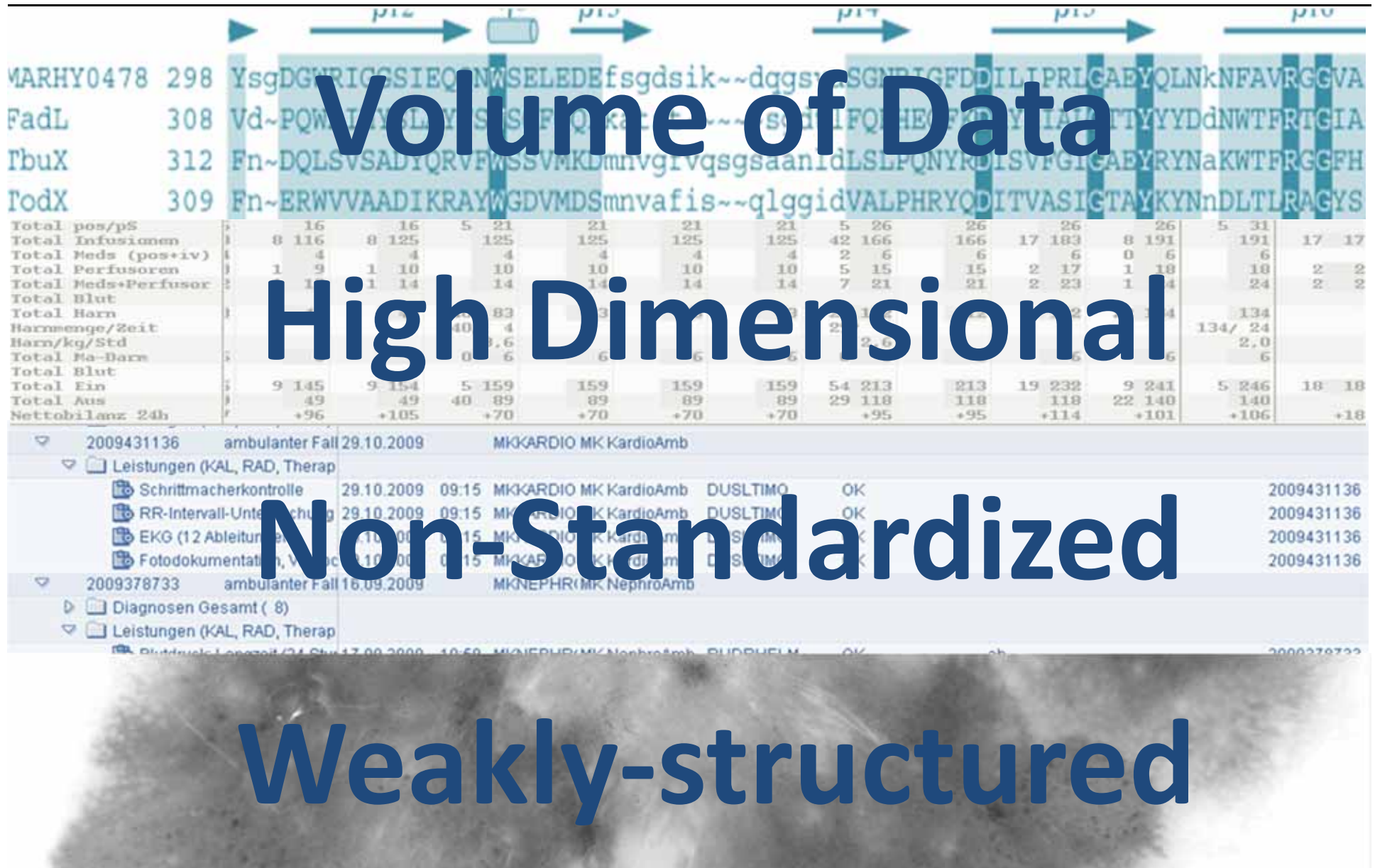
Holzinger, A. & Simonic, K.-M. (eds.) 2011. *Information Quality in e-Health*.

Lecture Notes in Computer Science LNCS 7058, Heidelberg, Berlin, New York: Springer.



Holzinger, A. & Simonic, K.-M. (Eds.) (2011) *Information Quality in e-Health. Lecture Notes in Computer Science LNCS 7058, Heidelberg, New York, Springer.*





Volume of Data

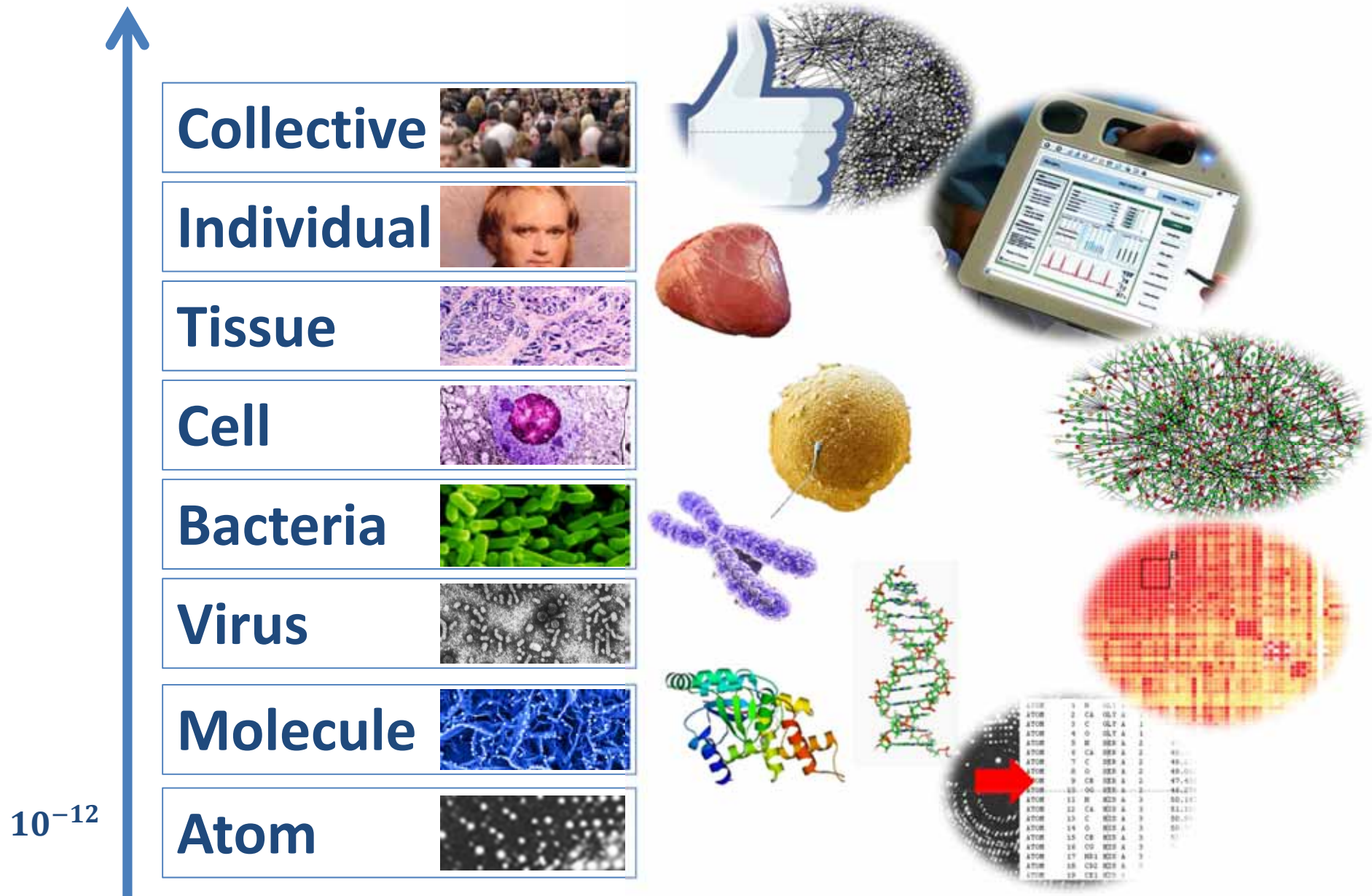
High Dimensional

Non-Standardized

Weakly-structured

Holzinger, A. (2011) Weakly Structured Data in Health-Informatics. In: *INTERACT 2011, Lisbon, IFIP*, 5-7.







Open Problems and Future Challenges

- 1. A unified controlled medical vocabulary (CMV);
- 2. A complete computer-based patient record that could serve as a regional/national/multinational resource and a format to allow exchange of records between systems;
- 3. The automatic coding of free-text reports, patient histories, discharge abstracts, etc.;
- 4. Automated analysis of medical records, yielding
 - a) the expected (most common) clinical presentation and course and the degree of clinical variability for patients with a given diagnosis;
 - b) the resources required in the care of patients compared by diagnosis, treatment protocol, clinical outcome, location, and physician;
- 5. A uniform, intuitive, anticipating user interface;
- 6. The human genome project;
- 7. A complete three-dimensional, digital representation of the body, including the brain, with graphic access to anatomic sections, etc.;
- 8. Techniques to ease the incorporation of new information management technologies into the infrastructure of organizations so that they can be used at the bedside or at the research bench;
- 9. A comprehensive, clinical decision support system.

- Grand new challenges from today's perspective include:
- 10. Closing the gap between Science and Practice
- 11. Data fusion and data integration in the clinical workplace
- 12. To provide a trade-off between Standardization and Personalization
- 13. An intuitive, unified and universal, adaptive and adaptable user interface
- 14. Integrated interactive Knowledge Discovery Methods particularly for the masses of still “unstructured data”
- 15. Mobile solutions for the bedside and the clinical bench
- A consequence of 14 and 15 will be the vision of “Watson” on the Smartphone. This goal was announced by IBM for the year 2020. The problem involved are the massive unstructured clinical data sets [1]

1. Holzinger, A., Stocker, C., Ofner, B., Prohaska, G., Brabenetz, A., & Hofmann-Wellenhof, R. (2013). Combining HCI, Natural Language Processing, and Knowledge Discovery - Potential of IBM Content Analytics as an assistive technology in the biomedical domain Springer Lecture Notes in Computer Science LNCS 7947 (pp. 13-24). Heidelberg, Berlin, New York: Springer.

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Books and Arts

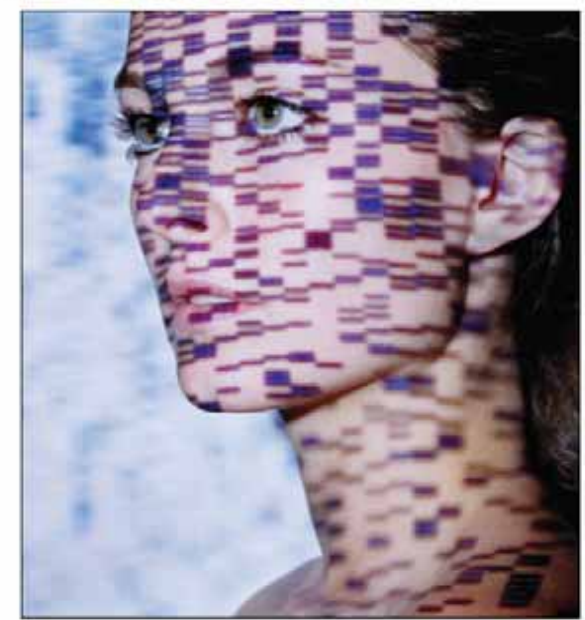
Nature **464**, 680 (1 April 2010) | doi:10.1038/464680a; Published online 31 March 2010

A reality check for personalized medicine

Muin J. Khoury¹, James Evans² & Wylie Burke³

Bringing genetic information into health care is welcome but its utility in the clinic needs to be rigorously reviewed, caution Muin J. Khoury, James Evans and Wylie Burke.

BOOK REVIEWED
Personal Genomics and Personalized Medicine
by Hamid Bolouri
Imperial College Press: 2010. 280 pp. £34

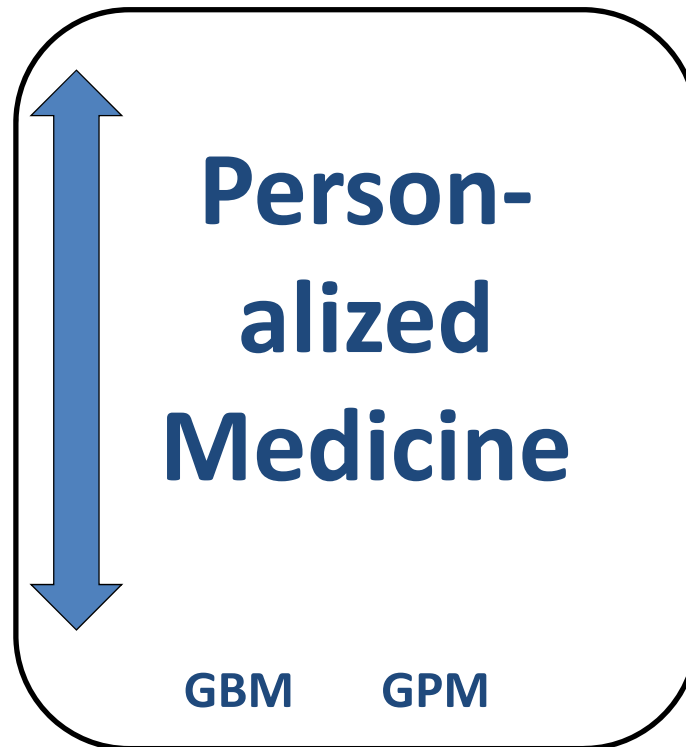


T. FLACH/STONE/GETTY

Genomic information: should it be treated in the same way as X-ray results?

EBM CPG

Standardized Medicine

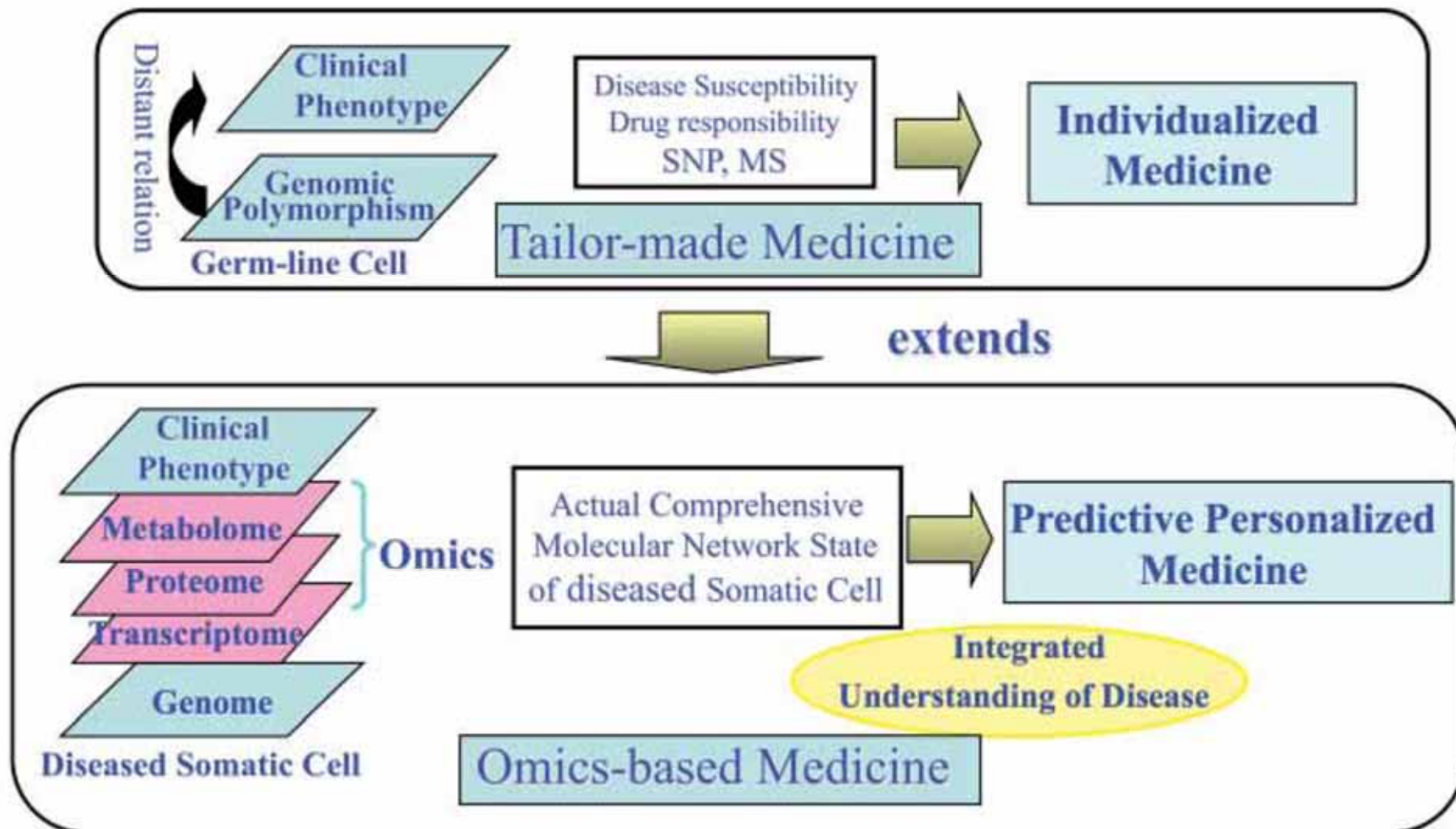


Pervasive Healthcare

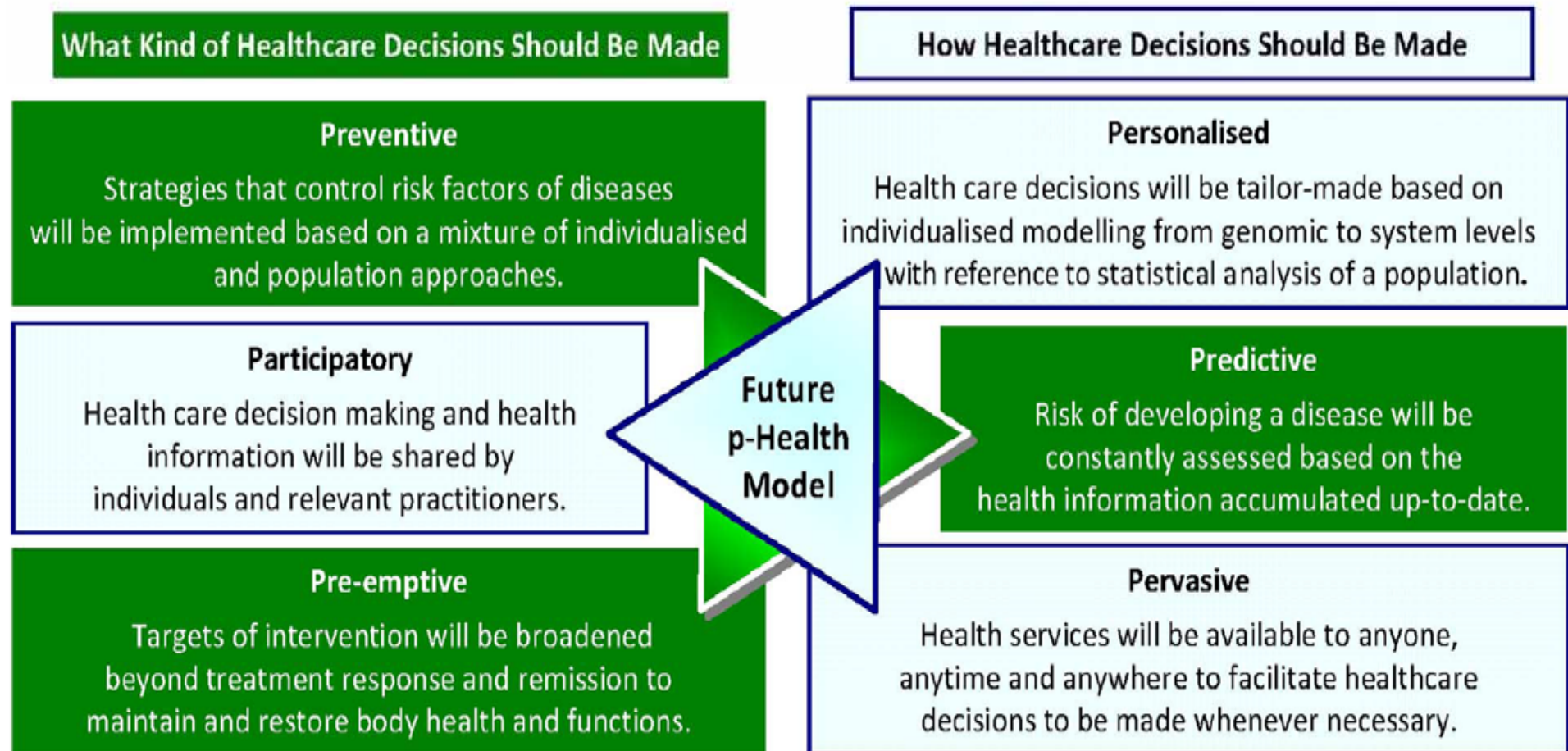
Preventive Health Integration

EBM = Evidence Based Medicine
CPG = Clinical Practice Guideline
GBM = Genome Based Medicine
GPM = Genetic Polymorphism

Tanaka, H. (2010)

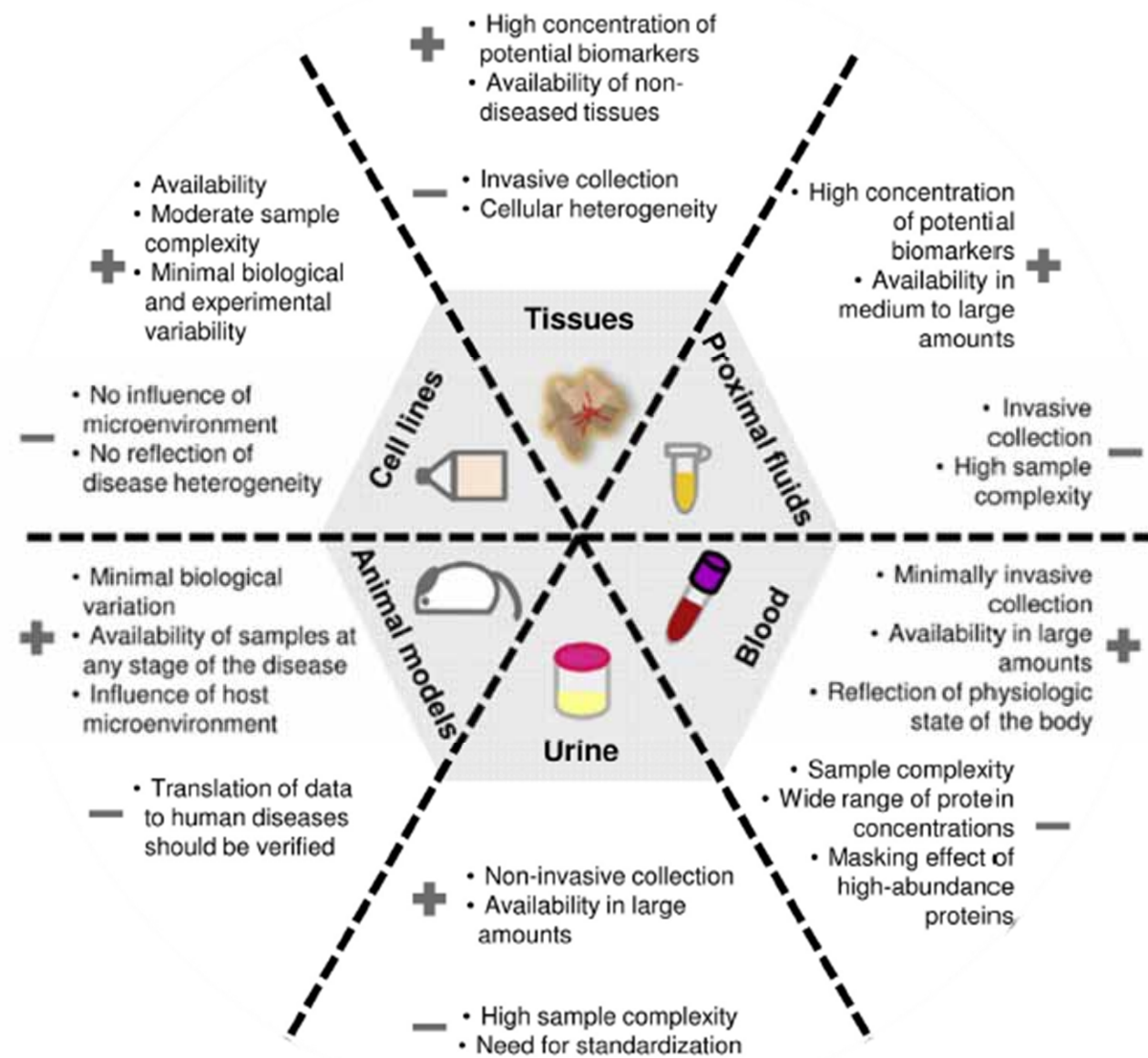


Tanaka, H. (2010) Omics-based Medicine and Systems Pathology A New Perspective for Personalized and Predictive Medicine. *Methods of Information In Medicine*, 49, 2, 173-185.



Zhang, Y. T. & Poon, C. C. Y. (2010) Editorial Note on Bio, Medical, and Health Informatics. *Information Technology in Biomedicine, IEEE Transactions on*, 14, 3, 543-545.

Drabovich, A. P., Pavlou, M. P., Batruch, I. & Diamandis, E. P. 2013. Chapter 2 - Proteomic and Mass Spectrometry Technologies for Biomarker Discovery. In: Haleem, J. I. & Timothy, D. V. (eds.) *Proteomic and Metabolomic Approaches to Biomarker Discovery*. Boston: Academic Press, pp. 17-37.



nature

www.nature.com/nature

458 | Issue no. 7234 | 5 March 2009

What price health?



Thank you!

08	Biomarkers are measured molecules which indicate the presence of an abnormal condition within a patient, and can be a gene (e.g., SNP), protein (e.g., prostate-specific antigen), or metabolite.	☐ Yes ☐ No	2 total
----	---	---	---------

06	☐ a) ... heterogeneity and weak <u>structurization</u> of the available data. Part of the definition of Biomedical Informatics is the ... ☐ ... effective use of biomedical data. ☐ ... motivation to improve computational capacities. ☐ ... effort to expand the technological capabilities. ☐ ... motivation to improve human health.	4 total
----	--	---------

02	The Von-Neumann Architecture is the fundamental computer organization structure of nearly all of our todays computing systems (e.g. in your PC, smartphone, microwave oven, car, etc.), please roughly sketch the Von Neumann Architecture and indicate the main parts:	1-28 1 each 6 total
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- Aebersold, R. & Mann, M. 2003. Mass spectrometry-based proteomics. *Nature*, 422, (6928), 198-207.
- Alagöz, F., Calero Valdez, A., Wilkowska, W., Ziefle, M., Dorner, S. & Holzinger, A. 2010. From Cloud Computing to Mobile Internet, From User Focus to Culture and Hedonism - The Crucible of Mobile Health Care and Wellness Applications. *ICPCA10 International Conference on Pervasive Computer Applications 2010*. Maribor (Slovenia): IEEE. 1-9.
- Anfinsen, C. B. 1973. Principles that Govern the Folding of Protein Chains. *Science*, 181, (4096), 223-230.
- Aral, S. 2011. Identifying Social Influence: A Comment on Opinion Leadership and Social Contagion in New Product Diffusion. *Marketing Science*, 30, (2), 217-223.
- Auinger, A., Ebner, M., Nedbal, D. & Holzinger, A. 2009. Mixing Content and Endless Collaboration – MashUps: Towards Future Personal Learning Environments. In: Stephanidis, C. (ed.) *Universal Access in Human-Computer Interaction HCI, Part III: Applications and Services, Lecture Notes in Computer Science LNCS 5616*. Berlin, Heidelberg, New York: Springer, pp. 14-23.
- Barabási, A. L., Gulbahce, N. & Loscalzo, J. 2011. Network medicine: a network-based approach to human disease. *Nature Reviews Genetics*, 12, (1), 56-68.
- Barabási, A. L. & Oltvai, Z. N. 2004. Network biology: understanding the cell's functional organization. *Nature Reviews Genetics*, 5, (2), 101-113.
- Bardram, J. E. Activity-Based Support for Mobility and Collaboration in Ubiquitous Computing. In: Baresi, L., ed. *Second International Conference on Ubiquitous Mobile Information and Collaboration Systems, 2004 Riga (Latvia)*. Springer, 169-184.
- Bianchi, D. W. 2012. From prenatal genomic diagnosis to fetal personalized medicine: progress and challenges. *Nat Med*, 18, (7), 1041-1051.
- Blum, B. I. & Duncan, K. 1990. *A History of Medical Informatics*, Reading (MA), Addison Wesley.
- Boal, D. 2012. *Mechanics of the Cell*, Cambridge University Press.
- Bobrow, D. G., Burchfiel, J. D., Murphy, D. L. & Tomlinson, R. S. 1972. TENEX, a paged time sharing system for the PDP-10. *Communications of the ACM*, 15, (3), 135-143.
- Boguski, M. S. & McIntosh, M. W. 2003. Biomedical informatics for proteomics. *Nature*, 422, (6928), 233-237.
- Boulton, G., Rawlins, M., Vallance, P. & Walport, M. 2011. Science as a public enterprise: the case for open data. *The Lancet*, 377, (9778), 1633-1635.
- Brooks, R. 2001. The relationship between matter and life. *Nature*, 409, (6818), 409-411.
- Brown, T. 2012. *Introduction to genetics: a molecular approach*, New York, Garland.
- Burgin, M. & Eberbach, E. 2012. Evolutionary computation and the processes of life: opening statement. *ACM Ubiquity*, 2012, (August), 1-13.
- Cavin, R., Lugli, P. & Zhirnov, V. 2012. Science and Engineering Beyond Moore's Law. *Proceedings of the IEEE*, 100, (13), 1720-1749.
- Chirikjian, G. S. 2011. Modeling loop entropy. *Methods in enzymology*, 487, 99.
- Cooper, S. B., Loewe, B. & Sorbi, A. 2008. *New Computational Paradigms: Changing Conceptions of What is Computable*, New York, Springer.
- Crick, F. 1970. Central Dogma of Molecular Biology. *Nature*, 227, (5258), 561-563.
- Crick, F. & Watson, J. 1953. Molecular structure of nucleic acids. *Nature*, 171, (4356), 737-738.
- Dill, K. A., Bromberg, S., Yue, K., Chan, H. S., Ftebig, K. M., Yee, D. P. & Thomas, P. D. 1995. Principles of protein folding—a perspective from simple exact models. *Protein Science*, 4, (4), 561-602.
- Drabovich, A. P., Pavlou, M. P., Batruch, I. & Diamandis, E. P. 2013. Chapter 2 - Proteomic and Mass Spectrometry Technologies for Biomarker Discovery. In: Haleem, J. I. & Timothy, D. V. (eds.) *Proteomic and Metabolomic Approaches to Biomarker Discovery*. Boston: Academic Press, pp. 17-37.
- Fisher, C., Lauría, E. & Chengalur-Smith, S. 2012. *Introduction to information quality*, Bloomington (IN), AuthorHouse.
- Ge, H., Walhout, A. J. M. & Vidal, M. 2003. Integrating 'omic' information: a bridge between genomics and systems biology. *TRENDS in Genetics*, 19, (10), 551-560.
- Gigerenzer, G. 2008. *Gut Feelings: Short Cuts to Better Decision Making* London, Penguin.
- Gromiha, M. M. 2010. *Protein Bioinformatics*, Amsterdam, Elsevier.
- Hawking, S. W., Penrose, R. & Atiyah, M. 1996. *The nature of space and time*, Princeton University Press Princeton.
- Hersey, A., Senger, S. & Overington, J. P. 2012. Open data for drug discovery: learning from the biological community. *Future Medicinal Chemistry*, 4, (15), 1865-1867.

- Holzinger, A. 2002. *Basiswissen IT/Informatik Band 1: Informationstechnik*. Wuerzburg, Vogel Buchverlag.
- Holzinger, A. 2003a. *Basiswissen IT/Informatik. Band 2: Informatik*, Wuerzburg, Vogel Buchverlag.
- Holzinger, A. 2003b. Finger Instead of Mouse: Touch Screens as a means of enhancing Universal Access. In: Carbonell, N. & Stephanidis, C. (eds.) *Universal Access: Theoretical Perspectives, Practice and Experience, Lecture Notes in Computer Science (LNCS 2615)* Berlin, Heidelberg, New York: Springer, pp. 387-397.
- Holzinger, A. 2013. Human-Computer Interaction and Knowledge Discovery (HCI-KDD): What is the benefit of bringing those two fields to work together? In: Cuzzocrea, A., Kittl, C., Simos, D. E., Weippl, E. & Xu, L. (eds.) *Multidisciplinary Research and Practice for Information Systems, Springer Lecture Notes in Computer Science LNCS 8127*. Heidelberg, Berlin, New York: Springer, pp. 319-328.
- Holzinger, A., Basic, L., Peischl, B. & Debevc, M. 2011. Handwriting Recognition on Mobile Devices: State of the art technology, usability and business analysis. *Proceedings of the 8th International Conference on electronic Business and Telecommunications*. Los Alamitos: IEEE, pp. 219-227.
- Holzinger, A., Dehmer, M. & Jurisica, I. 2014. Knowledge Discovery and interactive Data Mining in Bioinformatics - State-of-the-Art, future challenges and research directions. *BMC Bioinformatics*, 15, (S6), 11.
- Holzinger, A., Dorner, S., Födinger, M., Valdez, A. & Ziefle, M. 2010. Chances of Increasing Youth Health Awareness through Mobile Wellness Applications. In: Leitner, G., Hitz, M. & Holzinger, A. (eds.) *HCI in Work and Learning, Life and Leisure, Lecture Notes in Computer Science, LNCS 6389*. Berlin, Heidelberg: Springer, pp. 71-81.
- Holzinger, A., Emberger, W., Wassertheurer, S. & Neal, L. 2008. Design, Development and Evaluation of Online Interactive Simulation Software for Learning Human Genetics. *Elektrotechnik & Informationstechnik (e&i)*, 125, (5), 190-196.
- Holzinger, A., Kosec, P., Schwantzer, G., Debevc, M., Hofmann-Wellenhof, R. & Frühauf, J. 2011. Design and Development of a Mobile Computer Application to Reengineer Workflows in the Hospital and the Methodology to evaluate its Effectiveness. *Journal of Biomedical Informatics*, 44, (6), 968-977.
- Holzinger, A., Nischelwitzer, A., Friedl, S. & Hu, B. 2010. Towards life long learning: three models for ubiquitous applications. *Wireless Communications and Mobile Computing*, 10, (10), 1350-1365.
- Holzinger, A., Schaupp, K. & Eder-Halbedl, W. 2008. An Investigation on Acceptance of Ubiquitous Devices for the Elderly in an Geriatric Hospital Environment: using the Example of Person Tracking In: Miesenberger, K., Klaus, J., Zagler, W. & Karshmer, A. (eds.) *Computers Helping People with Special Needs, Lecture Notes in Computer Science, LNCS 5105*. Heidelberg, Berlin: Springer, pp. 22-29.
- Holzinger, A., Schlögl, M., Peischl, B. & Debevc, M. Preferences of Handwriting Recognition on Mobile Information Systems in Medicine: Improving handwriting algorithm on the basis of real-life usability research (Best Paper Award). ICE-B 2010 - ICETE The International Joint Conference on e-Business and Telecommunications, 2010 Athens (Greece). INSTICC, 14-21.
- Holzinger, A., Schwaberg, K. & Weitlaner, M. 2005. Ubiquitous Computing for Hospital Applications: RFID-Applications to enable research in Real-Life environments *29th Annual International Conference on Computer Software & Applications (IEEE COMPSAC)*. Edinburgh (UK): IEEE. 19-20. Holzinger, A., Searle, G., Peischl, B. & Debevc, M. 2012. An Answer to "Who needs a stylus?" On Handwriting Recognition on Mobile Devices. *e-Business and Telecommunications, Communications in Computer and Information Science, CCIS 314*. Heidelberg, Berlin, New York: Springer, pp. 156-167.
- Holzinger, A. & Simonik, K.-M. (eds.) 2011. *Information Quality in e-Health. Lecture Notes in Computer Science LNCS 7058*, Heidelberg, Berlin, New York: Springer.
- Holzinger, A., Stocker, C., Ofner, B., Prohaska, G., Brabenetz, A. & Hofmann-Wellenhof, R. 2013. Combining HCI, Natural Language Processing, and Knowledge Discovery - Potential of IBM Content Analytics as an assistive technology in the biomedical domain. *Springer Lecture Notes in Computer Science LNCS 7947*. Heidelberg, Berlin, New York: Springer, pp. 13-24.
- Holzinger, A., Stocker, C., Peischl, B. & Simonik, K.-M. 2012. On Using Entropy for Enhancing Handwriting Preprocessing. *Entropy*, 14, (11), 2324-2350.

- Hunter, L. 2009. *The Processes of Life: An introduction to molecular biology*, Cambridge (MA), MIT Press.
- Hurst, M. 2007. *Data Mining: Text Mining, Visualization and Social Media* [Online]. Available: http://datamining.typepad.com/data_mining/2007/01/the_blogosphere.html [Accessed 2011-05-10].
- Issaq, H. J. & Veenstra, T. D. 2013. Chapter 1 - Biomarker Discovery: Study Design and Execution. In: Haleem, J. I. & Timothy, D. V. (eds.) *Proteomic and Metabolomic Approaches to Biomarker Discovery*. Boston: Academic Press, pp. 1-16.
- Jeanquartier, F. & Holzinger, A. 2013. On Visual Analytics And Evaluation In Cell Physiology: A Case Study. In: Cuzzocrea, A., Kittl, C., Simos, D. E., Weippl, E. & Xu, L. (eds.) *Multidisciplinary Research and Practice for Information Systems LNCS 8127*. Heidelberg, Berlin: Springer, pp. 495-502.
- Jeong, H., Mason, S. P., Barabasi, A. L. & Oltvai, Z. N. 2001. Lethality and centrality in protein networks. *Nature*, 411, (6833), 41-42.
- Kiberstis, P. A. 2012. All Eyes on Epigenetics. *Science*, 335, (6069), 637.
- Klibanov, A. M. 2001. Improving enzymes by using them in organic solvents. *Nature*, 409, (6817), 241-246.
- Komaroff, A. L. 1979. The variability and inaccuracy of medical data. *Proceedings of the IEEE*, 67, (9), 1196-1207.
- Kuhn, K., Knoll, A., Mewes, H., Schwaiger, M., Bode, A., Broy, M., Daniel, H., Feussner, H., Gradinger, R. & Hauner, H. 2008. From Molecules to Populations. *Methods of Information in Medicine*, 47, (4), 283-295.
- Mahajan, V. & Schoeman, M. E. F. 1977. The use of computers in hospitals: an analysis of adopters and nonadopters. *Interfaces*, 95-107.
- Marth, J. D. 2008. A unified vision of the building blocks of life. *Nat Cell Biol*, 10, (9), 1015-1015.
- Mitchell, S., Spiteri, M. D., Bates, J. & Coulouris, G. Context-aware multimedia computing in the intelligent hospital. 2000 Denmark. ACM Press, 13-18.
- Mitreva, M. 2012. Structure, function and diversity of the healthy human microbiome. *Nature*, 486, 207-214.
- Mojzsis, S. J., Arrhenius, G., McKeegan, K. D., Harrison, T. M., Nutman, A. P. & Friend, C. R. L. 1996. Evidence for life on Earth before 3,800 million years ago. *Nature*, 384, (6604), 55-59.
- Molloy, J. C. 2011. The Open Knowledge Foundation: Open Data Means Better Science. *Plos Biology*, 9, (12).
- Moore, G. E. 1965. Cramming More Components Onto Integrated Circuits. *Electronics*, 38, (8), 114-117.
- Neumann, J. V. 1945. First Draft of a Report on the EDVAC. *University of Pennsylvania - Technical Report*, 49 pp.
- Ng, P. C., Murray, S. S., Levy, S. & Venter, J. C. 2009. An agenda for personalized medicine. *Nature*, 461, (7265), 724-726.
- Okumoto, S., Jones, A. & Frommer, W. B. 2012. Quantitative imaging with fluorescent biosensors. *Annual review of plant biology*, 63, 663-706.
- Patel, V. L., Kahol, K. & Buchman, T. 2011. Biomedical Complexity and Error. *Journal of Biomedical Informatics*, 44, (3), 387-389.
- Petz, G., Karpowicz, M., Fürschuß, H., Auinger, A., Štriteský, V. & Holzinger, A. 2013. Opinion Mining on the Web 2.0 – Characteristics of User Generated Content and Their Impacts. *Lecture Notes in Computer Science LNCS 7947*. Heidelberg, Berlin: Springer, pp. 35-46
- Petz, G., Karpowicz, M., Fürschuß, H., Auinger, A., Winkler, S., Schaller, S. & Holzinger, A. 2012. On Text Preprocessing for Opinion Mining Outside of Laboratory Environments. In: Huang, R., Ghorbani, A., Pasi, G., Yamaguchi, T., Yen, N. & Jin, B. (eds.) *Active Media Technology, Lecture Notes in Computer Science, LNCS 7669*. Berlin Heidelberg: Springer, pp. 618-629.
- Pevsner, J. 2009. *Bioinformatics and functional genomics*, Hoboken (NJ), John Wiley & Sons.
- Rabilloud, T., Chevallet, M., Luche, S. & Lelong, C. 2010. Two-dimensional gel electrophoresis in proteomics: past, present and future. *Journal of proteomics*, 73, (11), 2064-2077.
- Rouwkema, J., Rivron, N. C. & Van Blitterswijk, C. A. 2008. Vascularization in tissue engineering. *Trends in biotechnology*, 26, (8), 434-441.

- Rowen, L., Wong, G. K. S., Lane, R. P. & Hood, L. 2000. Intellectual property - Publication rights in the era of open data release policies. *Science*, 289, (5486), 1881-1881.
- Sackett, D. L., Rosenberg, W. M., Gray, J., Haynes, R. B. & Richardson, W. S. 1996. Evidence based medicine: what it is and what it isn't. *BMJ: British Medical Journal*, 312, (7023), 71.
- Schidlowski, M. 1988. A 3,800-million-year isotopic record of life from carbon in sedimentary rocks. *Nature*, 333, (6171), 313-318.
- Schotte, F., Lim, M. H., Jackson, T. A., Smirnov, A. V., Soman, J., Olson, J. S., Phillips, G. N., Wulff, M. & Anfinrud, P. A. 2003. Watching a protein as it functions with 150-ps time-resolved X-ray crystallography. *Science*, 300, (5627), 1944-1947.
- Schrödinger, E. 1944. *What Is Life? The Physical Aspect of the Living Cell*, Dublin, Dublin Institute for Advanced Studies at Trinity College.
- Shadbolt, N., O'hara, K., Berners-Lee, T., Gibbins, N., Glaser, H., Hall, W. & Schraefel, M. 2012. Open Government Data and the Linked Data Web: Lessons from data. gov. uk. *IEEE Intelligent Systems*, 16-24.
- Shehu, A. & Kavraki, L. E. 2012. Modeling structures and motions of loops in protein molecules. *Entropy*, 14, (2), 252-290.
- Shortliffe, E. H. 2011. Biomedical Informatics: Defining the Science and its Role in Health Professional Education. In: Holzinger, A. & Simonik, K.-M. (eds.) *Information Quality in e-Health. Lecture Notes in Computer Science LNCS 7058*. Heidelberg, New York: Springer, pp. 711-714.
- Simonik, K.-M. & Holzinger, A. 2010. Zur Bedeutung von Information in der Medizin. *OCG Journal*, 35, (1), 8.
- Sittig, D. F. 1994. Grand Challenges in Medical Informatics. *Journal of the American Medical Informatics Association*, 1, (5), 412-413.
- Southern, E. M. 1975. Detection of Specific Sequences among DNA fragments separated by Gel-Electrophoresis. *Journal of Molecular Biology*, 98, (3), 503-&.
- Sperelakis, N. 2012. *Cell Physiology Sourcebook: Essentials of Membrane Biophysics. Fourth Edition*, Amsterdam, Elseviere.
- Steinbuch, K. 1957. Informatik: Automatische Informationsverarbeitung. *Standard Electric Lorenz (SEL) Nachrichten*, (4), 171.
- Stelzl, U., Worm, U., Lalowski, M., Haenig, C., Brembeck, F. H., Goehler, H., Stroedicke, M., Zenkner, M., Schoenherr, A., Koeppen, S., Timm, J., Mintzlaff, S., Abraham, C., Bock, N., Kietzmann, S., Goedde, A., Toksöz, E., Droege, A., Krobitch, S., Korn, B., Birchmeier, W., Lehrach, H. & Wanker, E. E. 2005. A Human Protein-Protein Interaction Network: A Resource for Annotating the Proteome. *Cell*, 122, (6), 957-968.
- Tanaka, H. 2010. Omics-based Medicine and Systems Pathology A New Perspective for Personalized and Predictive Medicine. *Methods of Information In Medicine*, 49, (2), 173-185.
- Thompson, M. & Heneghan, C. 2012. BMJ Open Data Campaign: We need to move the debate on open clinical trial data forward. *British Medical Journal*, 345.
- Trent, R. J. 2012. *Molecular Medicine: Genomics to Personalized Healthcare. 4th Edition*, Amsterdam, Elsevier.
- Trygve, T. 2011. *Handbook of Epigenetics*, San Diego, Academic Press.
- Varshney, U. 2009. *Pervasive Healthcare Computing: EMR/EHR, Wireless and Health Monitoring*, New York, Springer.
- Walsh, J. E. 1960. Analyzing Medical Data: Some Statistical Considerations. *Medical Electronics, IRE Transactions on*, ME-7, (4), 362-366.
- Wassertheurer, S., Holzinger, A., Emberger, W. & Breitenecker, F. 2003. Design and Development of Interactive Online-Simulations for e-Learning. In: Hohmann, R. (ed.) *Frontiers in Simulation - 17th Symposium Simulationstechnik in Magdeburg (Germany)*. Delft: SCS European Publication, pp. 97-105.
- Weiser, M. 1991. The computer for the twenty-first century. *Scientific American*, 265, (3), 94-104.

- Westra, R., Tuyls, K., Saeys, Y. & Nowé, A. 2007. Knowledge Discovery and Emergent Complexity in Bioinformatics. *In: Tuyls, K., Westra, R., Saeys, Y. & Nowé, A. (eds.) Knowledge Discovery and Emergent Complexity in Bioinformatics*. Springer Berlin Heidelberg, pp. 1-9.
- Wiltgen, M. & Holzinger, A. 2005. Visualization in Bioinformatics: Protein Structures with Physicochemical and Biological Annotations. *In: Zara, J. & Sloup, J. (eds.) Central European Multimedia and Virtual Reality Conference (available in EG Eurographics Library)*. Prague: Czech Technical University (CTU), pp. 69-74.
- Wiltgen, M., Holzinger, A. & Tilz, G. P. 2007. Interactive Analysis and Visualization of Macromolecular Interfaces Between Proteins. *In: Holzinger, A. (ed.) HCI and Usability for Medicine and Health Care. Lecture Notes in Computer Science (LNCS 4799)*. Berlin, Heidelberg, New York: Springer, pp. 199-212.
- Xiao, W. Z. & Oefner, P. J. 2001. Denaturing high-performance liquid chromatography: A review. *Human Mutation*, 17, (6), 439-474.
- Yapijakis, C. 2009. Hippocrates of Kos, the Father of Clinical Medicine, and Asclepiades of Bithynia, the Father of Molecular Medicine. *In Vivo*, 23, (4), 507-514.
- Yildirim, P., Ekmekci, I. & Holzinger, A. 2013. On Knowledge Discovery in Open Medical Data on the Example of the FDA Drug Adverse Event Reporting System for Alendronate (Fosamax). *In: Holzinger, A. & Pasi, G. (eds.) Human-Computer Interaction and Knowledge Discovery in Complex, Unstructured, Big Data, Lecture Notes in Computer Science, LNCS 7947*. Berlin Heidelberg: Springer, pp. 195-206.
- Yip, L. Y. & Yong Chan, E. C. 2013. Chapter 8 - Gas Chromatography/Mass Spectrometry-Based Metabonomics. *In: Haleem, J. I. & Timothy, D. V. (eds.) Proteomic and Metabolomic Approaches to Biomarker Discovery*. Boston: Academic Press, pp. 131-144.
- Zhang, Y. T. & Poon, C. C. Y. 2010. Editorial Note on Bio, Medical, and Health Informatics. *Information Technology in Biomedicine, IEEE Transactions on*, 14, (3), 543-545.
- Ziefle, M., Röcker, C. & Holzinger, A. 2011. Medical Technology in Smart Homes: Exploring the User's Perspective on Privacy, Intimacy and Trust. *35th Annual IEEE Computer Software and Applications Conference Workshops COMPSAC 2011*. Munich: IEEE, pp. 410-415.