

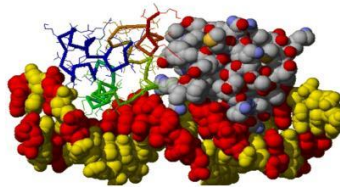
PERAN BIOINFORMATIKA DALAM PENINGKATAN PENELITIAN DAN PEMBELAJARAN BIOLOGI DI ERA REVOLUSI INDUSTRI 4.0

Prof. Widodo, S.Si., M.Si., Ph.D.Med.Sc.

Guru Besar dari Universitas Brawijaya

E-mail: widodo@ub.ac.id

**Peran Bioinformatika
dalam Peningkatan Penelitian dan Pembelajaran Biologi
di Era Revolusi Industri 4.0**



Prof. Widodo, S.Si., M.Si., Ph.D.Med.Sc.

Guru Besar dari Universitas Brawijaya

widodo@ub.ac.id

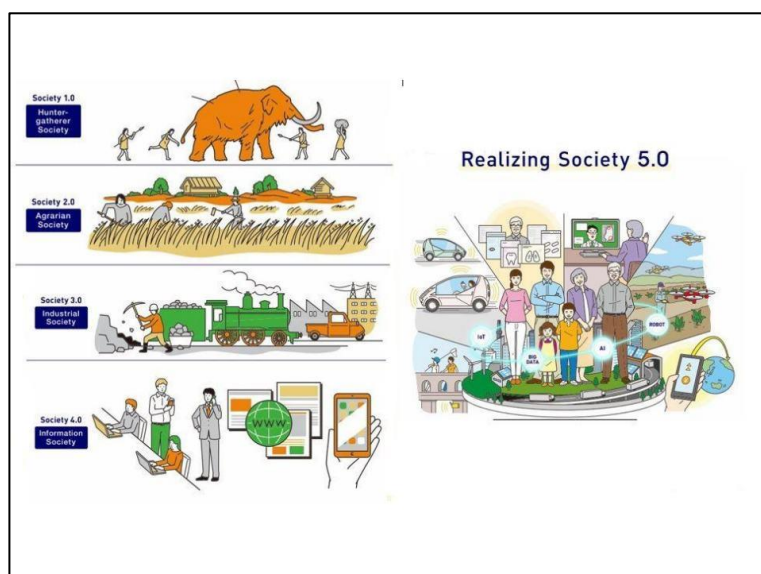
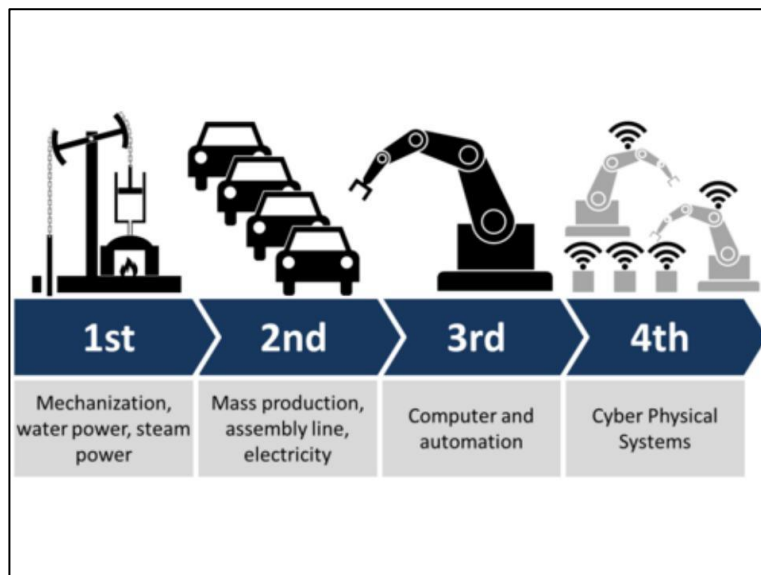
Latarbelakang

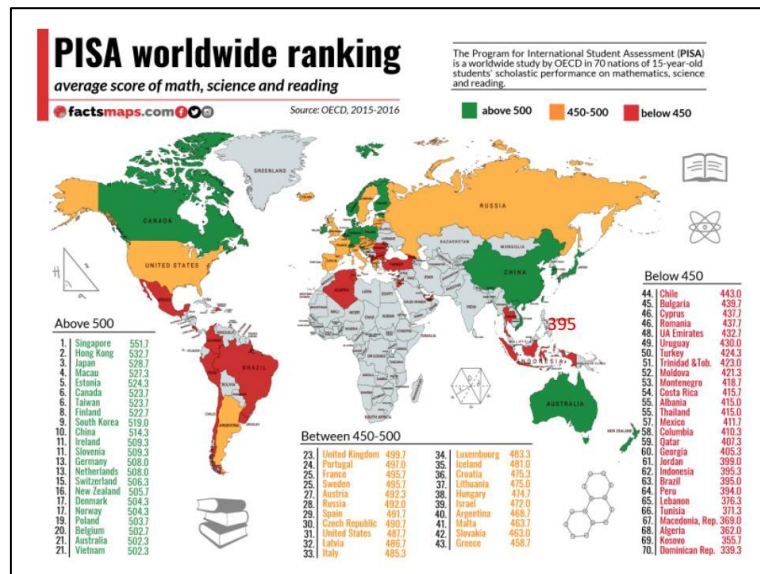
- Istilah "Industrie 4.0" berasal dari sebuah proyek dalam strategi teknologi canggih [pemerintah Jerman](#) yang mengutamakan [komputerisasi](#) pabrik.^[5]
- Istilah "Industrie 4.0" diangkat kembali di [Hannover Fair](#) tahun 2011.^[6]
- Pada Oktober 2012, Working Group on Industry 4.0 memaparkan rekomendasi pelaksanaan Industri 4.0 kepada pemerintah federal Jerman.
- Anggota kelompok kerja Industri 4.0 diakui sebagai bapak pendiri dan perintis Industri 4.0.
- Laporan akhir Working Group Industry 4.0 dipaparkan di Hannover Fair tanggal 8 April 2013.^[7]

https://id.wikipedia.org/wiki/Industri_4.0

Prinsip Industri 4.0:

- **interconnection,**
- **information transparency,**
- **technical assistance,**
- **decentralized decisions.**





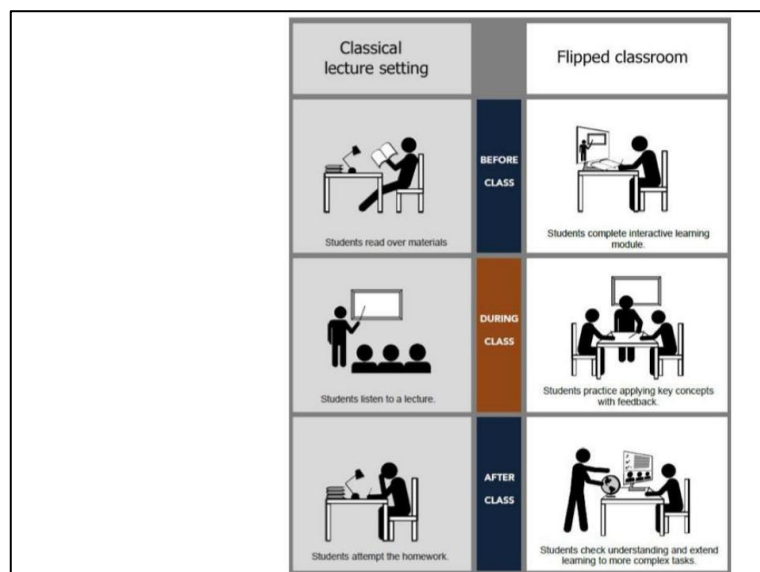
Tantangan yang dihadapi oleh Pendidikan dalam pengembangan kurikulum di era Revolusi Industri 4.0 dan society 5.0 adalah:

Menghasilkan lulusan yang memiliki kemampuan literasi baru meliputi

1. literasi data,
2. literasi teknologi
3. literasi manusia (HUMANISASI)
4. kemampuan memandang tanda- tanda akan terjadinya revolusi industri 5.0.

Capaian Pembelajaran Lulusan, CPL yang diperlukan di era industri 4.0 :

1. literasi data, kemampuan pemahaman untuk membaca, menganalisis, menggunakan data dan informasi (big data) di dunia digital;
2. literasi teknologi, kemampuan memahami cara kerja mesin, aplikasi teknologi (coding, artificial intelligence, dan engineering principle);
3. literasi manusia, kemampuan pemahaman tentang humanities, komunikasi dan desain; pemahaman akan tanda-tanda revolusi industri 4.0; pemahaman ilmu untuk diamalkan bagi kemaslahatan bersama secara lokal, nasional, dan global.
4. kemampuan memandang tanda- tanda akan terjadinya revolusi industri 5.0. Revolusi industri 5.0 dapat dipahami sebagai pasar kolaborasi manusia dengan sistem cerdas yang berbasis pada internet of things (IoT) atau sistem fisik cyber, dengan kemampuan memanfaatkan mesin-mesin cerdas lebih efisien dengan lingkungan yang lebih bersinergi (Rada, 2017).





Dimana peran Bioinformatic?

Manfaat Pendidikan Bioinformatik

1. Literasi data-BigDATA
2. Data Analyst
3. Literasi teknologi, coding & AI

AKSES DATABASE



DATA BASE :



DNA



PROTEIN



SENYAWA
KIMIA



SENYAWA
AKTIF JAMU



TAKSONOMI

SKILL



CORE
BIOINFORMATICS



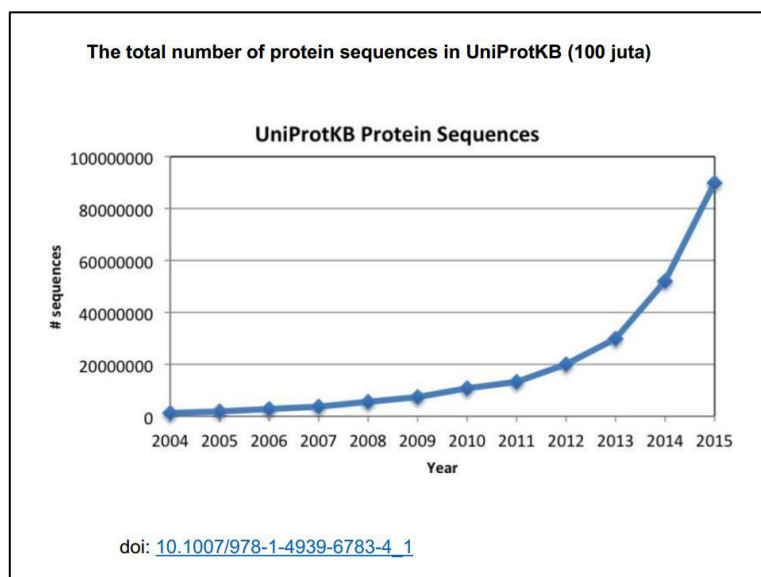
APPLICATION



DATA SCIENCE
& SOFTWARE DESIGN



PROFILES FROM
OTHER DOMAINS



Overview of Protein Bioinformatics Databases				
Category	DB Short Name	DB Name	URLs	Ref.
Sequence databases	CCDS	The Consensus CDS protein set database	https://www.ncbi.nlm.nih.gov/CCDS/CcdBrowse.cgi	[9]
	DDBJ	DNA Data Bank of Japan	http://www.ddbj.nig.ac.jp/	[10]
	ENA	European Nucleotide Archive	http://www.ebi.ac.uk/ena	[11]
	GenBank	GenBank nucleotide sequence database	https://www.ncbi.nlm.nih.gov/genbank/	[12]
	RefSeq [*]	NCBI Reference Sequence Database	https://www.ncbi.nlm.nih.gov/refseq/	[13]
	UniGene	Database of computationally identifies transcripts from the same locus	https://www.ncbi.nlm.nih.gov/uniGene/	[12]
2D gel databases	UniProtKB [*]	Universal Protein Resource (UniProt)	http://www.uniprot.org/	[14]
	COMPLUYEAST-2DPAGE	2-DE database at Universidad Complutense de Madrid, Spain	http://complueast2dpag.dacya.ucm.es/	[15]
	REPRODUCTION-2DPAGE	2-DE database at Nanjing Medical University, China	http://reprod.njmu.edu.cn/cgi-bin/2d/2d.cgi	[16]
	SWISS-2DPAGE	2-DE database at Swiss Institute of Bioinformatics, Switzerland	http://world-2dpag.expasy.org/swiss-2dpag/	[17]
	World-2DPAGE [*]	The World-2DPAGE database	http://world-2dpag.expasy.org/repository/	[18]
3D structure databases	DisProt	Database of Protein Disorder	http://www.disprot.org/	[19]
	MobiDB	Database of intrinsically disordered and mobile proteins	http://mobidb.bio.unipd.it/	[20]
	ModBase	Database of Comparative Protein Structure Models	http://modbase.compbio.ucsf.edu/modbase.cgi/index.cgi	[21]
	PDBe [*]	Protein Data Bank at Europe	http://www.ebi.ac.uk/pdbe/	[22]
	PDBj [*]	Protein Data Bank at Japan	http://pd bj.org/	[23]
	PDBsum	Pictorial database of 3D structures in the Protein Data Bank	http://www.ebi.ac.uk/pdbsum/	[24]

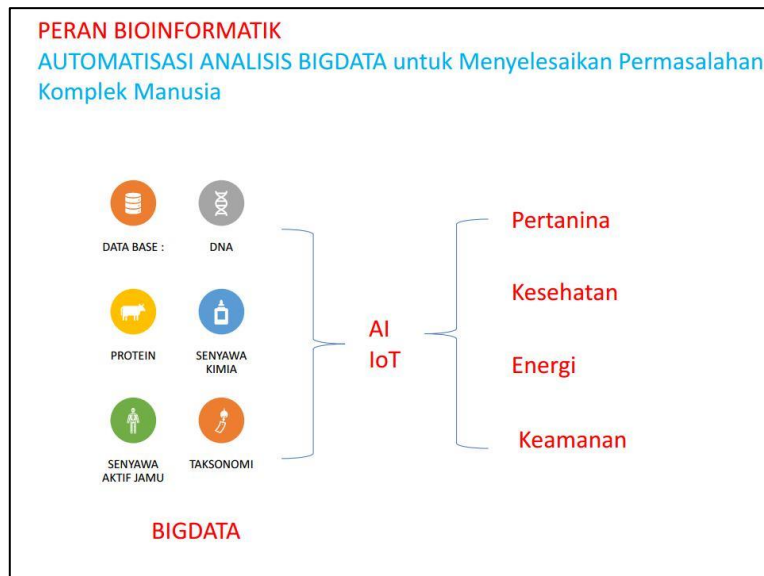
Chemistry databases	BindingDB	The Binding Database	http://www.bindingdb.org/	[28]
	ChEMBL [*]	Database of bioactive drug-like small molecules	https://www.ebi.ac.uk/chembl/	[29]
	DrugBank	Drug and Drug Target Database	http://www.drugbank.ca/	[30]
Enzyme and pathway databases	MetaCyc/BioCyc [*]	MetaCyc Database of Metabolic Pathways, BioCyc Collection of Pathway/Genome Databases	http://www.biocyc.org/	[31]
	BRENDA [*]	BRAunschweig ENzyme DAtabase	http://www.brenda-enzymes.org	[32]
	ENZYME	Enzyme nomenclature database	http://enzyme.expasy.org/	[33]
	Reactome [*]	A knowledgebase of biological pathways and processes	http://www.reactome.org/	[34]
	SABIO-RK	SABIO-RK: Biochemical Reaction Kinetics Database	http://sabiork.h-its.org/	[35]
	SignalLink	A signaling pathway resource with multi-layered regulatory networks	http://signalink.org/	[36]
	UniPathway	UniPathway: a resource for the exploration of metabolic pathways	http://www.unipathway.org	[37]
	Gene3D	Structural and Functional Annotation of Protein Families	http://gene3d.biochem.ucl.ac.uk/Gene3D/	[38]
Family and domain databases	HAMAP	High-quality Automated and Manual Annotation of Proteins	http://hamap.expasy.org/	[39]
	InterPro [*]	Integrated resource of protein families, domains and functional sites	http://www.ebi.ac.uk/interpro/	[40]
	PANTHER	The PANTHER Classification System	http://www.pantherdb.org/	[41]
	Pfam [*]	The Pfam protein families database	http://pfam.xfam.org/	[42]
	PIRSF [*]	A whole-protein classification database	http://pir.georgetown.edu/pirwww/dbinfo/pirsf.shtml	[43]
	PRINTS	Protein Motif fingerprint database	http://www.bioinf.manchester.ac.uk/dbbrowser/PRINTS/	[44]
	ProDom	Protein domain families database	http://prodom.prabi.fr/prodom/current/html/home.php	[45]



Gene expression databases	Bgee	Database for Gene Expression Evolution	http://bgee.unil.ch	[51]
	CleanEx	Database of gene expression profiles	http://cleanex.vital-it.ch/	[52]
	Genevisible	Search portal to normalized and curated expression data from Genevestigator	http://genevisible.com/search	[53]
	ExpressionAtlas [*]	Database of Differential and Baseline Expression	http://www.ebi.ac.uk/gxa/home	[54]
Genome annotation databases	Ensembl [*]	Ensembl Eukaryotic genome annotation database	http://www.ensembl.org/	[55]
	EnsemblBacteria	Ensembl Bacteria genome annotation database	http://bacteria.ensembl.org/	[56]
	EnsemblFungi	Ensembl Fungi genome annotation database	http://fungi.ensembl.org/	[56]
	EnsemblMetazoa	Ensembl Metazoa genome annotation database	http://metazoa.ensembl.org/	[56]
	EnsemblPlants	Ensembl Plants genome annotation database	http://plants.ensembl.org/	[56]
	EnsemblProtists	Ensembl Protists genome annotation database	http://protists.ensembl.org/	[56]
	Entrez Gene [*]	Database of Genes of Genomes in the Reference Sequence Collection	https://www.ncbi.nlm.nih.gov/gene	[57]
	KEGG	Kyoto Encyclopedia of Genes and Genomes	http://www.genome.jp/kegg/	[58]
	PATRIC	Bacterial Bioinformatics Resource Center	http://patricbrc.org/	[59]
	UCSC [*]	UCSC Genome Bioinformatics	http://genome.ucsc.edu	[60]
Organism specific databases	VectorBase	Bioinformatics Resource for Invertebrate Vectors of Human Pathogens	http://www.vectorbase.org/	[61]
	WBParaSite	WormBase ParaSite	http://parasite.wormbase.org	[62]
	ArachnoServer	ArachnoServer: Spider toxin database	http://www.arachnoserver.org	[63]
	CGD	Candida Genome Database	http://www.candidagenome.org/	[64]
	ConoServer	ConoServer: Cone snail toxin database	http://www.conoserver.org/	[65]
	CTD	Comparative Toxicogenomics Database	http://ctdbase.org/	[66]
	dictyBase	Central resource for Dictyostelid genomics	http://dictybase.org/	[67]
	EchoBASE	EchoBASE – an integrated post-genomic database for E. coli.	http://www.york.ac.uk/res/thomas/	[68]

Phylogenomic databases	eggNOG	Database of orthologous groups and functional annotation	http://eggnog.embl.de/	[98]
	HOGONOM	Database of Homologous Genes from Fully Sequenced Organisms	http://pbil.univ-lyon1.fr/databases/hogonom/home.php	[99]
	HOVERGEN	Homologous Vertebrate Genes Database	http://pbil.univ-lyon1.fr/databases/hovergen.html	[100]
	InParanoid	Eukaryotic Ortholog Groups with inparalogs	http://inparanoid.sbc.su.se/	[101]
	KO	Kyoto Encyclopedia of Genes and Genomes Orthology	http://www.genome.jp/kegg/	[102]
	OMA [*]	The OMA orthology database	http://omabrowser.org/	[103]
	OrthoDB	Database of Orthologous Groups	http://cegg.unige.ch/orthodb6	[104]
	PhylomeDB	Database for complete catalogs of gene phylogenies (phylogenies)	http://phylomedb.org/	[105]
Polymorphism and mutation databases	TreeFam	Database of animal gene trees	http://www.treefam.org	[106]
	BioMuta	Single-nucleotide variation and disease association database	https://hive.biochemistry.gwu.edu/tools/biomuta/	[107]
	dbSNP [*]	Database of Short Genetic Variations	https://www.ncbi.nlm.nih.gov/SNP/	[12]
	DMDM	Domain Mapping of Disease Mutations	http://bioinf.umbc.edu/dmdm/	[108]
Protein-protein interaction databases	BioGRID	The Biological General Repository for Interaction Datasets	http://thebiogrid.org	[109]
	DIP	Database of Interacting Proteins	http://dip.doe-mbi.ucla.edu/	[110]
	IntAct [*]	IntAct Molecular Interaction Database	http://www.ebi.ac.uk/intact/	[111]
	MINT	The Molecular INteraction database	http://mint.bio.uniroma2.it/mint/	[112]
	STRING	Search Tool for the Retrieval of Interacting Genes/Proteins	http://string-db.org	[113]
Proteomic databases	MaxQB	The MaxQuant DataBase	http://maxqb.biochem.mpg.de/mxdb/	[114]
	PaxDb	Protein Abundance Across Organisms	http://pax-db.org	[115]

 CORE BIOINFORMATICS	 APPLICATION	 DATA SCIENCE & SOFTWARE DESIGN	 PROFILES FROM OTHER DOMAINS
- Bridge the gap between biology and computer science - Typically have Master's degrees in bioinformatics - Sound understanding of biology and biological data, and experts in translating and interpreting large amounts of biological data into valuable insights with help from advanced mathematical, statistical and computational techniques	- Variety of educational backgrounds such as molecular biology, biochemistry or biotechnology - Apply standard bioinformatics software tools to analyse and solve scientific problems relevant to their field - Basic understanding of programming and algorithm design which supports their data analysis and interpretation	- Educational background within computer science, but do not necessarily have any knowledge within biology - Know how to organise, maintain and store large and complex data sets in a safe manner - Can create sound and user-friendly software tools for data analysis	- Profiles from other domains applicable to bioinformatics have an educational background within areas such as physics, statistics or mathematics - Solid experience with how to find mathematical and statistical answers and solutions to highly complex research questions based on large datasets

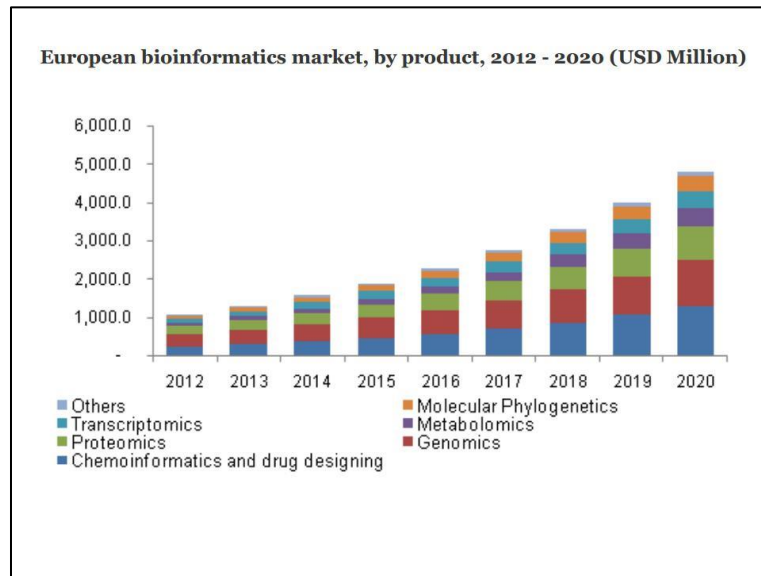


Peran Bioinformatik untuk Penelitian

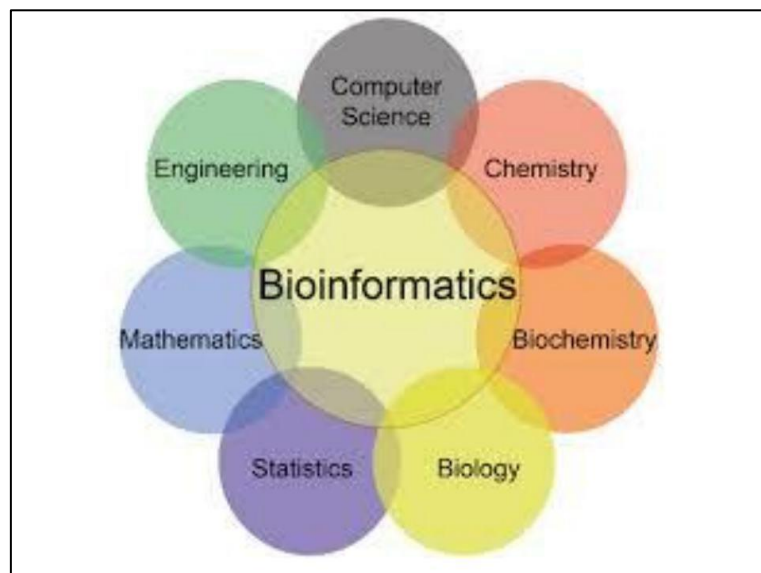
1. *Tool* untuk memahami data komplek
2. *Melahirkan* Pengetahuan interdisiplin
3. Speed up teknologi baru

Role of Bioinformatics in Biotechnology

- Climate Change Studies
- Waste Cleanup
- Bioenergy/Biofuels
- Biodefense
- Forensic Science
- Veterinary Science
- Crop Improvement
- Evolutionary Studies/Phylogenetics
- Drug Discovery
- Cheminformatics
- Transcriptomics
- Proteomics
- Genomics

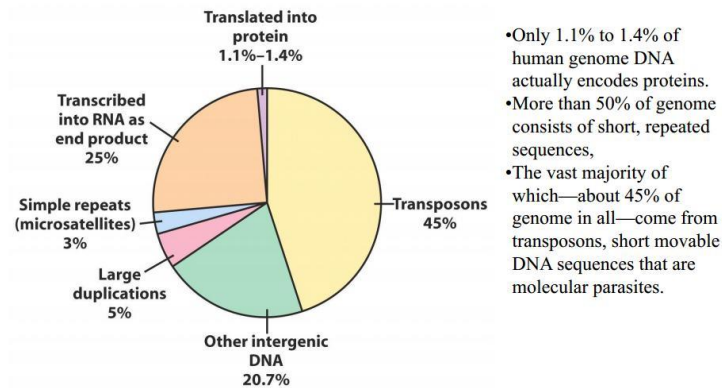


APA itu Bioinformatik ?



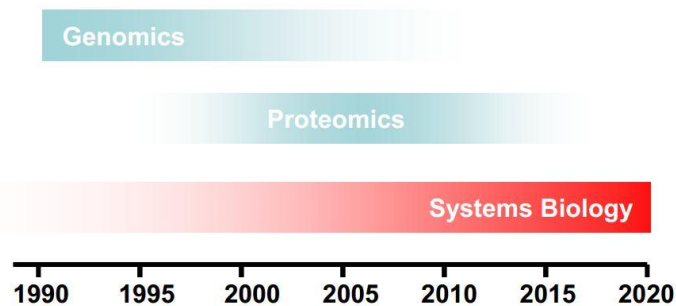


Snapshot of the human genome



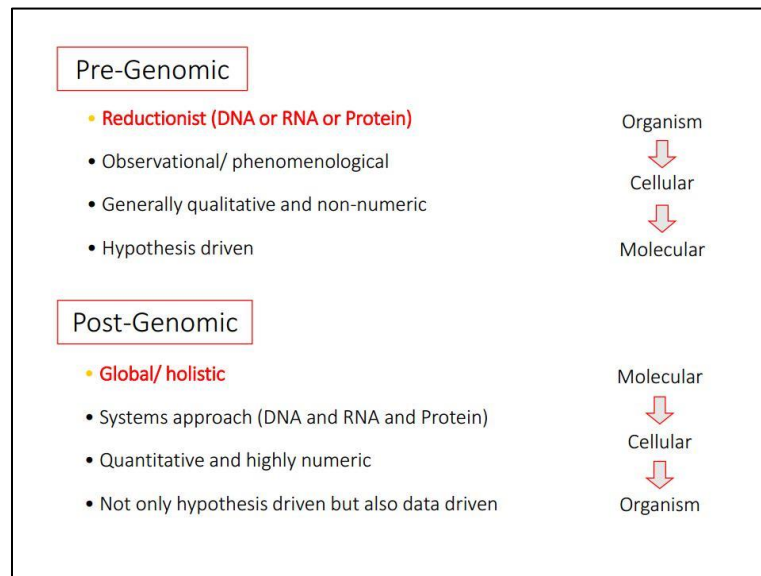
27

Genomics, Proteomics & Systems Biology



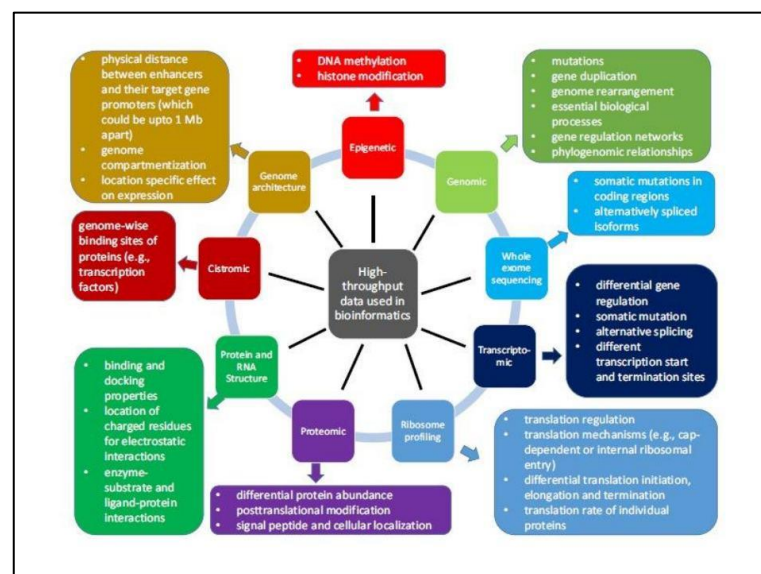
What is Systems Biology?

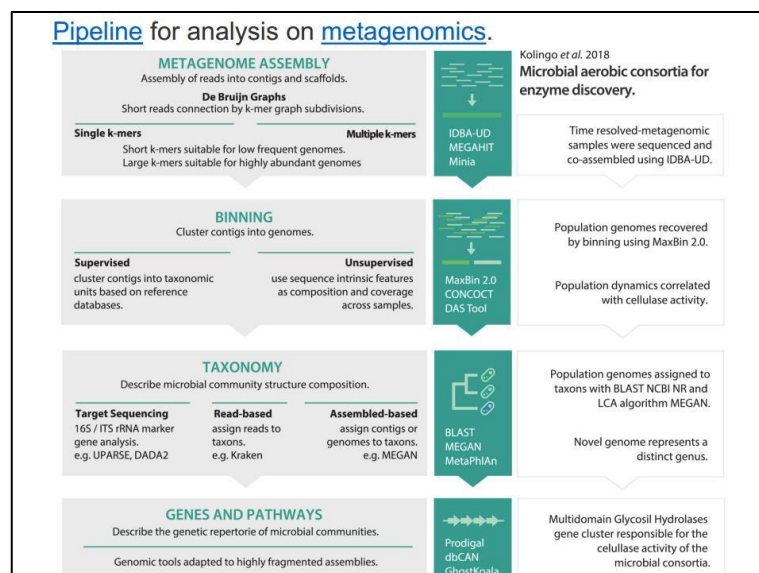
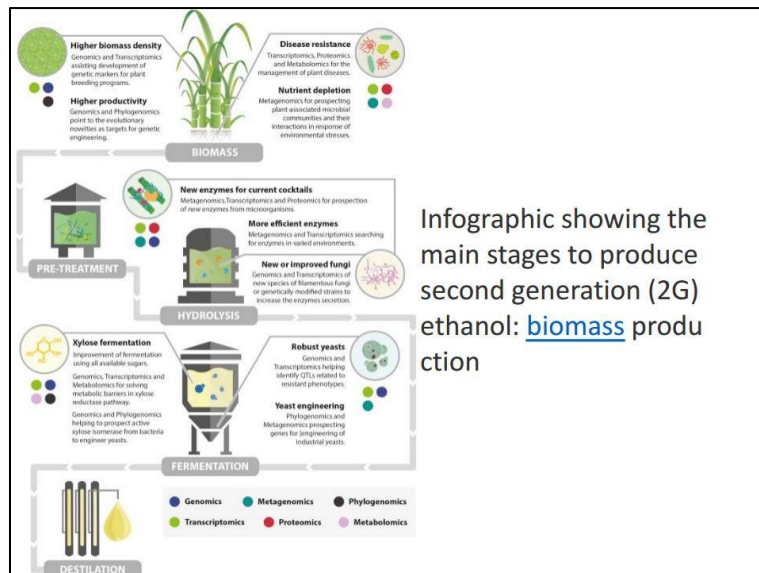
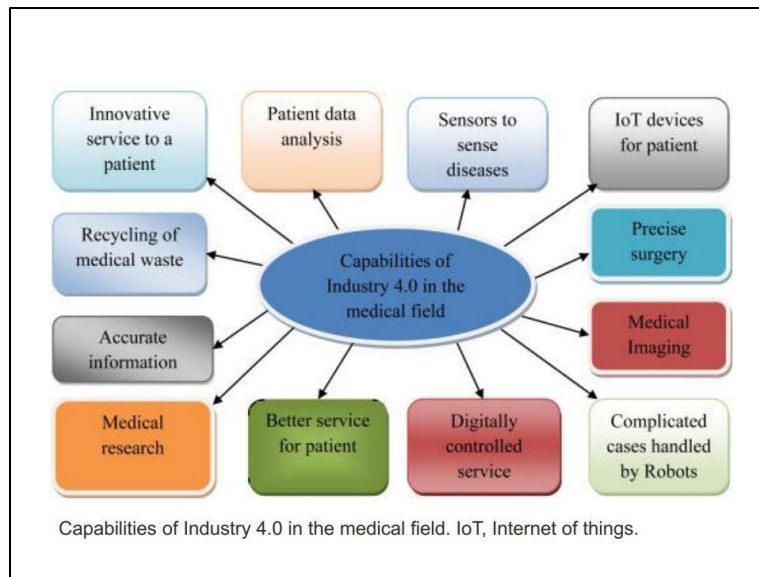
- **Systems Biology** - Systems biology is the study of an organism, viewed as an *integrated* and *interacting network* of genes, proteins and biochemical reactions. The integration of genomics, proteomics, metabolomics & modeling
- **The Goal:** *Predictive, Preventative and Personalized Medicine, precision agriculture*

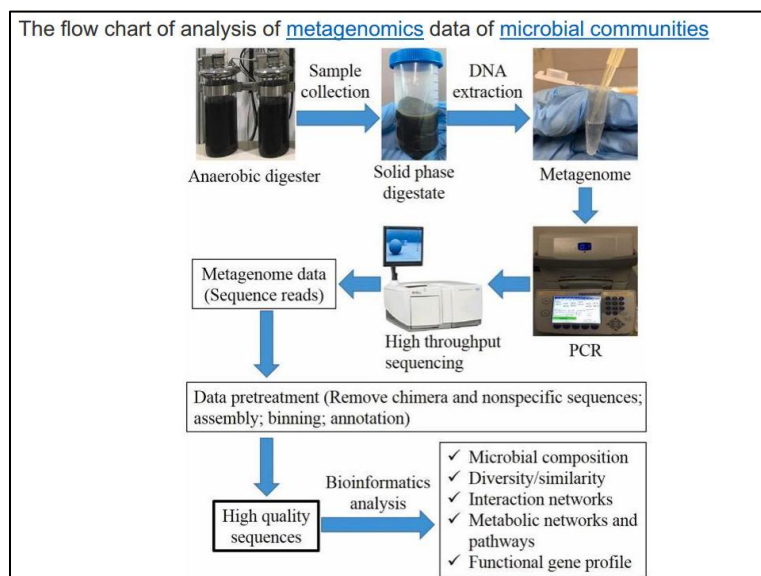
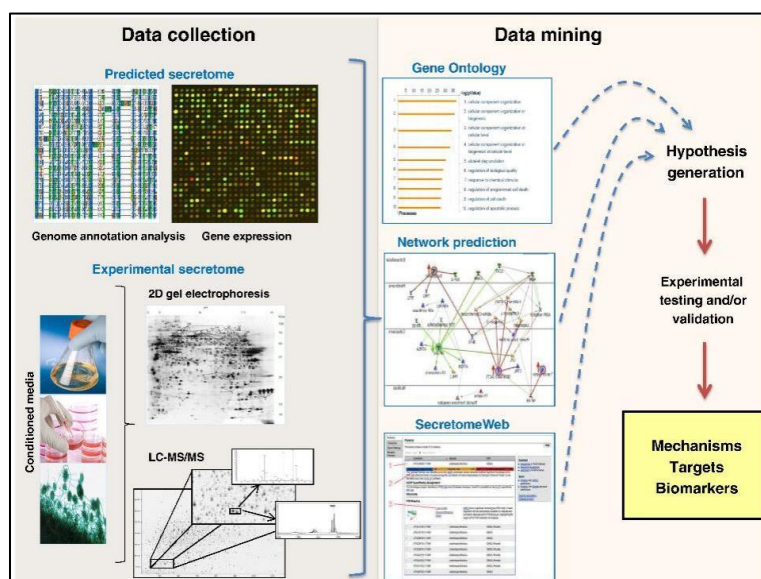


Applications of Bioinformatics

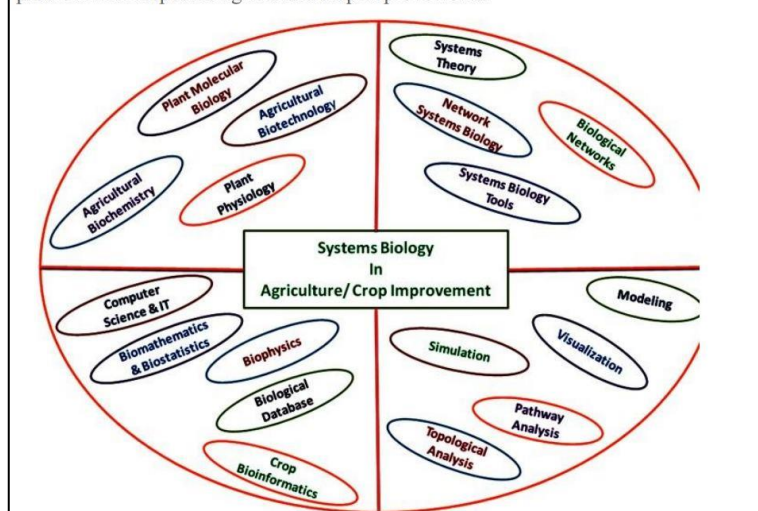
- Drug design
- Gene therapy
- Sequence analysis of proteins and DNA-Disease
- Molecular structure related to Disease
- Analysis and simulation of metabolic networks
- SNP & Disease
- Pathway Analysis (Pathomechanism)
- Diagnostic
- Prognostic
- Vaccine Design
- Molecular Mechanis







Systems Biology: An integration of interdisciplinary approaches for solving biological problem with respect to agriculture/crop improvements.





CASE STUDY

INDONESIA

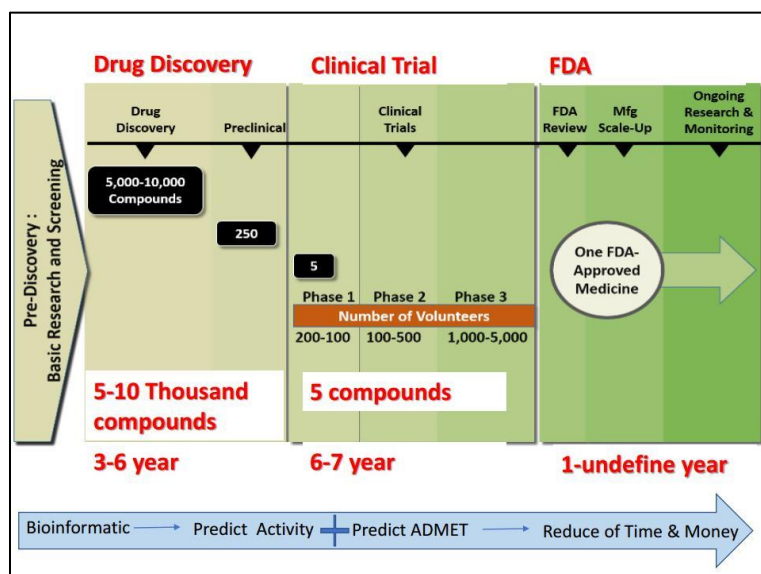


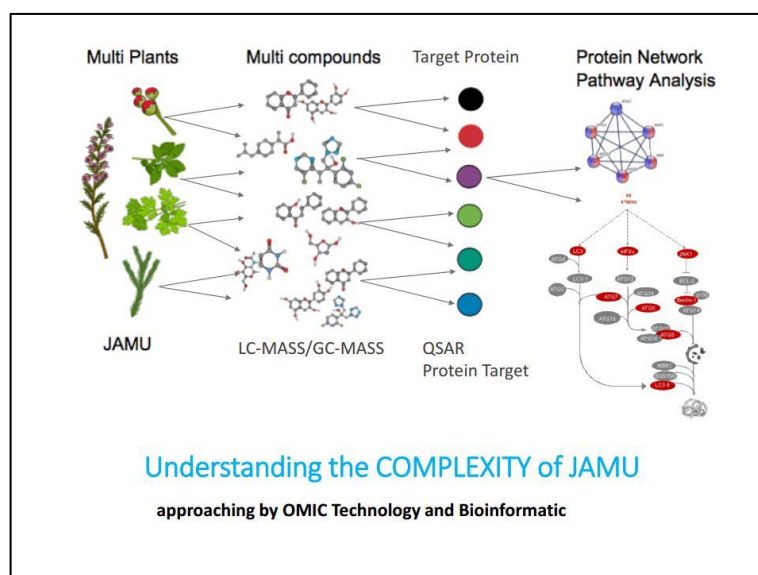
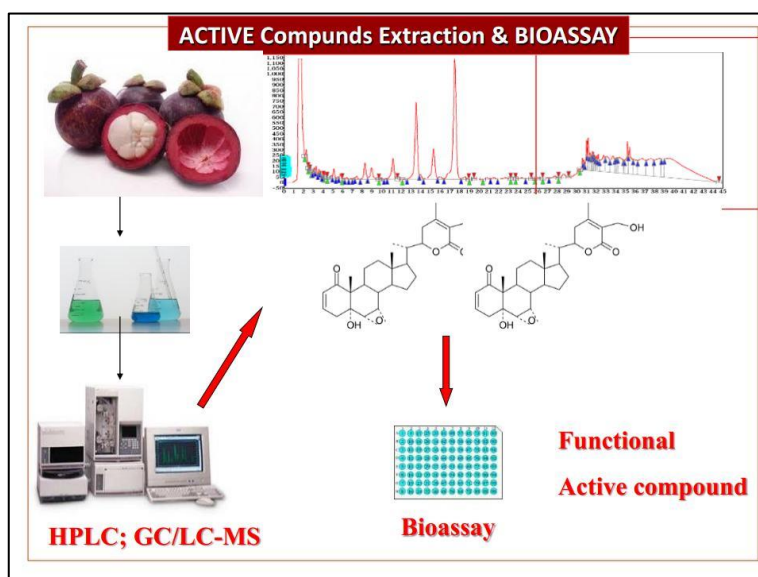
Mammals: 667
Birds: 1604
Reptiles: 749
Plants: 30,000

7.000 plants have Medicinal activity
(Erdelen et al.,1999)

**HUGE POTENCY :
RICHEST BIODIVERSITY**





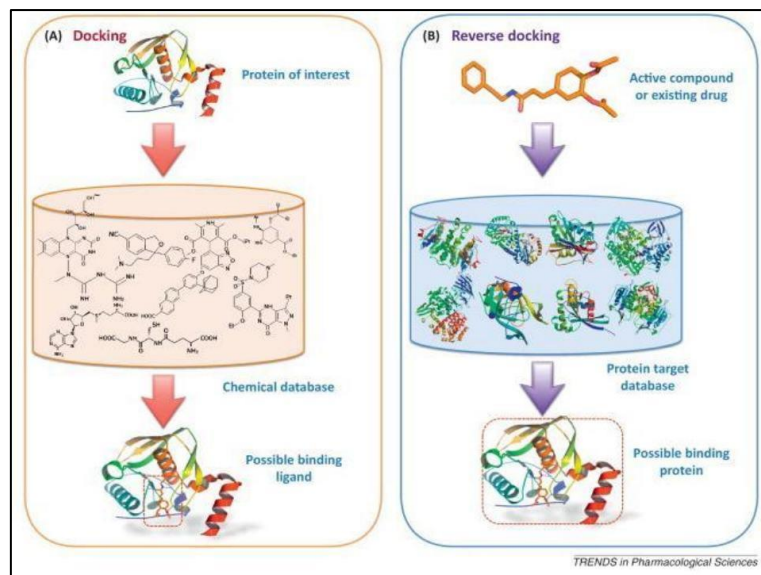
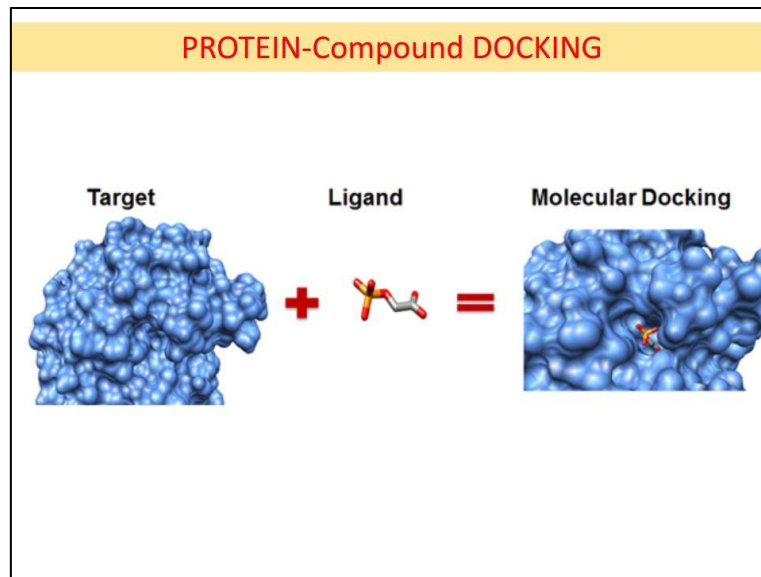


IN SILICO SCREENING

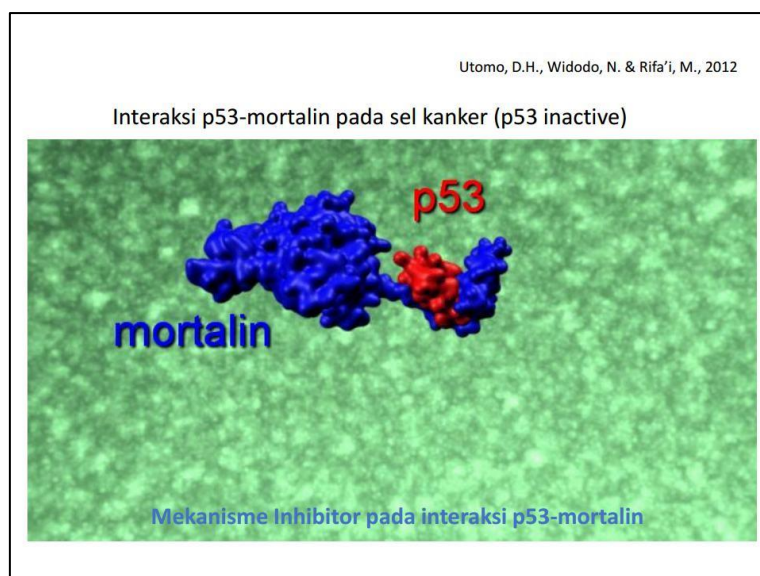
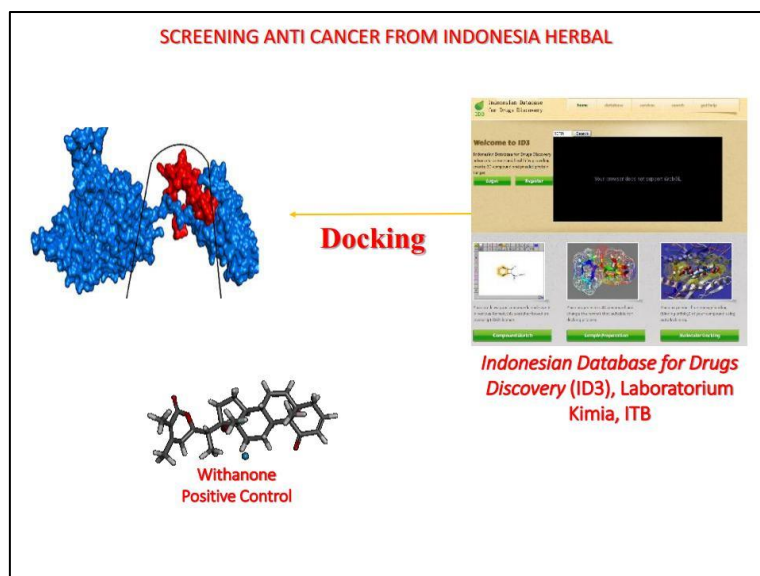
Modality

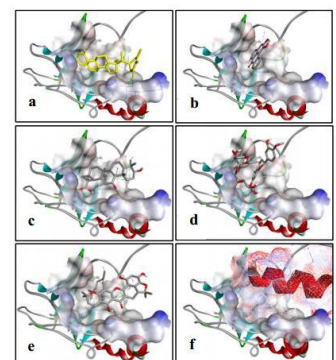
1. Database:
 - a. 8206 in Drug Bank
 - b. Over 35 million in ZINC
 - c. 82.6 million in PUBCHEM
2. Robust Method in GCMS
3. Predict Function
4. Predict Toxicity





CASE STUDY





Gambar 1. Pola interaksi antara senyawa aktif (dalam bentuk stik) dengan mortalin (kartun). Senyawa withanone (a), Liriodenine (b), glycyrrhethinic acid (c), Hesperidin (d), dan Artoindonesianin A (e) menempel di daerah pengikat p53 (*p53-binding domain*) pada protein mortalin (f), sehingga dimungkinkan dapat menghambat interaksi antara p53 dengan mortalin.

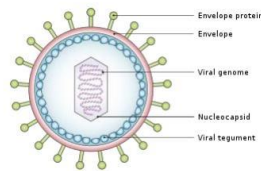
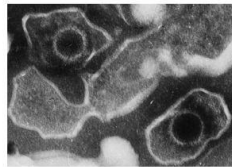
Tabel 2. Kemampuan pengikatan antara senyawa-senyawa aktif dengan protein mortalin. Empat senyawa aktif dari tanaman Indonesia memiliki binding affinity yang lebih baik di banding control positif, Withanone.

Senyawa Aktif	Binding Affinity (kcal/mol)
Hesperidin	-8.7
Artoindonesianin A	-8.6
Glycyrrhethinic Acid	-8.2
Liriodenine	-8.2
Withanone	-8.0
Gossypol	-7.8
Aloesin	-7.5
Eleagic Acid	-7.4
Rutin	-7.4
Diosbulbin B	-7.3
Betulinic Acid	-7.3
Geniposide	-7.0
Allamandin	-6.7
Resveratrol	-5.9
Anethole	-5.3
Eugenol	-5.3
Allicin	-4.0



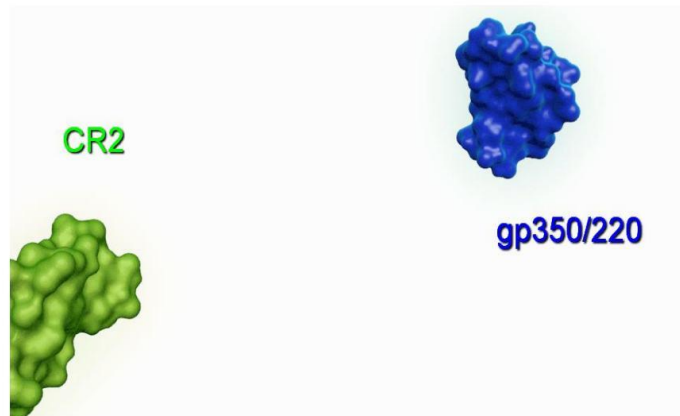
CASE STUDY

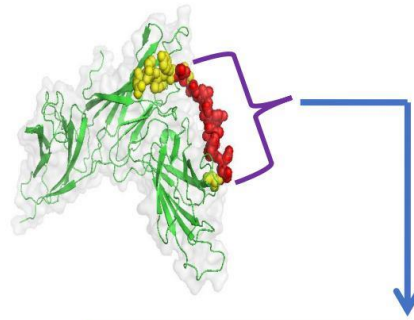
Introduction



- Epstein-Barr virus (EBV) usually infects children and commonly does not show direct symptoms after infection.
- This virus infected many people, especially in the Asia region, including Indonesia.
- EBV has a very vital role in nasopharyngeal carcinoma (NPC) progression.
- One of the efforts to prevent the NPC is vaccination.

Protein gp350/220 is capsid of EBV bind into CR2, during infection process





Epitope, potential for vaccine

We have identified epitope of Gp350/220 protein that potential for the vaccine of EBV (Sitompul et al., 2012).

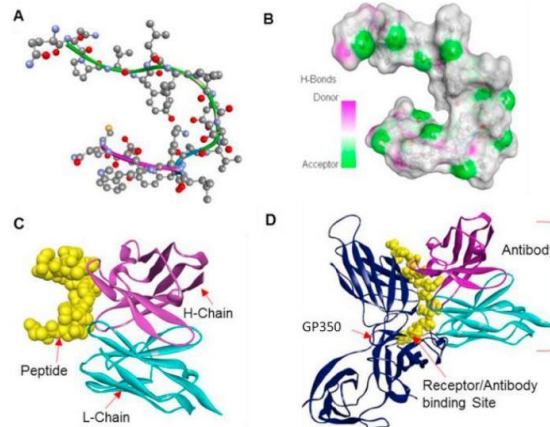


Fig. 1. The tertiary structure of EBVepitope (A) showed a coil structure with many hydrogen donors/acceptors (B). The epitope could bind 72A1 mAb in both the heavy- (purple) and light-chain (cyan) regions (C), which was similar to the binding pattern between the intact GP350/220 protein and 72A1 mAb (D). (For interpretation of the

TERIMAKASIH