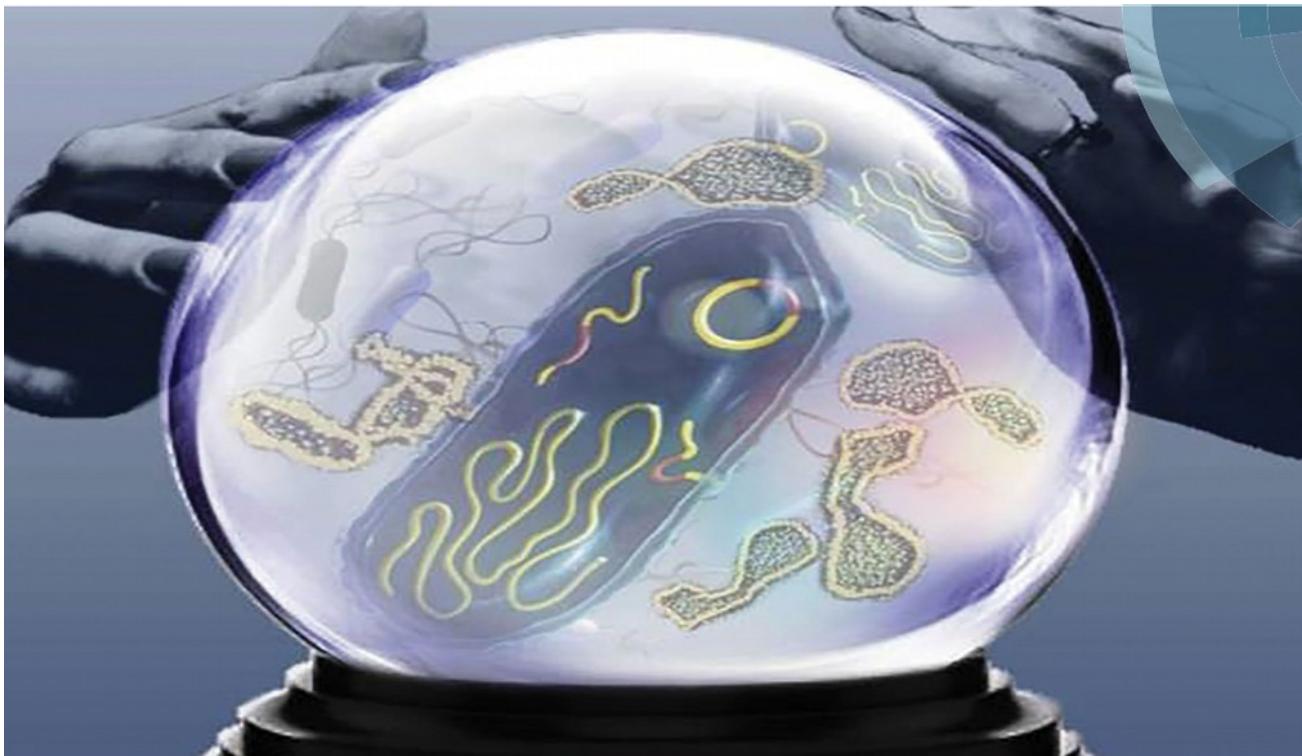


Introduction: Machine Learning MultiOmics Integration

Physalia course, online via zoom

Nikolay Oskolkov, MRG Group Leader, LIOS, Riga, Latvia



@NikolayOskolkov



@oskolkov.bsky.social



Personal homepage:
<https://nikolay-oskolkov.com>

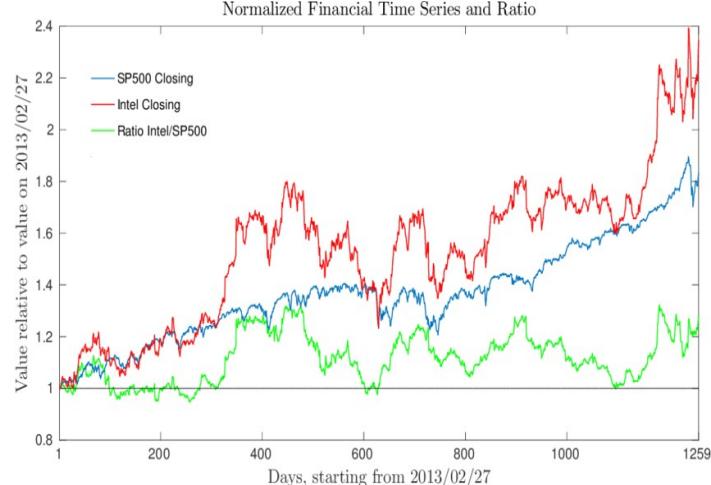
Image adapted from Molecular Omics, Issue 1, 2018

Introduction: High Dimensional Omics Data

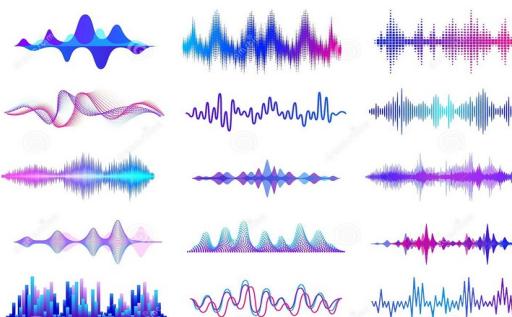
Tabular

A screenshot of Microsoft Excel showing a complex tabular dataset. The table has numerous columns with headers such as '#', 'JENKINS', 'Yr', 'Mon', 'Day', 'Time', 'Serial', 'Footprint', 'Core Speed', 'Core Scale', 'Mem', 'Mem Size', 'Mem Speed', 'Absorb', 'Videt', 'Videt Speed', 'Videt Scale', 'Temp', 'Temp Scale', 'Code', 'Pulse', 'Relative Humidity', 'Status', 'Cloud Cover', and 'Cloud Scale'. The data consists of numerical values and some text entries.

Time Series



Sound

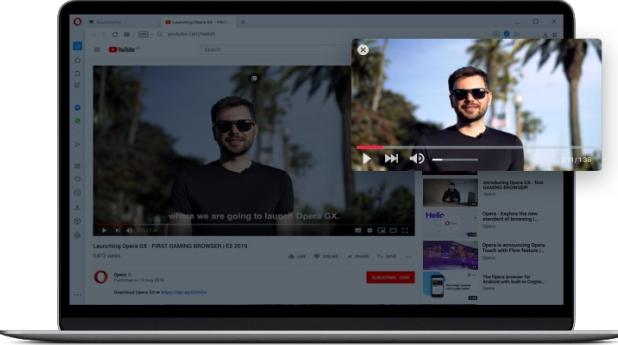


Image



DATA

Video



Text

Editing Wikipedia articles on
Medicine



Editing Wikipedia can be daunting for novices, especially if you're editing for the first time or for the first time as a class assignment. This guide is designed to assist students who have never edited Wikipedia before to understand some of the basic concepts and respond to it, and it would be polite for you to acknowledge the time they volunteer to help you learn how to edit.

Be accurate

Wikipedia is a resource millions

of people use to make medical

decisions, so it's vitally important

to be as accurate as possible

for medical information than the

second best source, like the

WHO. But with great power comes

great responsibility!

Understand the guidelines

While many of the guidelines

in this guide are in the media

area, we have developed additional

guidelines to ensure that the

content is medically sound.

Take extra time to read

and understand these guidelines.

When you edit, remember

your changes must fit these special

guidelines, or your edits are likely to be undone by other editors

as they clean up after you. That

takes time, so don't worry

about creating content. If you're not

comfortable with these guidelines,

talk to your instructor

about an alternative off-wiki

assignment.

Should I? Don't I?

Everyone wants to make the

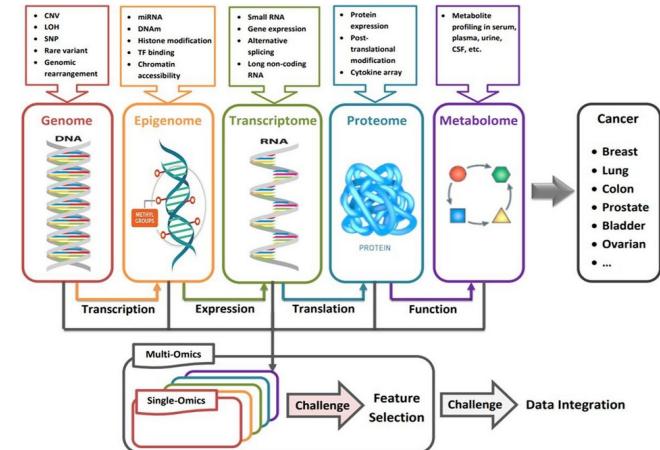
encyclopedia fun. Take the time

to understand the rules, and soon you'll be

contributing to a valuable resource you'll be

warning others about.

Omics Data

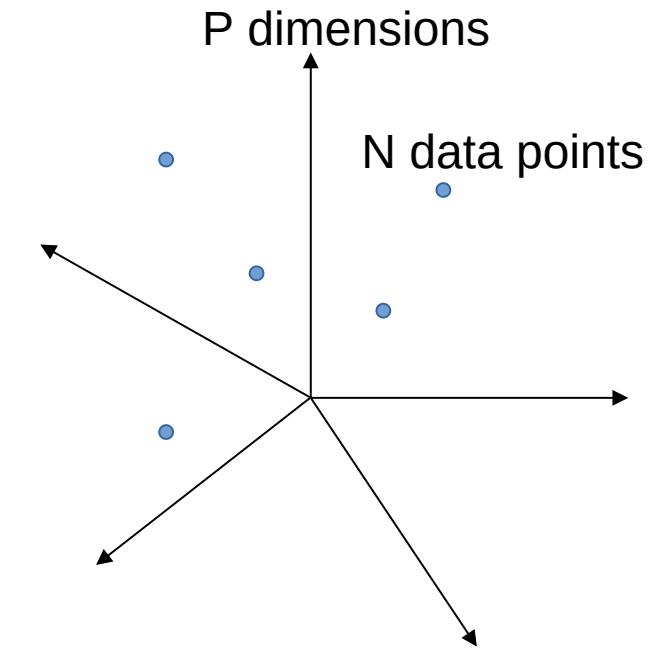


Statistical observations:
e.g. samples, cells etc.

Features: genes, proteins,
microbes, metabolites etc.

N

0	3	1	0	2	3	8	1	1	3
1	1	0	0	7	1	2	2	3	3
1	2	2	0	0	6	7	1	2	2
1	2	3	10	0	4	6	1	0	5
3	2	2	1	4	3	2	1	6	0
7	4	4	5	3	9	6	1	6	1
7	1	1	5	2	8	9	1	3	6
5	0	1	6	2	0	0	0	1	5
1	6	3	3	4	6	2	0	1	1
1	2	2	4	1	1	3	0	8	2



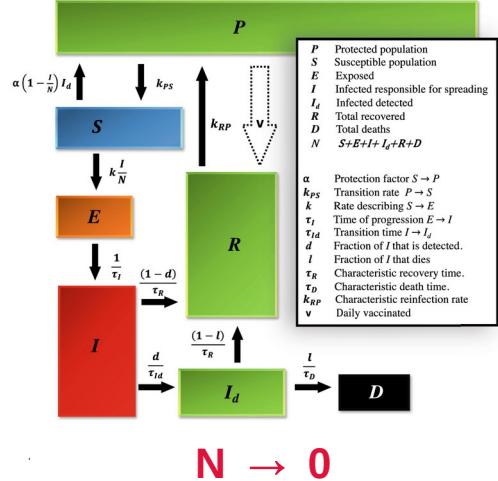
High Dimensional Data:
P >> N

For a robust statistical analysis, one should properly “sample” the P-dimensional space, hence large sample size is required, $N \gg P$

P is the number of features (genes, proteins, genetic variants etc.)
N is the number of observations (samples, cells, nucleotides etc.)

Biology / Biomedicine

Mathematical modeling



Bayesianism



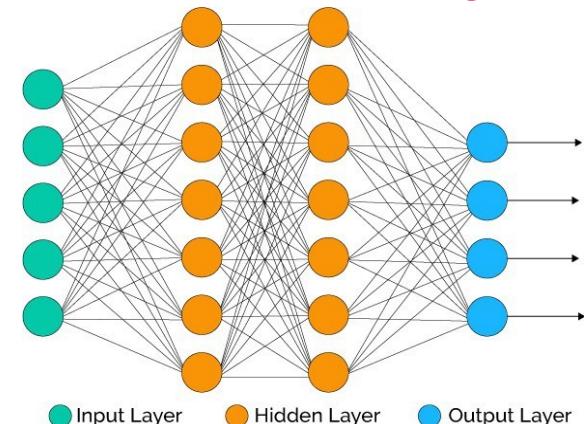
$N \ll P$

Frequentism



$N \approx P$

Machine Learning



Hypothesis-driven

The Curse of Dimensionality

Amount of Data

Data-driven

Ex.1

$$Y = \alpha + \beta X$$

$$\beta = (X^T X)^{-1} X^T Y$$

$$(X^T X)^{-1} \sim \frac{1}{\det(X^T X)} \dots \rightarrow \infty, \quad n \ll p$$

$$\text{Ex.2} \quad E[\hat{\sigma}^2] = \frac{n-p}{n} \sigma^2$$

Biased ML variance estimator in HD-space

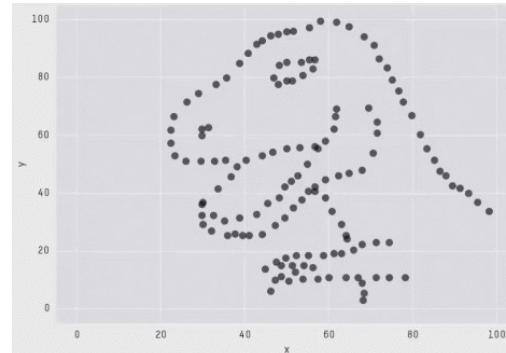
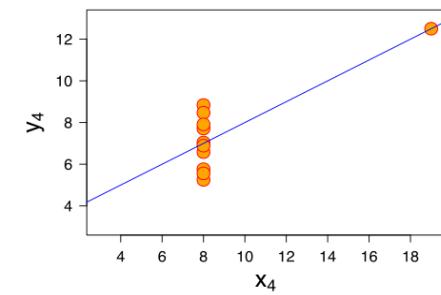
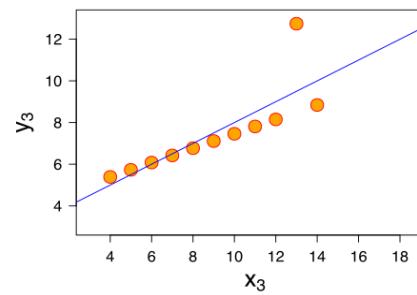
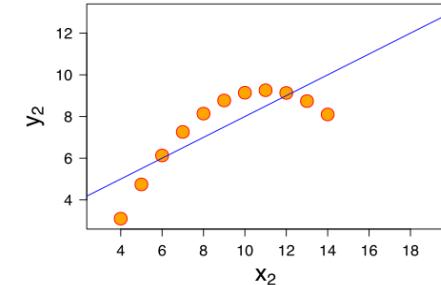
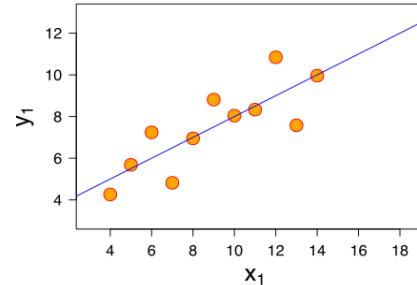
- Maximum likelihood based
- Focus on summary statistics
- Focus too much on p-values

$$L(x_i | \mu, \sigma^2) = \frac{1}{\sqrt{2\pi\sigma^2}} \exp^{-\frac{\sum_{i=1}^N (x_i - \mu)^2}{2\sigma^2}}$$

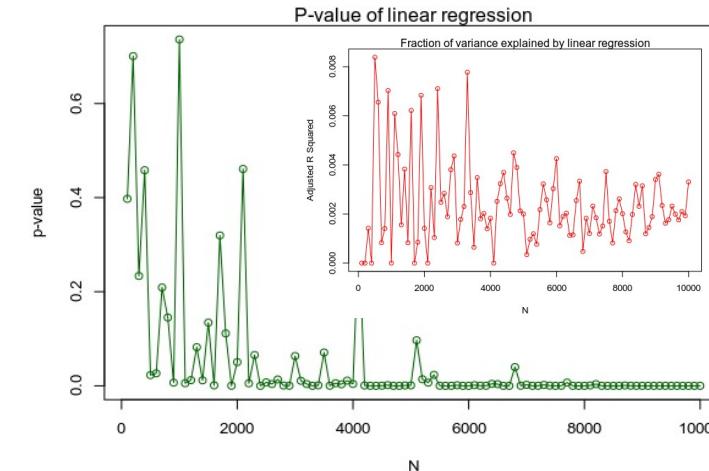
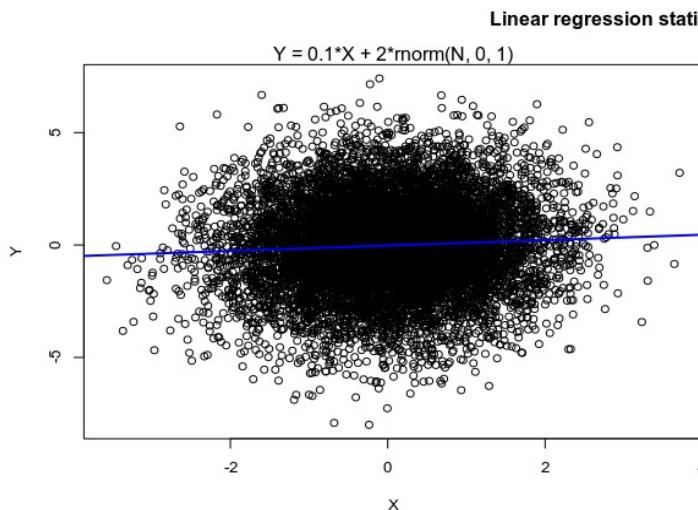
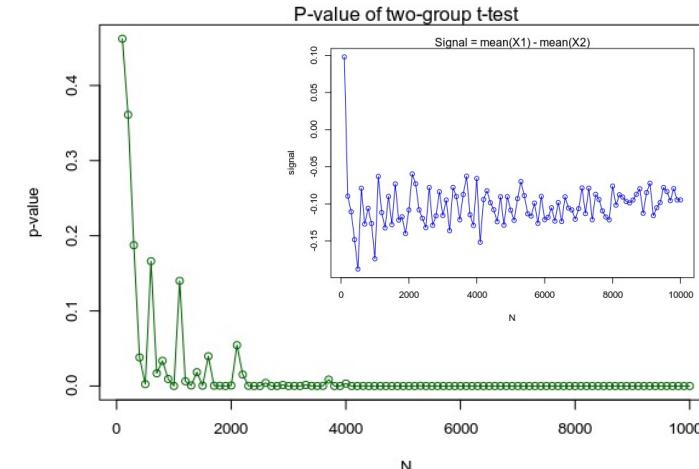
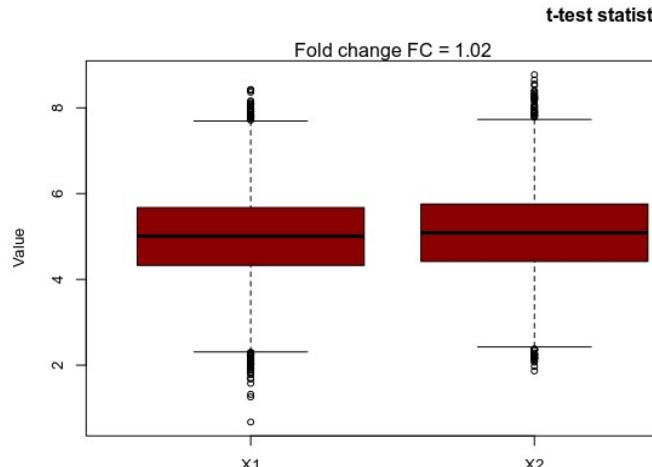
$$\frac{\partial L(x_i | \mu, \sigma^2)}{\partial \mu} = 0; \quad \frac{\partial L(x_i | \mu, \sigma^2)}{\partial \sigma^2} = 0$$

$$\mu = \frac{1}{N} \sum_{i=0}^N x_i - \text{mean estimator}$$

$$\sigma^2 = \frac{1}{N} \sum_{i=0}^N (x_i - \mu)^2 - \text{variance estimator}$$



X Mean:	54.2659224
Y Mean:	47.8313999
X SD :	16.7649829
Y SD :	26.9342120
Corr. :	-0.0642526

**nature**

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nature | 20 March 2019

Scientists rise up against statistical significance

Valentin Amrhein, Sander Greenland, Blake McShane and more than 800 signatories call for an end to hyped claims and the dismissal of possibly crucial effects.

Valentin Amrhein, Sander Greenland & Blake McShane



It is questionable whether
p-value is the best metric
for ranking features

```

1 n <- 20 # number of samples
2 p <- 2 # number of features / dimensions
3 Y <- rnorm(n)
4 X <- matrix(rnorm(n * p), n, p)
5 summary(lm(Y ~ X))

```

Call:
lm(formula = Y ~ X)

Residuals:

	Min	1Q	Median	3Q	Max
	-2.0522	-0.6380	0.1451	0.3911	1.8829

Coefficients:

	Estimate	Std. Error	t value	Pr(> t)
(Intercept)	0.14950	0.22949	0.651	0.523
X1	-0.09405	0.28245	-0.333	0.743
X2	-0.11919	0.24486	-0.487	0.633

Residual standard error: 1.017 on 17 degrees of freedom
 Multiple R-squared: 0.02204, Adjusted R-squared: -0.09301
 F-statistic: 0.1916 on 2 and 17 DF, p-value: 0.8274

Going to higher dimensions →

```

1 n <- 20 # number of samples
2 p <- 10 # number of features / dimensions
3 Y <- rnorm(n)
4 X <- matrix(rnorm(n * p), n, p)
5 summary(lm(Y ~ X))

```

Call:
lm(formula = Y ~ X)

Residuals:

	Min	1Q	Median	3Q	Max
	-1.0255	-0.4320	0.1056	0.4493	1.0617

Coefficients:

	Estimate	Std. Error	t value	Pr(> t)
(Intercept)	0.54916	0.26472	2.075	0.0679 .
X1	0.30013	0.21690	1.384	0.1998
X2	0.68053	0.27693	2.457	0.0363 *
X3	-0.10675	0.26010	-0.410	0.6911
X4	-0.21367	0.33690	-0.634	0.5417
X5	-0.19123	0.31881	-0.600	0.5634
X6	0.81074	0.25221	3.214	0.0106 *
X7	0.09634	0.24143	0.399	0.6992
X8	-0.29864	0.19004	-1.571	0.1505
X9	-0.78175	0.35408	-2.208	0.0546 .
X10	0.83736	0.36936	2.267	0.0496 *

Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

Residual standard error: 0.8692 on 9 degrees of freedom
 Multiple R-squared: 0.6592, Adjusted R-squared: 0.2805
 F-statistic: 1.741 on 10 and 9 DF, p-value: 0.2089

Going to even higher dimensions →

```

1 n <- 20 # number of samples
2 p <- 20 # number of features / dimensions
3 Y <- rnorm(n)
4 X <- matrix(rnorm(n * p), n, p)
5 summary(lm(Y ~ X))

```

Call:
lm(formula = Y ~ X)

Residuals:

ALL 20 residuals are 0: no residual degrees of freedom!

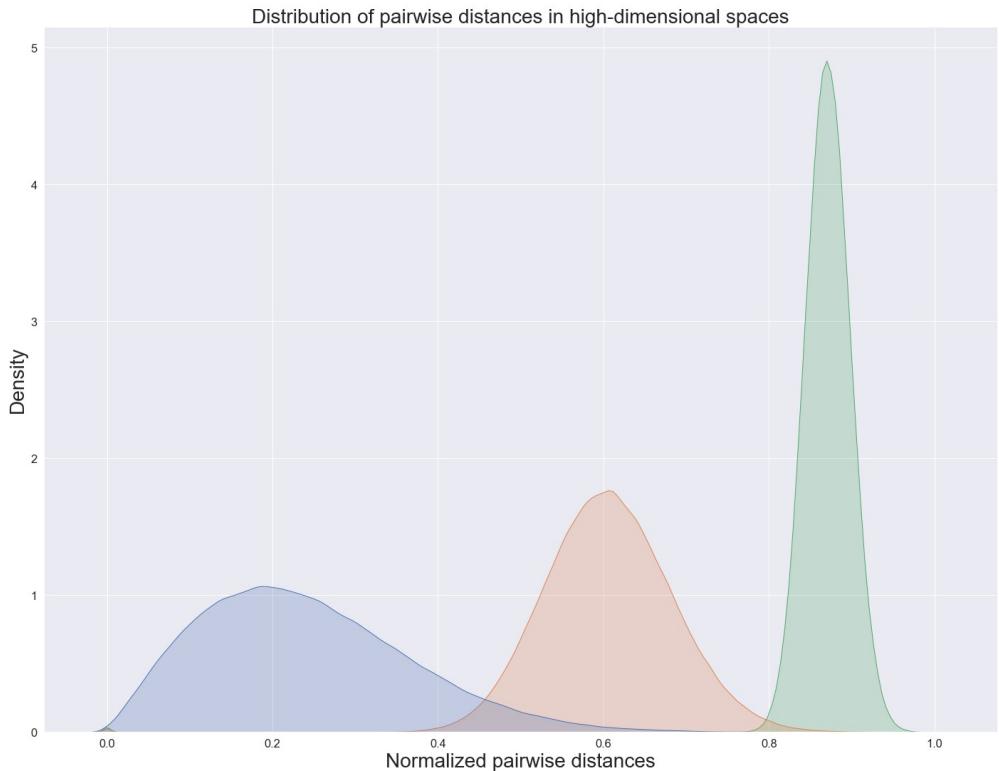
Coefficients: (1 not defined because of singularities)

	Estimate	Std. Error	t value	Pr(> t)
(Intercept)	1.34889	NaN	NaN	NaN
X1	0.66218	NaN	NaN	NaN
X2	0.76212	NaN	NaN	NaN
X3	-1.35033	NaN	NaN	NaN
X4	-0.57487	NaN	NaN	NaN
X5	0.02142	NaN	NaN	NaN
X6	0.40290	NaN	NaN	NaN
X7	0.03313	NaN	NaN	NaN
X8	-0.31983	NaN	NaN	NaN
X9	-0.92833	NaN	NaN	NaN
X10	0.18091	NaN	NaN	NaN
X11	-1.37618	NaN	NaN	NaN
X12	2.11438	NaN	NaN	NaN
X13	-1.75103	NaN	NaN	NaN
X14	-1.55673	NaN	NaN	NaN
X15	0.01112	NaN	NaN	NaN
X16	-0.50943	NaN	NaN	NaN
X17	-0.47576	NaN	NaN	NaN
X18	0.31793	NaN	NaN	NaN
X19	1.43615	NaN	NaN	NaN
X20	NA	NA	NA	NA

Residual standard error: NaN on 0 degrees of freedom
 Multiple R-squared: 1, Adjusted R-squared: NaN
 F-statistic: NaN on 19 and 0 DF, p-value: NA

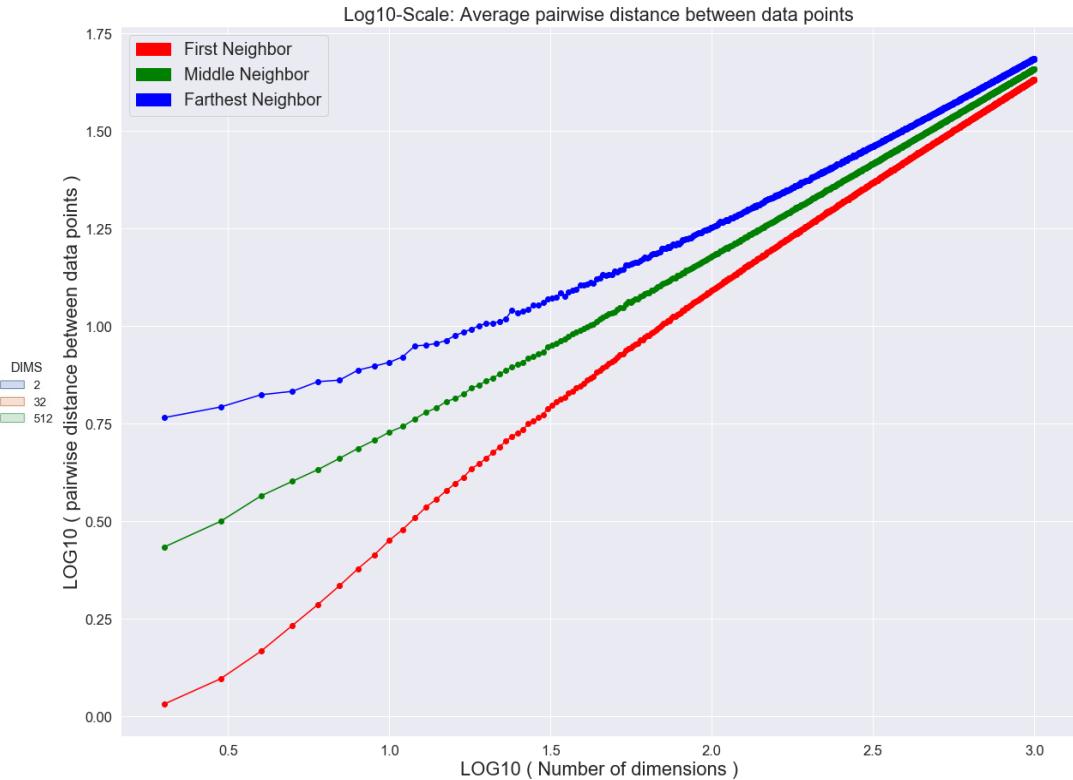
This is another way we face the Curse of Dimensionality in computational biology

More on the Curse of Dimensionality

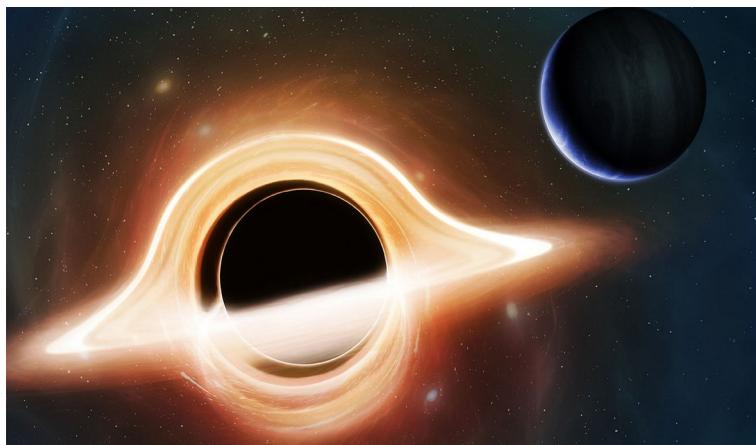
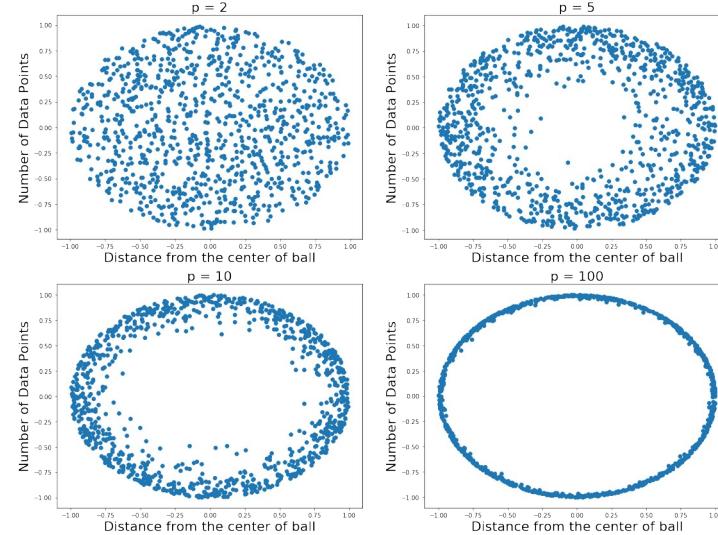
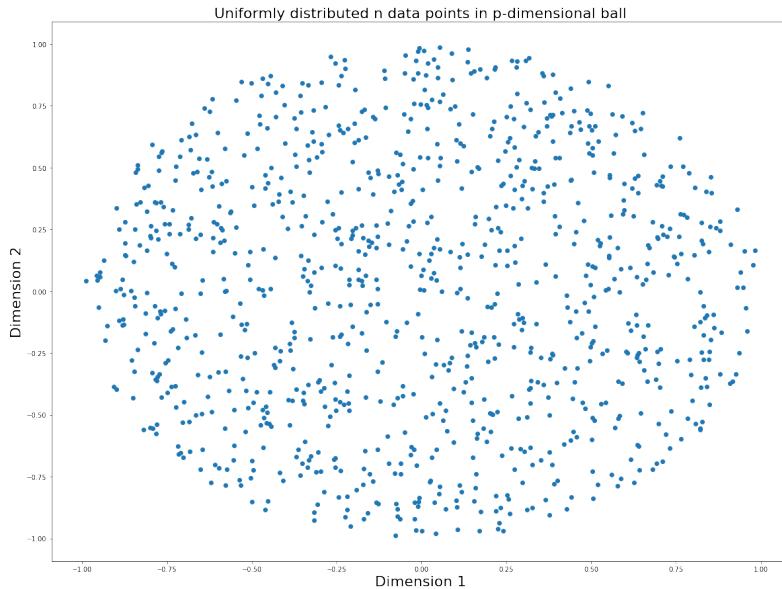


Data points become far from each other and equidistant in high dimensions

In high-dimensional space we can not separate cases and controls any more



The differences between closest and farthest data point neighbours disappears in high-dimensional spaces: can't run cluster analysis



High-dimensional data can be viewed as having a **"hole in the middle"**, hence the concept of mean / centroid loses its validity, hence we can't use Gaussian distribution

The curse(s) of dimensionality

There is such a thing as too much of a good thing.

Naomi Altman and Martin Krzywinski

We generally think that more information is better than less. However, in the 'big data' era, the sheer number of variables that can be collected from a single sample can be problematic. This embarrassment of riches is called the 'curse of dimensionality'¹ (CoD) and manifests itself in a variety of ways. This month, we discuss four important problems of dimensionality as it applies to data sparsity^{1,2}, multicollinearity³, multiple testing⁴ and overfitting⁵. These effects are amplified by poor data quality, which may increase with the number of variables.

Throughout, we use n to indicate the sample size from the population of interest and p to indicate the number of observed variables, some of which may have missing values for some samples. For example, we may have $n = 1,000$ subjects and $p = 200,000$ single-nucleotide polymorphisms (SNPs).

First, as the dimensionality p increases, the 'volume' that the samples may occupy grows rapidly. We can think of each of the n

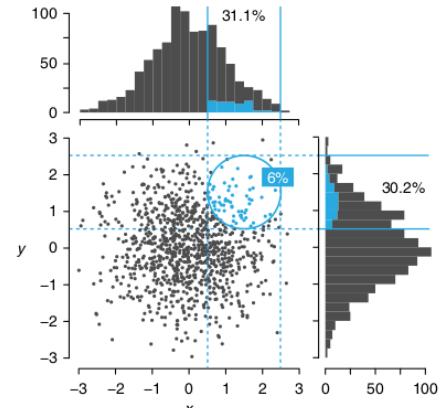


Fig. 1 | Data tend to be sparse in higher dimensions. Among 1,000 (x, y) points in which both x and y are normally distributed with a mean of 0 and s.d. $= 1$, only 6% fall within σ of $(x, y) = (1.5, 1.5)$ (blue circle). However, when the data are projected into a lower dimension—shown by histograms—about 30% of the points (all bins within blue solid lines) fall within ± 1.5 plus

A and 100 to have the minor allele a. If we tabulate on two SNPs, A and B, we will expect only ten samples to exhibit both minor alleles with genotype ab. With SNPs A, B and C, we expect only one sample to have genotype abc, and with four or more SNPs, we expect empty cells in our table. We need a much larger sample size to observe samples with all the possible genotypes. As p increases, we may quickly find that there are no samples with similar values of a predictor.

Even with just five SNPs, our ability to predict and classify the samples is impeded because of the small number of subjects that have similar genotypes. In situations where there are many gene variants, this effect is exacerbated, and it may be very difficult to find affected subjects with similar genotypes and hence to predict or classify on the basis of genetic similarity.

If we treat the distance between points (e.g., Euclidian distance) as a measure of similarity, then we interpret greater distance as greater dissimilarity. As n increases, this

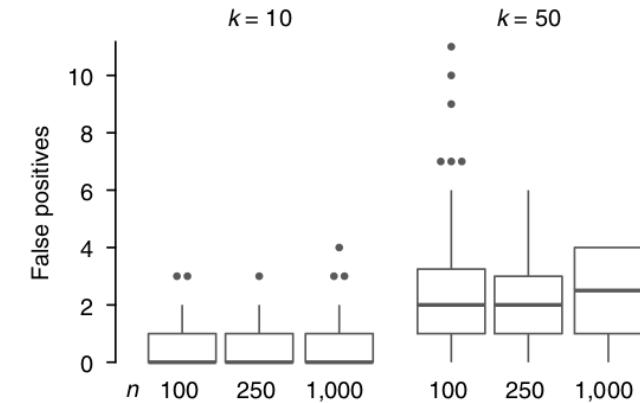


Fig. 3 | The number of false positives increases with each additional predictor. The box plots show the number of false positive regression-fit P values (tested at $\alpha = 0.05$) of 100 simulated multiple regression fits on various numbers of samples ($n = 100, 250$ and $1,000$) in the presence of one true predictor and $k = 10$ and 50 extraneous uncorrelated predictors. Box plots show means (black center lines), 25th and 75th percentiles (box edges), and minimum and maximum values (whiskers). Outliers (dots) are jittered.

Correcting for multiple testing does not solve the problem of too many false-positive hits

$$\begin{pmatrix} 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \\ 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \end{pmatrix}$$

Metabolomics
 $N \approx P$

$$\begin{pmatrix} 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \\ 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \end{pmatrix}$$

Proteomics
 $N \approx P$

$$\begin{pmatrix} 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \\ 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 \end{pmatrix}$$

Metagenomics
 $N \approx P$

manageable

P

$$N \begin{pmatrix} 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 \\ 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 1 & 0 & 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 \end{pmatrix}$$

Transcriptomics
 $N \ll P$
(Single cell: $N \leq P$)

challenging

P

$$N \begin{pmatrix} 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 \\ 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 \end{pmatrix}$$

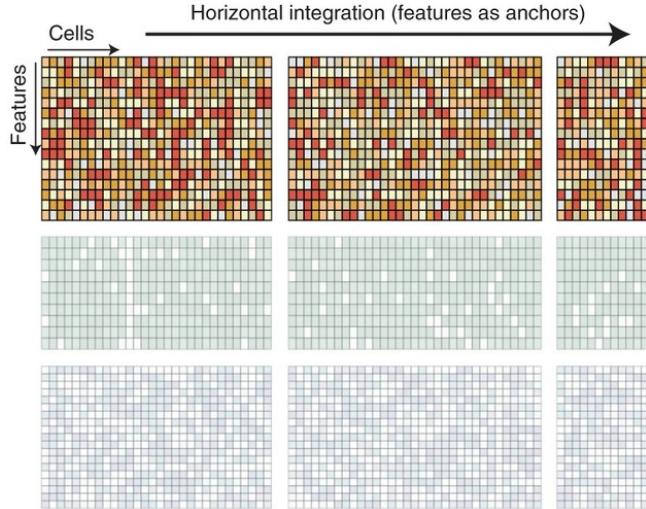
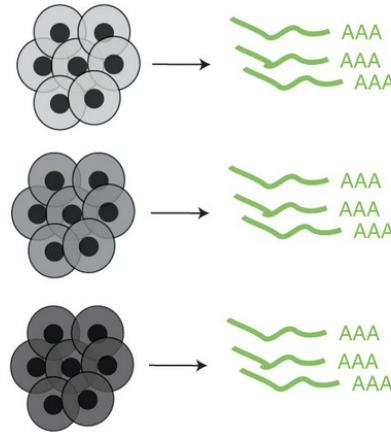
Genomics
 $N \llll P$

P

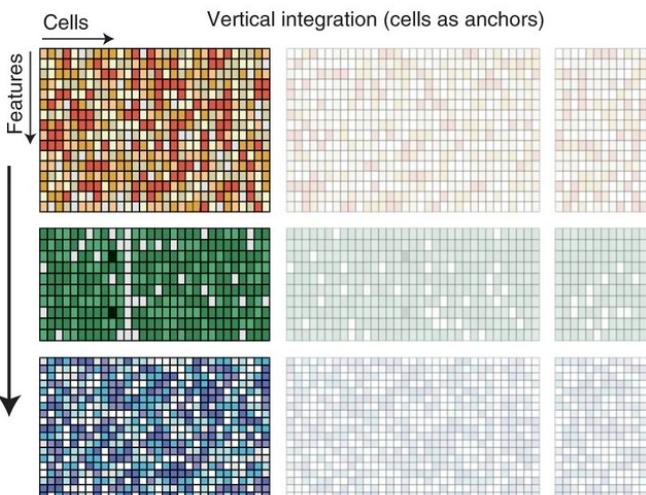
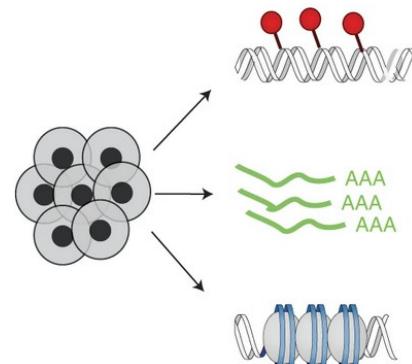
$$N \begin{pmatrix} 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 \\ 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 \\ 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 1 & 0 & 1 & 1 & 1 & 0 & 1 \end{pmatrix}$$

Methylomics
 $N \llll P$

Multi-Omics Data Integration

a

Similar to batch correction problem

b

Here I will focus on integration across features

Statistical observations:
e.g. samples, cells etc.

	N									
0	3	1	0	2	3	8	1	1	1	3
1	1	0	0	7	1	2	2	3	3	3
1	2	2	0	0	6	7	1	2	2	2
1	2	3	10	0	4	6	1	0	5	5
3	2	2	1	4	3	2	1	6	0	0
7	4	4	5	3	9	6	1	6	1	1
7	1	1	5	2	8	9	1	3	6	6
5	0	1	6	2	0	0	0	1	5	5
1	6	3	3	4	6	2	0	1	1	1
1	2	2	4	1	1	3	0	8	2	2

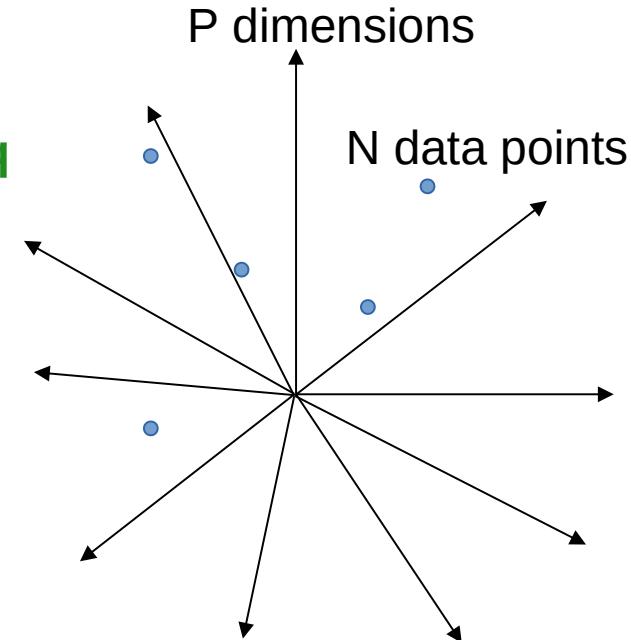
Features: genes, proteins,
microbes, metabolites etc.

	N									
0	3	1	0	2	3	8	1	1	1	3
1	1	0	0	7	1	2	2	3	3	3
1	2	2	0	0	6	7	1	2	2	2
1	2	3	10	0	4	6	1	0	5	5
3	2	2	1	4	3	2	1	6	0	0
7	4	4	5	3	9	6	1	6	1	1
7	1	1	5	2	8	9	1	3	6	6
5	0	1	6	2	0	0	0	1	5	5
1	6	3	3	4	6	2	0	1	1	1
1	2	2	4	1	1	3	0	8	2	2

P₂

Omic1:
RNAseq

Omic2:
BSseq

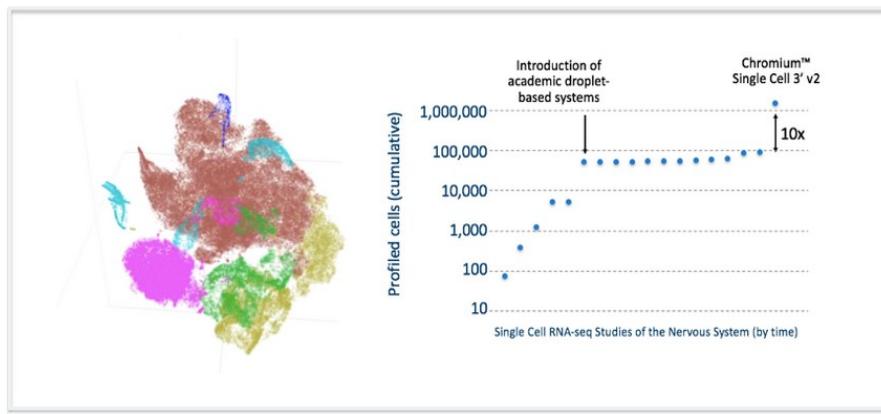


P₁ + P₂ >> N integration across features leads to even more high-dimensional data

Big Data in Single Cell Genomics

CAREERS BLOG 10X UNIVERSITY

10X GENOMICS SOLUTIONS & PRODUCTS RESEARCH & APPLICATIONS EDUCATION & RESOURCES

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Our 1.3 million single cell dataset is ready 0 KUDOS



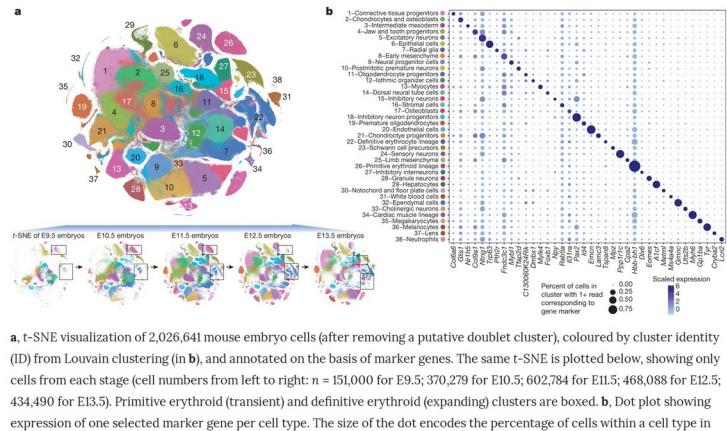
POSTED BY: grace-10x, on Feb 21, 2017 at 2:28 PM

At ASHG last year, we announced our 1.3 Million Brain Cell Dataset, which is, to date, the largest dataset published in the single cell RNA-sequencing (scRNA-seq) field. Using the Chromium™ Single Cell 3' Solution (v2 Chemistry), we were able to sequence and profile 1,308,421 individual cells from embryonic mice brains. Read more in our application note [Transcriptional Profiling of 1.3 Million Brain Cells with the Chromium™ Single Cell 3' Solution](#).

MENU nature

Fig. 2: Identifying the major cell types of mouse organogenesis.

From: The single-cell transcriptional landscape of mammalian organogenesis



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EXPLORER NOW

A next-generation platform to re-analyze published single-cell sequencing data

Single Cell Analysis

5,500,000 cells will be indexed into BioTuring Single-cell Data Repository this September

by @biomarkers • August 30, 2019

f | t | in

Human Cell Atlas, single-cell data

We are glad to announce that we will upscale the current single-cell database in BioTuring Single-cell Browser to 5,500,000 cells this September. With this release, we will double the current number of publications indexed in BioTuring Single-cell Browser, and cross the number of cells hosted on available public single-cell data repositories like Human Cell Atlas (HCA) and Broad Institute's Single-cell Portal.

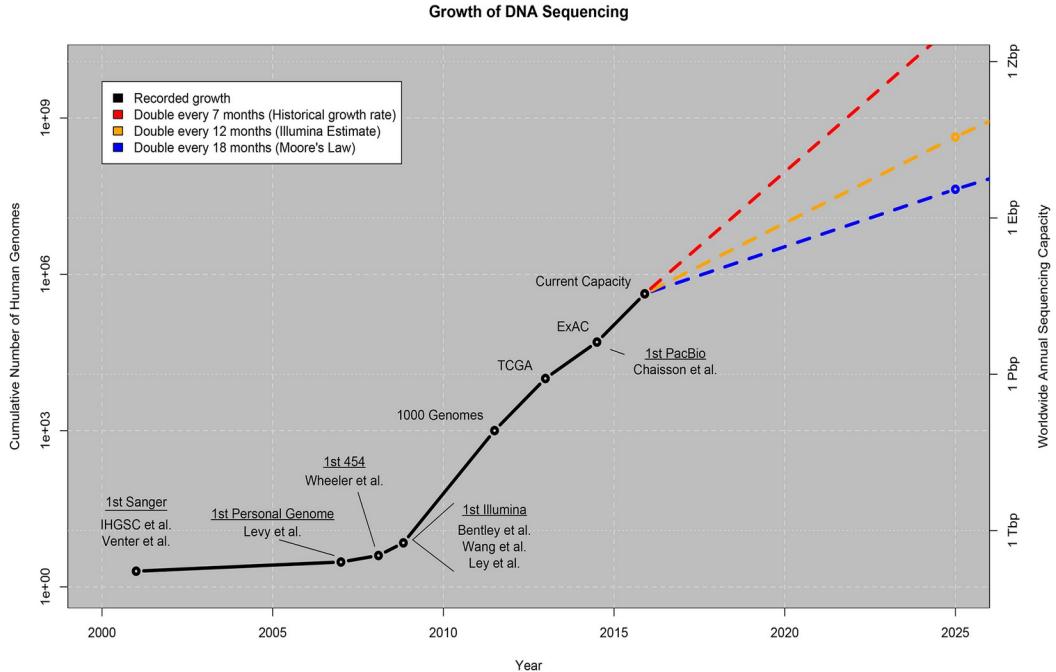
RECENT POSTS

A new tool to interactively visualize single-cell objects (Seurat, Scanpy, SingleCellExperiments,...)

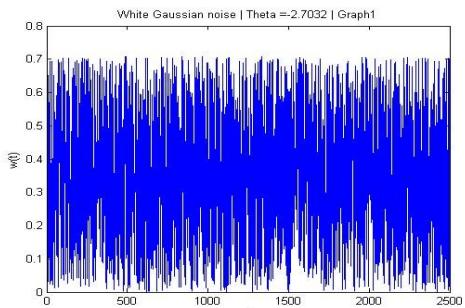
September 26, 2019

5,500,000 cells will be indexed into BioTuring Single-cell Data Repository this September

August 30, 2019



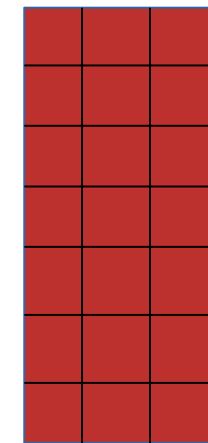
Stephens et al., (2015). Big Data: Astronomical or Genomical? PLoS Biology 13(7)



A file with **White Noise** can also take a lot of disk space

Genomics / WGS: Little Data

$$N_1 \sim 10^3$$

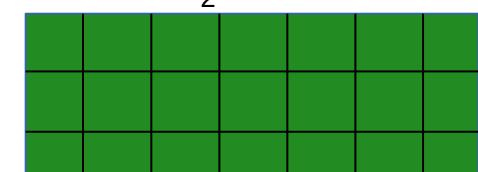


$$P_1 \sim 10^6$$

$$N_1 * P_1 = N_2 * P_2 = 10^9$$

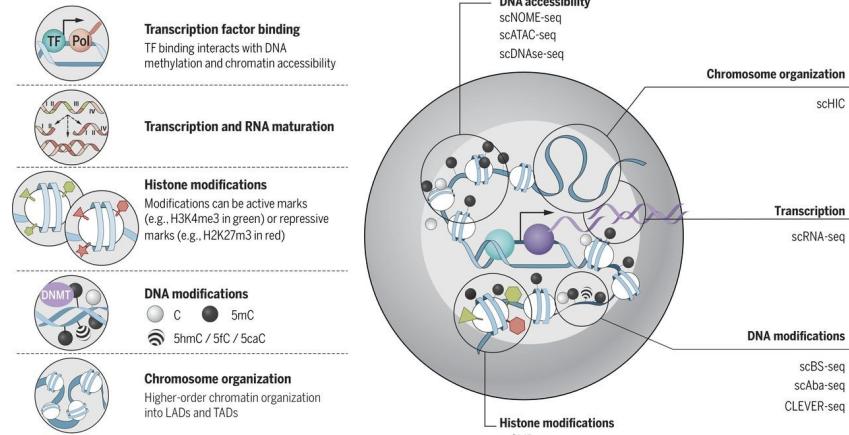
scRNASeq: Big Data

$$N_2 \sim 10^6$$

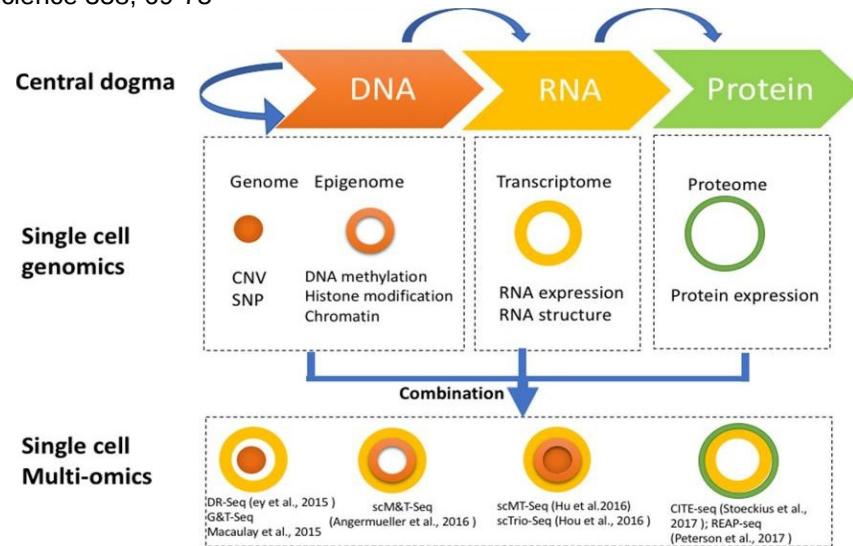


$$P_2 \sim 10^3$$

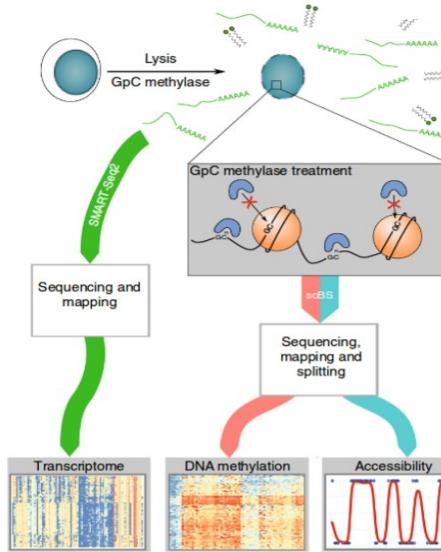
Multi-Omics in Single Cell Genomics



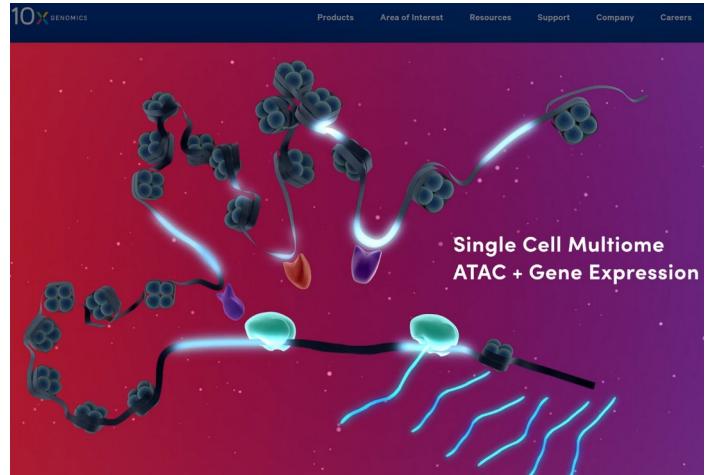
Kelsey et al., 2017, Science 358, 69-75



Hu et al., 2018, Frontier in Cell and Developmental Biology 6, 1-13



Clark et al., 2018, Nature Communications 9, 781

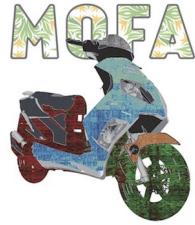
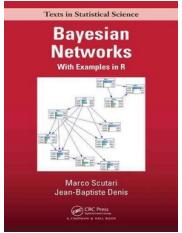


How to define and evaluate multi-Omics data integration?

OnPLS

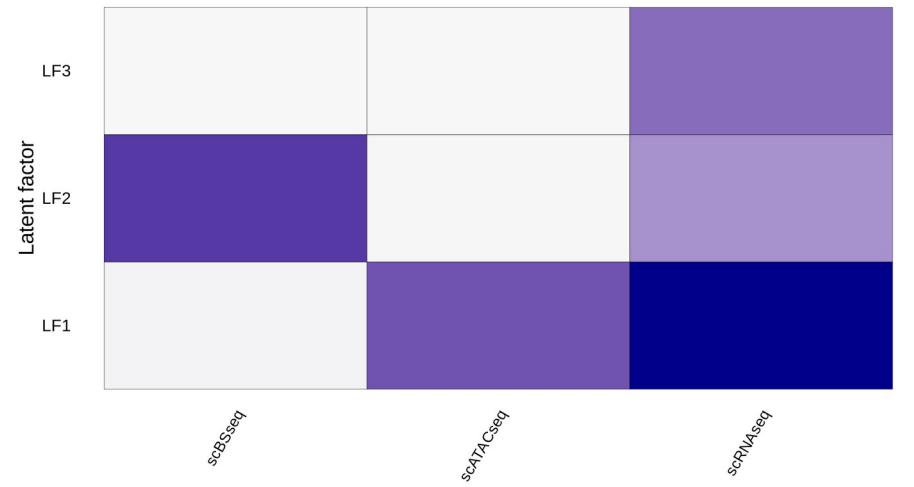
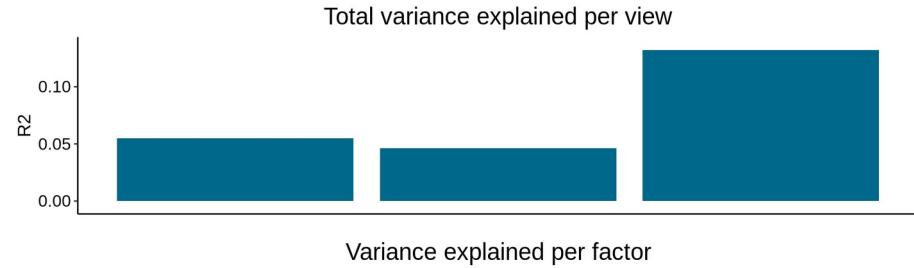
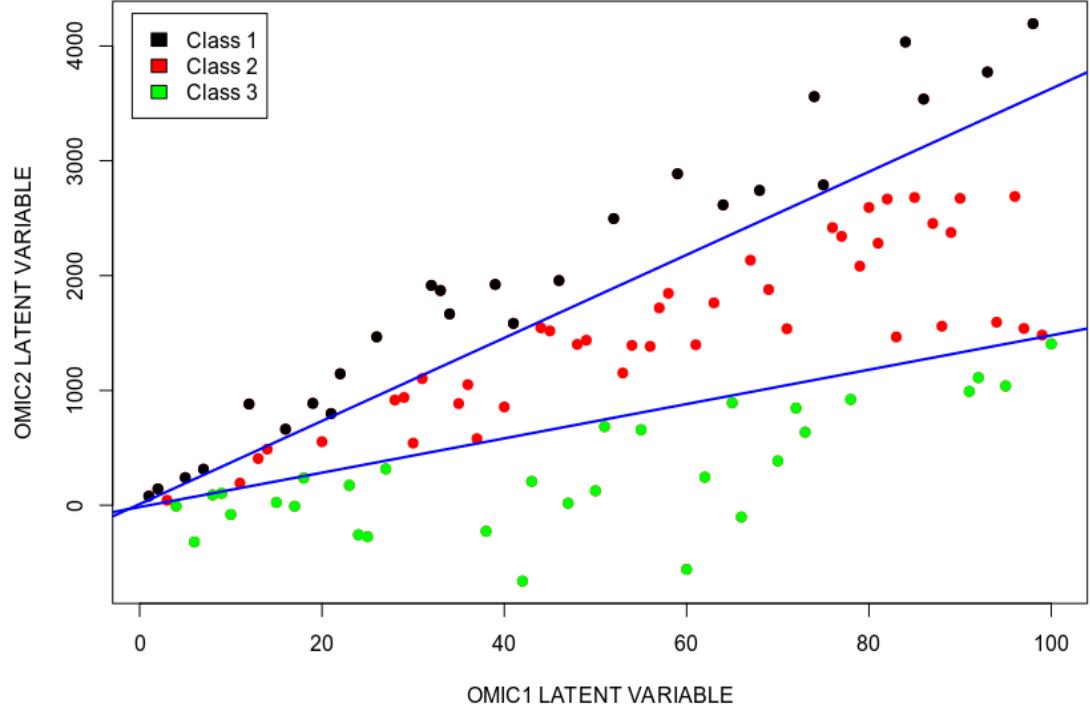
JIVE

DISCO



Clustering of Clusters

Idea Behind Omics Integration:
See Patterns Hidden in Individual Omics





How I Evaluate Omics Integration



TEXT (78%)



A	R	N	D	C	Q	E	G	H	I	L	K	M	F	P	S	T	W	Y	V
A	-4	-1	-2	0	-1	-1	0	-2	-1	-1	-1	0	-3	-2	0	-2	-2	0	-1
R	-1	5	-2	0	-1	0	0	-2	-1	-1	-1	0	-4	-2	-3	-2	-2	-1	-3
N	0	6	1	-3	0	0	0	-2	-1	-3	-3	0	-2	-1	0	-1	-2	-3	-2
D	-2	-2	1	6	-3	0	2	-1	-1	-3	-3	-1	-1	-4	-3	-3	-2	-3	-1
C	0	-3	-3	9	-3	-4	-3	-3	-1	-1	-3	-1	-2	-3	-1	-1	-2	-2	-1
Q	-1	1	0	0	-3	5	2	-2	0	-3	-2	1	0	-3	-1	0	-1	-2	-1
E	-1	1	0	0	-3	2	2	-2	0	-3	-2	1	0	-3	-1	0	-1	-2	-1
G	0	-2	0	-4	-3	2	5	-2	0	-3	-2	1	0	-3	-1	0	-1	-2	-1
F	-1	-2	0	1	-3	0	0	-2	-1	-2	-3	-1	-2	-3	-1	0	-1	-2	-1
H	-2	0	1	-1	-3	0	0	-2	8	-3	-3	-1	-2	-1	-2	-2	-2	-3	-1
I	-1	-3	-3	-3	-1	-3	-4	-3	-4	2	-3	1	0	-3	-2	-1	-3	-1	3
L	1	-1	-2	-3	-4	-1	-2	-3	-4	-3	-2	-1	-2	-1	-2	-1	-2	-1	-1
K	-1	-1	-2	-3	-4	-1	-2	-3	-4	-3	-2	-1	-2	-1	-2	-1	-2	-1	-1
M	-1	-1	-2	-3	-4	-1	-2	-3	-4	-3	-2	-1	-2	-1	-2	-1	-2	-1	-1
F	-2	-2	-3	-3	-2	-3	-3	-2	-3	-1	-2	-4	7	-1	-1	-4	-3	-2	-1
P	-1	-2	-2	-1	-3	-1	-1	-2	-3	-1	-2	-4	7	-1	-1	-4	-3	-2	-1
S	1	-1	1	0	-1	0	0	-1	-2	-2	0	-1	-2	-1	4	1	-3	-2	-2
T	0	-1	0	-1	-1	-1	-1	-2	-2	-1	-2	-1	-2	-1	4	1	-3	-2	-2
W	-3	-3	-4	-2	-3	-2	-3	-2	-3	-2	-3	-2	-3	-2	-3	-2	-3	-2	-3
Y	-2	-2	-2	-3	-2	-3	-2	-1	-2	-3	-2	-2	-1	-2	-3	-2	-2	-3	-1
V	0	-3	-3	-3	-1	-2	-2	-3	-3	-2	-1	-2	-2	0	-3	-1	-2	-3	-1

Predict Facebook user interests

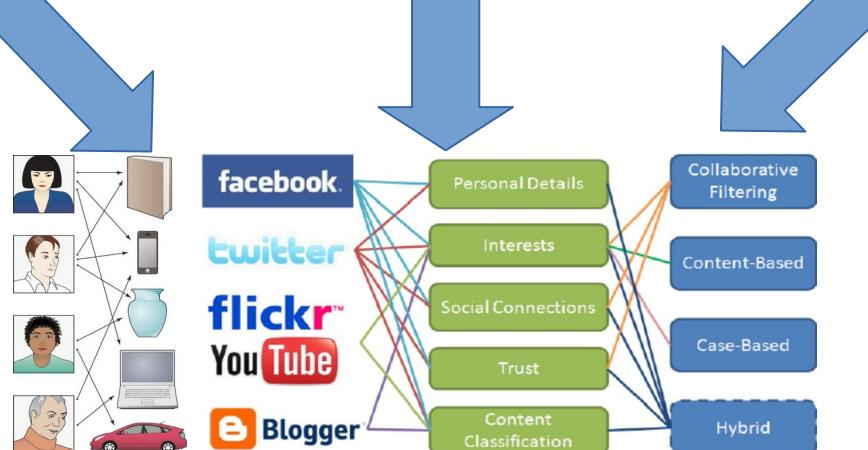
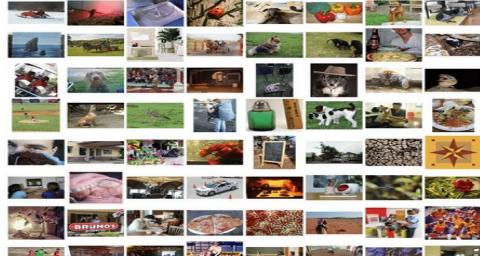
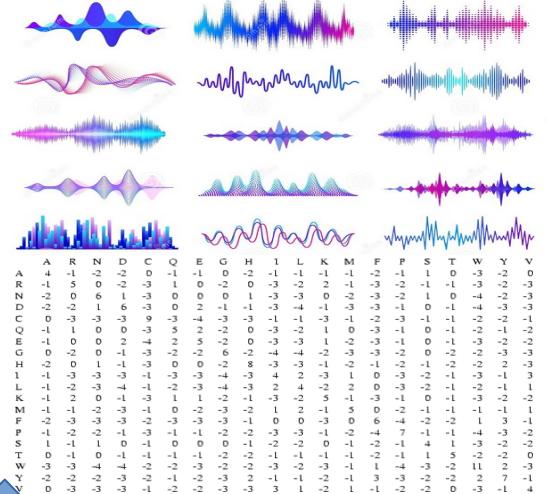


IMAGE (83%)



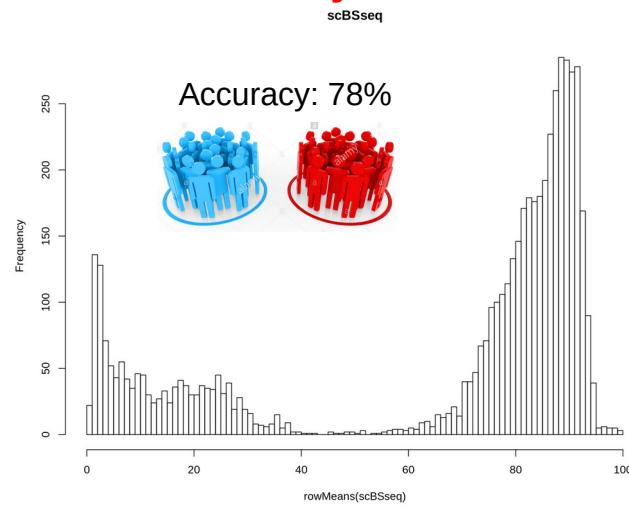
SOUND (75%)



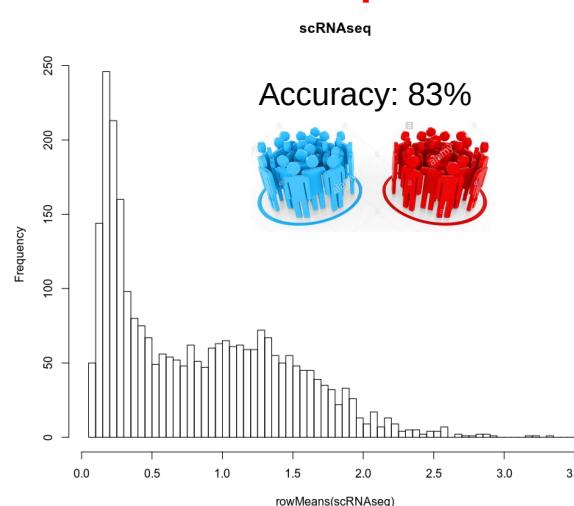
Data Integration Accuracy: 96%

How I Evaluate Omics Integration

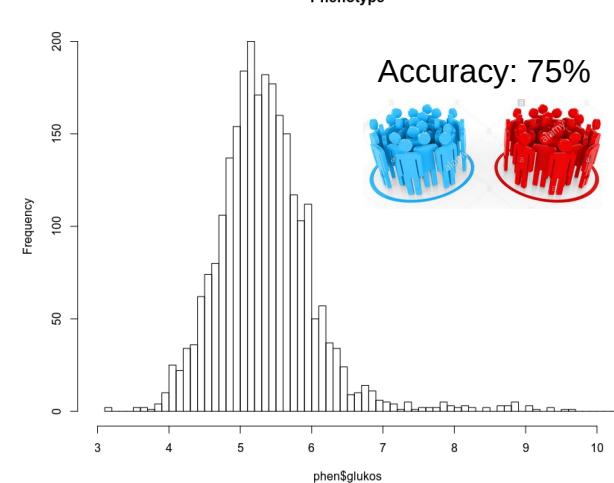
Methylation



Gene Expression

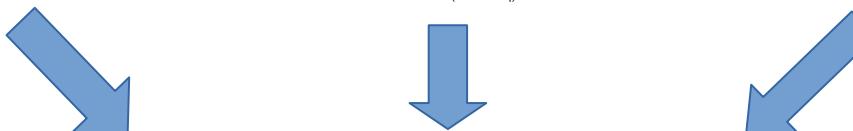


Phenotype



1) Convert to common space:

Neural Networks, SNF, UMAP



2) Explicitly model distributions:

MOFA, Bayesian Networks

3) Extract common variation:

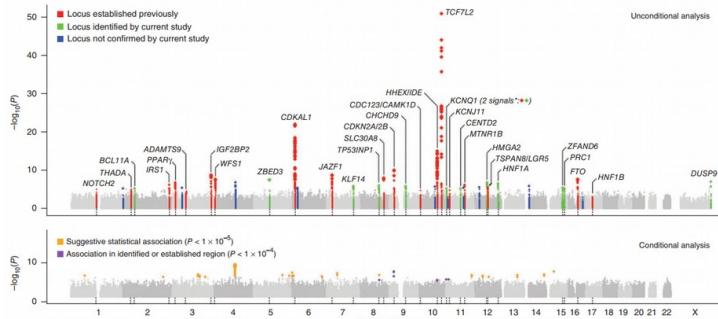
PLS, CCA, Factor Analysis



Data Integration Accuracy: 96%

Prediction as a Criterion of Success

Statistics searches for candidates



Consequence

NEWS FEATURE PERSONAL GENOMES NATURE/Vol 456/6 November 2008



The case of the missing heritability

B. Maher, Nature 456, 18-21 (2008)

Machine Learning optimizes prediction



Letter | Published: 31 July 2019

A clinically applicable approach to continuous prediction of future acute kidney injury

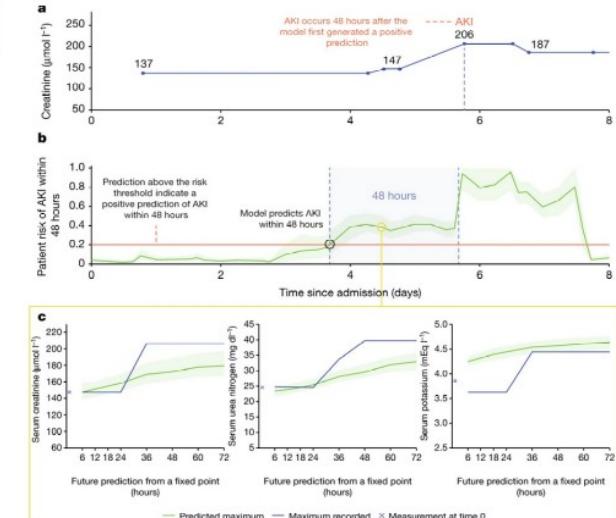
Nenad Tomalec, Kavir Glorot, [...] Shahril Mohamed

Nature 572, 116–119 (2019) | Download Citation |

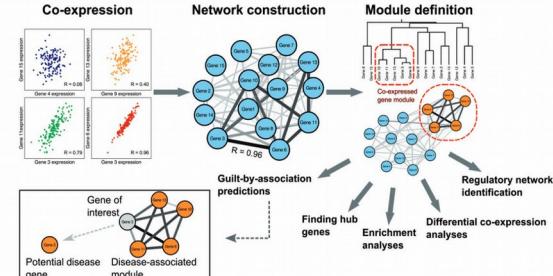
Abstract

The early prediction of deterioration could have an important role in supporting healthcare professionals, as an estimated 11% of deaths in hospital follow a failure to promptly recognize and treat deteriorating patients¹. To achieve this goal requires predictions of patient risk that are continuously updated and accurate, and delivered at an individual level with sufficient context and enough time to act. Here we develop a deep learning approach for the continuous risk prediction of future deterioration in patients, building on recent work that models adverse events from electronic health records^{2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17} and using acute kidney injury – a common and potentially life-threatening condition¹⁸ – as an exemplar. Our model was developed on a large, longitudinal dataset of electronic health records that cover diverse

From: A clinically applicable approach to continuous prediction of future acute kidney injury



Consequence



- 1) Biological data are **high-dimensional** and notoriously difficult to analyze
- 2) Integration across Omics is often sensitive to the **Curse of Dimensionality**
- 3) Integrating across Omics we expect to discover **novel patterns** in the data
- 4) Increased **prediction accuracy** is an indication of successful data integration
- 5) **Single cell Omics** are promising for integration in terms of statistical power



*Knut och Alice
Wallenbergs
Stiftelse*



**LUNDS
UNIVERSITET**