# Statistical Natural Language Processing A refresher on probability theory

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# Why probability theory?

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Short answer: practice proved otherwise.

### Slightly long answer

- Many linguistic phenomena are better explained as tendencies, rather than fixed rules
- Probability theory captures many characteristics of (human) cognition, language is not an exception

# What is probability?

- Probability is a measure of (un)certainty
- We quantify the probability of an event with a number between 0 and 1
  - 0 the event is impossible
  - 0.5 the event is as likely to happen as it is not
    - 1 the event is certain
- The set of all possible *outcomes* of a trial is called *sample space*  $(\Omega)$
- An *event* (E) is a set of outcomes

### Axioms of probability state that

- 1.  $P(E) \in \mathbb{R}$ ,  $P(E) \geqslant 0$
- 2.  $P(\Omega) = 1$
- 3. For disjoint events  $E_1$  and  $E_2$ ,  $P(E_1 \cup E_2) = P(E_1) + P(E_2)$

# What you should already know



- $P(\{\bullet\}) = 4/9$
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- $P(\{\bullet\}) = 1/9$
- $P(\{\bullet, \bullet\}) = 8/9$
- $P(\{\bullet, \bullet, \bullet\}) = 1$

# What you should already know



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$$P(\{\bullet\bullet\}) = 16/81$$

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• 
$$P(\{\bullet\bullet, \bullet\bullet\}) = 20/81$$

# Where do probabilities come from



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Two major (rival) ways of assigning probabilities to events are

- Frequentist (objective) probabilities: probability of an event is its relative frequency (in the limit)
- Bayesian (subjective) probabilities: probabilities are degrees of belief

- A random variable is a variable whose value is subject to uncertainties
- A random variable is always a number
- Think of a random variable as mapping between the outcomes of a trial to (a vector of) real numbers (a real valued function on the sample space)
- Example outcomes of uncertain experiments
  - height or weight of a person
  - length of a word randomly chosen from a corpus
  - whether an email is spam or not
  - the first word of a book, or first word uttered by a baby

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Note: not all of these are numbers

#### mapping outcomes to real numbers

- Continuous
  - frequency of a sound signal: 100.5, 220.3, 4321.3 ...
- Discrete
  - Number of words in a sentence: 2, 5, 10, ...
  - Whether a review is negative or positive:

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Outcome	Noun	Verb	Adj	Adv	
Value	1	2	3	4	

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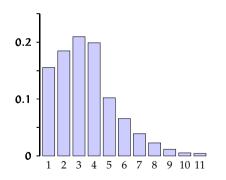
- The POS tag of a word:

Outcome	Noun	Verb	Adj	Adv	
Value <b>or</b>	10000	01000	00100	00010	

### Probability mass function

#### Example: probabilities for sentence length in words

• *Probability mass function (PMF)* of a *discrete* random variable (X) maps every possible (x) value to its probability (P(X = x)).



χ	P(X = x)
1	0.155
2	0.185
3	0.210
4	0.194
5	0.102
6	0.066
7	0.039
8	0.023
9	0.012
10	0.005
11	0.004

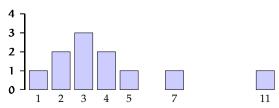
# Populations, distributions, samples

- A probability distribution characterizes a random variable
- We can define a distribution with a vector or table of probabilities, if we have a finite sample space
- Otherwise, we use (parametric) functions to map the (infinite) set of outcomes to probabilities
- Probability distributions characterize possibly infinite *populations*
- In most cases we have to work with samples

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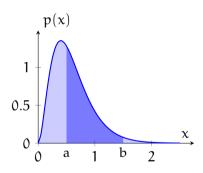
A sample from the distribution on the previous slide:



# Probability density function (PDF)

- Continuous variables have *probability density functions*
- p(x) is not a probability (note the notation: we use lowercase p for PDF)
- Area under p(x) sums to 1
- P(X = x) = 0
- Non zero probabilities are possible for ranges:

$$P(a \leqslant x \leqslant b) = \int_{a}^{b} p(x) dx$$



### Cumulative distribution function

• 
$$F_X(x) = P(X \le x)$$

1.0

1.2 3 4 5 6 7 8 9 10 11

Sentence length

Length	Prob.	C. Prob.
1	0.16	0.16
2	0.18	0.34
3	0.21	0.55
4	0.19	0.74
5	0.10	0.85
6	0.07	0.91
7	0.04	0.95
8	0.02	0.97
9	0.01	0.99
10	0.01	0.99
11	0.00	1.00
	•	

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10 / 57

# Expected value

• Expected value (mean) of a random variable X is,

$$E[X] = \mu = \sum_{i=1}^{n} P(x_i)x_i = P(x_1)x_1 + P(x_2)x_2 + \ldots + P(x_n)x_n$$

More generally, expected value of a function of X is

$$E[f(X)] = \sum_{x} P(x)f(x)$$

- Expected value is a measure of central tendency
- Note: it is not the 'most likely' value
- Expected value is linear

$$E[aX + bY] = aE[X] + bE[Y]$$

### Variance and standard deviation

• Variance of a random variable X is,

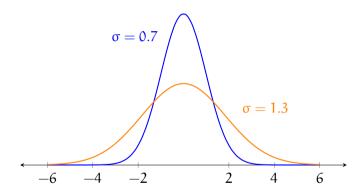
$$Var(X) = \sigma^2 = \sum_{i=1}^{n} P(x_i)(x_i - \mu)^2 = E[X^2] - (E[X])^2$$

- It is a measure of spread, divergence from the central tendency
- The square root of variance is called standard deviation

$$\sigma = \sqrt{\left(\sum_{i=1}^{n} P(x_i) x_i^2\right) - \mu^2}$$

- Standard deviation is in the same units as the values of the random variable
- Variance is not linear:  $\sigma_{X+Y}^2 \neq \sigma_X^2 + \sigma_Y^2$  (neither the  $\sigma$ )

# Example: two distributions with different variances



# Short divergence: Chebyshev's inequality

For any probability distribution, and k > 1,

$$P(|x - \mu| > k\sigma) \leqslant \frac{1}{k^2}$$

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Distance from μ	2σ	3σ	5σ	10σ	100σ
Probability	0.25	0.11	0.04	0.01	0.0001

14 / 57

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Distance from μ	2σ	3σ	5σ	10σ	100σ
Probability	0.25	0.11	0.04	0.01	0.0001

This also shows why standardizing values of random variables,

$$z = \frac{x - \mu}{\sigma}$$

makes sense (the normalized quantity is often called the z-score).

### Median and mode of a random variable

Median is the mid-point of a distribution. Median of a random variable is defined as the number m that satisfies

$$P(X \le m) \geqslant \frac{1}{2}$$
 and  $P(X \geqslant m) \geqslant \frac{1}{2}$ 

- Median of 1, 4, 5, 8, 10 is 5
- Median of 1, 4, 5, 7, 8, 10 is 6

### Median and mode of a random variable

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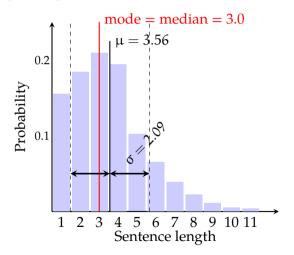
- Median of 1, 4, 5, 8, 10 is 5
- Median of 1, 4, 5, 7, 8, 10 is 6

Mode is the value that occurs most often in the data.

- Modes appear as peaks in probability mass (or density) functions
- Mode of 1, 4, 4, 8, 10 is 4
- Modes of 1, 4, 4, 8, 9, 9 are 4 and 9

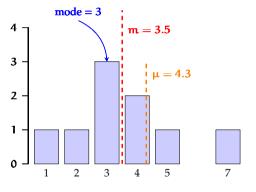
### Mode, median, mean, standard deviation

Visualization on sentence length example



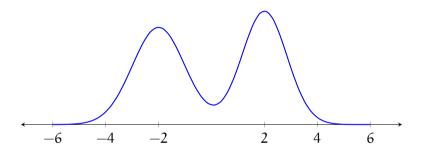
### Mode, median, mean

#### sensitivity to extreme values





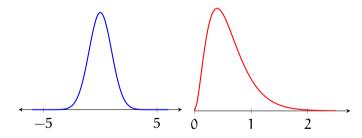
### Multimodal distributions



- A distribution is multimodal if it has multiple modes
- Multimodal distributions often indicate confounding variables

### Skew

- Another important property of a probability distribution is its *skew*
- symmetric distributions have no skew
- positively skewed distributions have a long tail on the right
- negatively skewed distributions have a long left tail

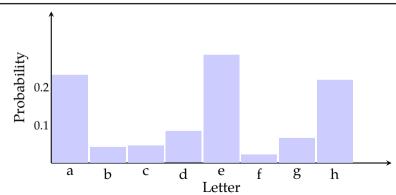


# Another example distribution

#### A probability distribution over letters

• An alphabet with 8 letters and their probabilities of occurrance;

				1			-	
Lett.	a	b	С	d	e	f	g	h
Prob.	0.23	0.04	0.05	0.08	0.29	0.02	0.07	0.22



### Probability distributions

- A distributions on a finite set of outcomes can be defined by a vector (or table) of probabilities
- Some random variables (approximately) follow a distribution that can be parametrized with a number of parameters
- For example, Gaussian (or normal) distribution is conventionally parametrized by its mean  $(\mu)$  and variance  $(\sigma^2)$
- Common notation we use for indicating that a variable X follows a particular distribution is

$$X \sim Normal(\mu, \sigma^2)$$
 or  $X \sim \mathcal{N}(\mu, \sigma^2)$ .

• For the rest of this lecture, we will revise some of the important probability distributions

# Probability distributions (cont)

- A probability distribution is called *univariate* if it was defined on scalars
- multivariate probability distributions are defined on vectors
- Probability distributions are abstract mathematical objects (functions that map events/outcomes to probabilities)
- A probability distribution is generative device: it can generate samples
- In most problems, we only have access to a samples
- Learning (or *inference*) is often cast as finding an (approximate) distribution from a sample

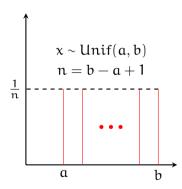
## Uniform distribution (discrete)

- A uniform distribution assigns equal probabilities to all values in range [a, b], where a and b are the parameters of the distribution
- Probabilities of the values outside range is 0

• 
$$\mu = \frac{b+a}{2}$$

• 
$$\sigma_2 = \frac{(b-a+1)^2-1}{12}$$

There is also an analogous continuous uniform distribution



#### Bernoulli distribution

Bernoulli distribution characterizes simple random experiments with two outcomes

- Coin flip: heads or tails
- Spam detection: spam or not
- Predicting gender: female or male

We denote (arbitrarily) one of the possible values with 1 (often called a success), the other with 0 (often called a failure)

$$P(X = 1) = p$$

$$P(X = 0) = 1 - p$$

$$P(X = k) = p^{k}(1 - p)^{1 - k}$$

$$\mu_{X} = p$$

$$\sigma_{X}^{2} = p(1 - p)$$

#### Binomial distribution

Binomial distribution is a generalization of Bernoulli distribution to n trials, the value of the random variable is the number of 'successes' in the experiment

$$P(X = k) = \binom{n}{k} p^{k} (1 - p)^{n-k}$$
$$\mu_{X} = np$$
$$\sigma_{X}^{2} = np(1 - p)$$

Remember that  $\binom{n}{k} = \frac{n!}{k!(n-k)!}$ .

## Categorical distribution

- Extension of Bernoulli to k mutually exclusive outcomes
- For any k-way event, the probability distribution is parametrized by k parameters  $p_1, \ldots, p_k$  (k-1) independent parameters) where

$$\sum_{i=1}^{k} p_i = 1$$

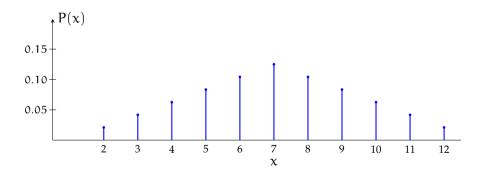
$$E[x_i] = p_i$$

$$Var(x_i) = p_i(1 - p_i)$$

• Similar to Bernoulli–binomial generalization, *multinomial* distribution is the generalization of categorical distribution to n trials

## Categorical distribution example

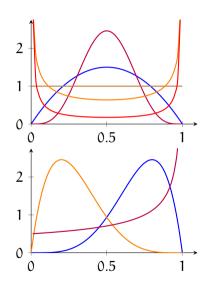
sum of the outcomes from roll of two fair dice



#### Beta distribution

- Beta distribution is defined in range [0, 1]
- It is characterized by two parameters  $\alpha$  and  $\beta$

$$p(x) = \frac{x^{\alpha - 1} (1 - x)^{\beta - 1}}{\frac{\Gamma(\alpha)\Gamma(\beta)}{\Gamma(\alpha + \beta)}}$$



#### Beta distribution

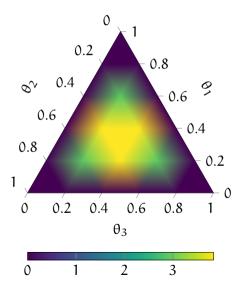
where do we use it

- A common use is the random variables whose values are probabilities
- Particularly important in Bayesian methods as a conjugate prior of Bernoulli and Binomial distributions
- The *Dirichlet distribution* generalizes Beta distribution to k-dimensional vectors whose components are in range (0, 1) and  $||x||_1 = 1$ .
- Dirichlet distribution is used often in NLP, e.g., *latent Dirichlet allocation* is a well know method for topic modeling

29 / 57

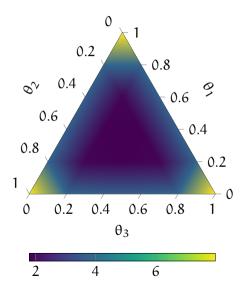
### Example Dirichlet distributions

 $\theta = (2, 2, 2)$ 



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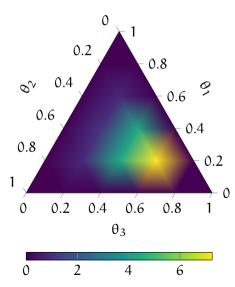
 $\theta = (0.8, 0.8, 0.8)$ 



30 / 57

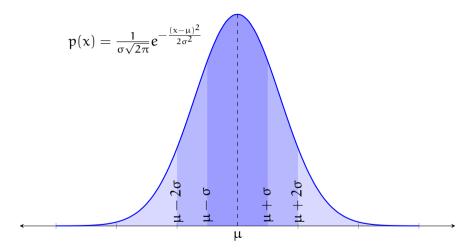
### Example Dirichlet distributions

 $\theta = (2, 2, 4)$ 



30 / 57

### Gaussian (normal) distribution



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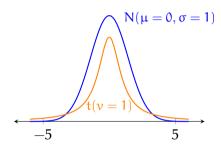
#### Short detour: central limit theorem

Central limit theorem states that the sum of a large number of independent and identically distributed variables (i.i.d.) is normally distributed.

- Expected value (average) of means of samples from any distribution will be distributed normally
- Many (inference) methods in statistics and machine learning work because of this fact

#### Student's t-distribution

- T-distribution is another important distribution
- It is similar to normal distribution, but it has heavier tails
- It has one parameter: degree of freedom (v)



## Joint and marginal probability

Two or more random variables form a *joint probability distribution*.

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Two or more random variables form a *joint probability distribution*.

#### An example with letter bigrams:

	a	b	c	d	e	f	g	h
a	0.04	0.02	0.02	0.03	0.05	0.01	0.02	0.06
b	0.01	0.00	0.00	0.00	0.01	0.00	0.00	0.01
c	0.02	0.00	0.00	0.00	0.01	0.00	0.00	0.01
d	0.02	0.00	0.00	0.01	0.02	0.00	0.01	0.02
e	0.06	0.02	0.01	0.03	0.08	0.01	0.01	0.07
f	0.00	0.00	0.00	0.00	0.01	0.00	0.00	0.01
$\mathbf{g}$	0.01	0.00	0.00	0.01	0.02	0.00	0.01	0.02
h	0.08	0.00	0.00	0.01	0.10	0.00	0.01	0.02

34 / 57

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c	0.02	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.05
d	0.02	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.08
e	0.06	0.02	0.01	0.03	0.08	0.01	0.01	0.07	0.29
f	0.00	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.02
$\mathbf{g}$	0.01	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.07
h	0.08	0.00	0.00	0.01	0.10	0.00	0.01	0.02	0.22
	0.23	0.04	0.05	0.08	0.29	0.02	0.07	0.22	

## Expected values of joint distributions

$$E[f(X,Y)] = \sum_{x} \sum_{y} P(x,y)f(x,y)$$

## Expected values of joint distributions

$$\begin{split} E[f(X,Y)] &= \sum_{x} \sum_{y} P(x,y) f(x,y) \\ \mu_{X} &= E[X] = \sum_{x} \sum_{y} P(x,y) x \\ \mu_{Y} &= E[Y] = \sum_{x} \sum_{y} P(x,y) y \end{split}$$

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We can simplify the notation by vector notation, for  $\mu = (\mu_x, \mu_y)$ ,

$$\mu = \sum_{\mathbf{x} \in \mathsf{XY}} \mathsf{xP}(\mathbf{x})$$

where vector  $\mathbf{x}$  ranges over all possible combinations of the values of random variables  $\mathbf{X}$  and  $\mathbf{Y}$ .

### Variances of joint distributions

$$\sigma_X^2 = \sum_x \sum_y P(x,y) (x - \mu_X)^2$$

$$\sigma_Y^2 = \sum_x \sum_y P(x,y) (y-\mu_Y)^2$$

### Variances of joint distributions

$$\begin{split} \sigma_X^2 &= \sum_x \sum_y P(x,y)(x-\mu_X)^2 \\ \sigma_Y^2 &= \sum_x \sum_y P(x,y)(y-\mu_Y)^2 \\ \sigma_{XY} &= \sum_x \sum_y P(x,y)(x-\mu_X)(y-\mu_Y) \end{split}$$

• The last quantity is called *covariance* which indicates whether the two variables vary together or not

### Variances of joint distributions

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Again, using vector/matrix notation we can define the *covariance matrix* ( $\Sigma$ ) as

$$\Sigma = E[(x - \mu)^2]$$

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36 / 57

### Covariance and the covariance matrix

$$\mathbf{\Sigma} = \begin{bmatrix} \sigma_{\mathbf{X}}^2 & \sigma_{\mathbf{XY}} \\ \sigma_{\mathbf{YX}} & \sigma_{\mathbf{Y}}^2 \end{bmatrix}$$

- The main diagonal of the covariance matrix contains the variances of the individual variables
- Non-diagonal entries are the covariances of the corresponding variables
- Covariance matrix is symmetric ( $\sigma_{XY} = \sigma_{YX}$ )
- For a joint distribution of k variables we have a covariance matrix of size  $k \times k$

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#### Correlation

Correlation is a normalized version of covariance

$$r = \frac{\sigma_{XY}}{\sigma_{X}\sigma_{Y}}$$

Correlation coefficient (r) takes values between -1 and 1

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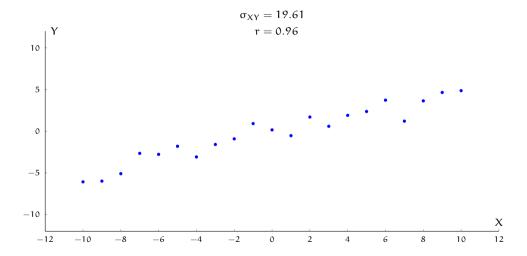
$$r = \frac{\sigma_{XY}}{\sigma_X \sigma_Y}$$

Correlation coefficient (r) takes values between -1 and 1

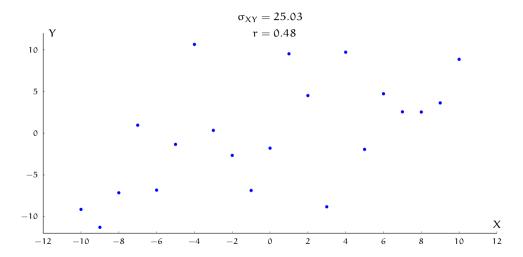
- 1 Perfect positive correlation.
- (0,1) positive correlation: x increases as y increases.
  - 0 No correlation, variables are independent.
- (-1,0) negative correlation: x decreases as y increases.
  - -1 Perfect negative correlation.

Note: like covariance, correlation is a symmetric measure.

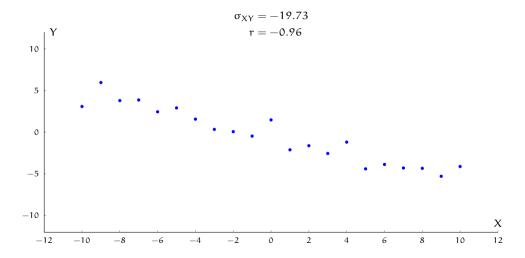
### Correlation: visualization (1)



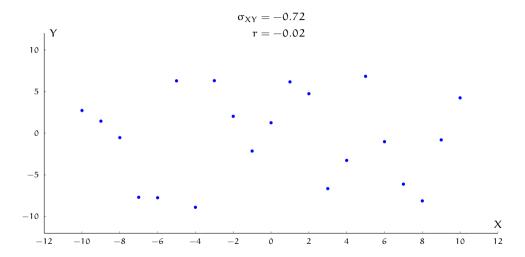
## Correlation: visualization (2)



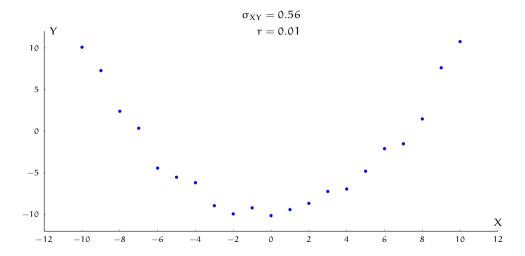
### Correlation: visualization (3)



## Correlation: visualization (4)



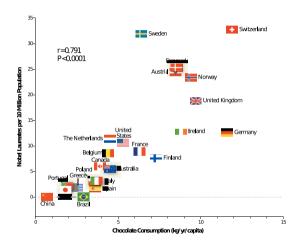
## Correlation: visualization (5)



# Correlation and independence

- Statistical (in)dependence is an important concept (in ML)
- The correlation (or covariance) of independent random variables is 0
- The reverse is not true: 0 correlation does not imply independence
- Correlation measures a linear dependence (relationship) between two variables, a non-linear dependence is not measured by correlation

## Short divergence: correlation and causation



From Messerli (2012).

# Conditional probability

In our letter bigram example, given that we know that the first letter is **e**, what is the probability of second letter being **d**?

	a	b	c	d	e	f	$\mathbf{g}$	h	
a	0.04	0.02	0.02	0.03	0.05	0.01	0.02	0.06	0.23
b	0.01	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.04
c	0.02	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.05
d	0.02	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.08
e	0.06	0.02	0.01	0.03	0.08	0.01	0.01	0.07	0.29
f	0.00	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.02
$\mathbf{g}$	0.01	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.07
h	0.08	0.00	0.00	0.01	0.10	0.00	0.01	0.02	0.22
	0.23	0.04	0.05	0.08	0.29	0.02	0.07	0.22	

$$P(L_1 = e, L_2 = d) = 0.025940365$$

$$P(L_1 = e) = 0.28605090$$

## Conditional probability

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b	0.01	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.04
c	0.02	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.05
d	0.02	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.08
e	0.06	0.02	0.01	0.03	0.08	0.01	0.01	0.07	0.29
f	0.00	0.00	0.00	0.00	0.01	0.00	0.00	0.01	0.02
$\mathbf{g}$	0.01	0.00	0.00	0.01	0.02	0.00	0.01	0.02	0.07
h	0.08	0.00	0.00	0.01	0.10	0.00	0.01	0.02	0.22
	0.23	0.04	0.05	0.08	0.29	0.02	0.07	0.22	
	1)	1 4250	10265					D/I	<b>\</b>

$$P(L_1 = e, L_2 = d) = 0.025940365$$

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$$P(L_2 = d | L_1 = e) = \frac{P(L_1 = e, L_2 = d)}{P(L_1 = e)}$$

## Conditional probability (2)

In terms of probability mass (or density) functions,

$$P(X \mid Y) = \frac{P(X, Y)}{P(Y)}$$

If two variables are independent, knowing the outcome of one does not affect the probability of the other variable:

$$P(X | Y) = P(X) \qquad P(X, Y) = P(X)P(Y)$$

More notes on notation/interpretation:

- P(X = x, Y = y) Probability that X = x and Y = y at the same time (joint probability)
  - P(Y = y) Probability of Y = y, for any value of  $X(\sum_{x \in X} P(X = x, Y = y))$ (marginal probability)
- $P(X = x \mid Y = y)$  Probability of X = x, given Y = y (conditional probability)

# Bayes' rule

$$P(X \mid Y) = \frac{P(Y \mid X)P(X)}{P(Y)}$$

- This is a direct result of the axioms of the probability theory
- It is often useful as it 'inverts' the conditional probabilities
- The term P(X), is called prior
- The term P(Y | X), is called likelihood
- The term P(X | Y), is called posterior

We use a test t to determine whether a patient has COVID-19 (c)

• If a patient has c test is positive 99% of the time: P(t | c) = 0.99

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- What is the probability that a patient has c given t?

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$$P(c \mid t) = \frac{P(t \mid c)P(c)}{P(t)} = \frac{P(t \mid c)P(c)}{P(t \mid c)P(c) + P(t \mid \neg c)P(\neg c)}$$

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$$P(c \mid t) = \frac{P(t \mid c)P(c)}{P(t)} = \frac{P(t \mid c)P(c)}{P(t \mid c)P(c) + P(t \mid \neg c)P(\neg c)} = 0.09$$

#### Chain rule

We rewrite the relation between the joint and the conditional probability as

$$P(X, Y) = P(X \mid Y)P(Y)$$

We can also write the same quantity as,

$$P(X,Y) = P(Y | X)P(X)$$

For more than two variables, one can write

$$P(X, Y, Z) = P(Z | X, Y)P(Y | X)P(X) = P(X | Y, Z)P(Y | Z)P(Z) = ...$$

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In general, for any number of random variables, we can write

$$P(X_1, X_2, ..., X_n) = P(X_1 | X_2, ..., X_n) P(X_2, ..., X_n)$$

If two random variables are conditionally independent:

$$P(X,Y \mid Z) = P(X \mid Z)P(Y \mid Z)$$

If two random variables are conditionally independent:

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This is often used for simplifying the statistical models. For example in spam filtering with *naive Bayes* classifier, we are interested in

$$P(w_1, w_2, w_3 \mid \text{spam})$$

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$$P(w_1, w_2, w_3 | spam) = P(w_1 | w_2, w_3, spam)P(w_2 | w_3, spam)P(w_3 | spam)$$

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$$P(w_1, w_2, w_3 | spam) = P(w_1 | w_2, w_3, spam)P(w_2 | w_3, spam)P(w_3 | spam)$$

with the assumption that occurrences of words are independent of each other given we know the email is spam or not,

$$P(w_1, w_2, w_3 \mid \text{spam}) = P(w_1 \mid \text{spam})P(w_2 \mid \text{spam})P(w_3 \mid \text{spam})$$

#### Continuous random variables

#### some reminders

The rules and quantities we discussed above apply to continuous random variables with some differences

- For continuous variables, P(X = x) = 0
- We cannot talk about probability of the variable being equal to a single real number
- But we can define probabilities of ranges
- For all formulas we have seen so far, replace summation with integrals
- Probability of a range:

$$P(a < X < b) = \int_{a}^{b} p(x) dx$$

52 / 57

#### Multivariate continuous random variables

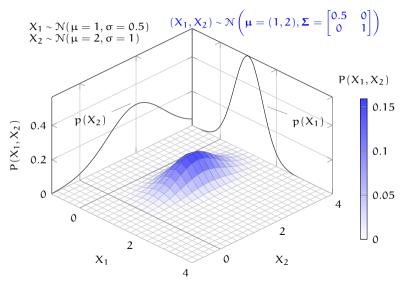
• Joint probability density

$$p(X,Y) = p(X \mid Y)p(Y) = p(Y \mid X)p(X)$$

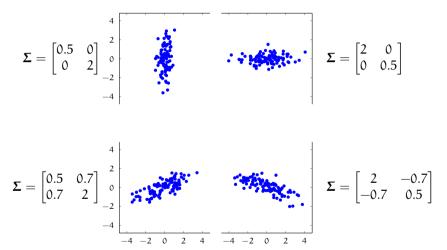
• Marginal probability

$$P(X) = \int_{-\infty}^{\infty} p(x, y) dy$$

#### Multivariate Gaussian distribution



## Samples from bi-variate normal distributions



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## Summary: some keywords

- Probability, sample space, outcome, event
- Random variables: discrete and continuous
- Probability mass function
- Probability density function
- Cumulative distribution function
- Expected value
- Variance / standard deviation
- Median and mode
- Skewness of a distribution

- Joint and marginal probabilities
- Covariance, correlation
- Conditional probability
- Bayes' rule
- Chain rule
- Some well-known probability distributions:

Bernoulli binomial categorical multinomial beta Dirichlet

Gaussian Student's t

#### Next

Wed Information theory
Mon ML Intro / regression
Wed Classification

# References and further reading

- MacKay (2003) covers most of the topics discussed in a way quite relevant to machine learning. The complete book is available freely online (see the link below)
- See Grinstead and Snell (2012) a more conventional introduction to probability theory. This book is also freely available
- For an influential, but not quite conventional approach, see Jaynes (2007)



Chomsky, Noam~(1968).~"Quine's~empirical~assumptions".~In: Synthese~19.1, pp.~53-68.~doi:~10.1007/BF00568049



Grinstead, Charles Miller and James Laurie Snell (2012). Introduction to probability. American Mathematical Society. ISBN: 9780821894149. URL: http://www.dartmouth.edu/~chance/teaching\_aids/books\_articles/probability\_book/book.html.



Jaynes, Edwin T (2007). Probability Theory: The Logic of Science. Ed. by G. Larry Bretthorst. Cambridge University Press. ISBN: 978-05-2159-271-0.



MacKay, David J. C. (2003). Information Theory, Inference and Learning Algorithms. Cambridge University Press. ISBN: 978-05-2164-298-9. URL: http://www.inference.phy.cam.ac.uk/itprnn/book.html.



Messerli, Franz H (2012). "Chocolate consumption, cognitive function, and Nobel laureates". In: The New England journal of medicine 367.16, pp. 1562–1564.