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### **Multinomial Distribution - Motivation**

Suppose we modified assumption (1) of the binomial distribution to allow for more than two outcomes.

For example, suppose that for the family with parents that are heterozygote carriers of a recessive trait, we are interested in knowing the probability of

 $Q_1$ : One of their n=3 offspring will be unaffected (AA), 1 will be affected (aa) and one will be a carrier (Aa),

 $Q_2$ :All of their offspring will be carriers,

 $Q_3$ :Exactly two of their offspring will be affected (aa) and one will be a carrier.

#### **Multinomial Distribution - Motivation**

For each child, we can represent these possibilities with three indicator variables for the *i*-th child as

 $Y_{il} = 1$  if unaffected (AA), & 0 otherwise  $Y_{i2} = 1$  if carrier (Aa), & 0 otherwise  $Y_{i3} = 1$  if affected (aa), & 0 otherwise

Notice only one of the three  $Y_{il}$ ,  $Y_{i2}$ ,  $Y_{i3}$  can be equal to 1, so  $\Sigma_j Y_{ij} = 1$ .

For the binomial distribution with 2 outcomes, there are  $2^n$  unique outcomes in n trials. In the family with n=3 children, there are  $2^3 = 8$  unique outcomes.

For the multinomial distribution with n trials and only 3 outcomes, the number of unique outcomes is  $3^n$ . For our small family, that's  $3^3$ =27 outcomes.

#### **Possible Outcomes**

Combinations: As with the binomial, there are different ways to arrange possible outcomes from a total of *n* objects (trials) if order doesn't matter. For the multinomial distribution, the combinations are summarized as

$$C_{k}^{n} = \frac{n!}{k_{1}! k_{2} ... k_{J}!}$$

where the  $k_j$  (j=1,2,...,J) correspond to the totals for the different outcomes.

E.g. (*n*=2 offspring)

### Child number

1	2	Outcomes		
$\overline{AA}$	AA	2 unaffected, 0 carrier, 0 affected		
AA	Aa	1 unaffected, 1 carrier, 0 affected		
Aa	AA	1 unaffected, 1 carrier, 0 affected		
AA	aa	1 unaffected, 0 carrier, 1 affected		
aa	AA	1 unaffected, 0 carrier, 1affected		
Aa	Aa	0 unaffected, 2 carrier, 0 affected		
aa	Aa	0 unaffected, 1 carrier, 1 affected		
Aa	aa	0 unaffected, 1 carrier, 1 affected		
aa	aa	0 unaffected, 0 carrier, 2 affected		

For the case of n=2 offspring (i.e., trials), what are the probabilities of these outcomes?

E.g. (
$$n=2$$
,  $k_1$ =unaffected,  $k_2$ =carrier,  $k_3$ =affected)

### Child number

1	2	Outcomes	# ways
$\mathbf{p}_1$	$p_1$	$k_1 = 2, k_2 = 0, k_3 = 0$	<b>←</b> ——1 •
$\mathbf{p}_1$	$p_2$	$k_1 = 1, k_2 = 1, k_3 = 0$	2
$\mathbf{p}_2$	$\mathbf{p}_1$	$k_1 = 1, k_2 = 1, k_3 = 0$	
$\mathbf{p}_1$	$\mathbf{p}_3$	$k_1 = 1, k_2 = 0, k_3 = 1$	2
$p_3$	$\mathbf{p}_1$	$k_1 = 1, k_2 = 0, k_3 = 1$	
$p_2$	$p_2$	$k_1 = 0, k_2 = 2, k_3 = 0$	<b>←</b> 1
$p_3$	$p_2$	$k_1 = 0, k_2 = 1, k_3 = 1$	2
$p_2$	$p_3$	$k_1 = 0, k_2 = 1, k_3 = 1$	+
$p_3$	$p_3$	$k_1 = 0, k_2 = 0, k_3 = 2$	← 1

For each possible outcome, the probability  $Pr[Y_1=k_1, Y_2=k_2, Y_3=k_3]$  is

$$p_1^{k1}p_2^{k2}p_3^{k3}$$

There are  $\frac{n!}{k_j!}$  sequences for each probability, so in general...

#### **Multinomial Probabilities**

What is the probability that a multinomial random variable with **n** trials and success probabilities  $p_1$ ,  $p_2$ , ...,  $p_J$  will yield exactly  $k_1$ ,  $k_2$ , ...  $k_J$  successes?

$$P(Y_1 = k_1, Y_2 = k_2, ..., Y_J = k_J) = \frac{n!}{k_1! k_2! ... k_J!} p_1^{k_1} p_2^{k_2} \cdots p_J^{k_J}$$

### **Assumptions:**

- 1) J possible outcomes only one of which can be a success (1) a given trial.
- 2) The probability of success for each possible outcome,  $p_i$ , is the same from trial to trial.
- 3) The outcome of one trial has no influence on other trials (independent trials).
- 4) Interest is in the (sum) total number of "successes" over all the trials.

$$k_1 \mid k_2 \mid k_3 \mid k_4 \mid \cdot \cdot \cdot \mid k_{J-1} \mid k_J$$

 $n = \sum_{i} k_{i}$  is the total number of trials.

### **Multinomial Probabilities - Examples**

Returning to the original questions:

 $Q_1$ : One of n=3 offspring will be unaffected (AA), one will be affected (aa) and one will be a carrier (Aa) (recessive trait, carrier parents)?

**Solution:** For a given child, the probabilities of the three outcomes are:

$$p_1 = \Pr[AA] = 1/4,$$
  
 $p_2 = \Pr[Aa] = 1/2,$   
 $p_3 = \Pr[aa] = 1/4.$ 

We have

$$P(Y_1 = 1, Y_2 = 1, Y_3 = 1) = \frac{3!}{1!1!1!} p_1^1 p_2^1 p_3^1$$

$$= \frac{(3)(2)(1)}{(1)(1)} \left(\frac{1}{4}\right)^1 \left(\frac{1}{2}\right)^1 \left(\frac{1}{4}\right)^1$$

$$= \frac{3}{16} = 0.1875.$$

## Quiz

1. What is the probability that all three offspring will be carriers?

2. What is the probability that exactly two offspring will be affected and one a carrier?

#### **Solutions**

 $Q_2$ : What is the probability that all three offspring will be carriers?

$$P(Y_1 = 0, Y_2 = 3, Y_3 = 0) = \frac{3!}{0!3!0!} p_1^0 p_2^3 p_3^0$$

$$= \frac{(3)(2)(1)}{(3)(2)(1)} \left(\frac{1}{4}\right)^0 \left(\frac{1}{2}\right)^3 \left(\frac{1}{4}\right)^0$$

$$= \frac{1}{8} = 0.125.$$

 $Q_3$ : What is the probability that exactly two offspring will be affected and one a carrier?

$$P(Y_1 = 0, Y_2 = 1, Y_3 = 2) = \frac{3!}{0!1!2!} p_1^0 p_2^1 p_3^2$$

$$= \frac{(3)(2)(1)}{(2)(1)} \left(\frac{1}{4}\right)^0 \left(\frac{1}{2}\right)^1 \left(\frac{1}{4}\right)^2$$

$$= \frac{3}{32} = 0.09375.$$

### **Example - Mean and Variance**

It turns out that the (marginal) outcomes of the multinomial distribution are binomial. We can immediately obtain the means for each outcome (i.e., the j<sup>th</sup> cell)

MEAN: 
$$E[k_j] = E\left[\sum_{i=1}^{n} Y_{ij}\right] = \sum_{i=1}^{n} E[Y_{ij}]$$
  
=  $\sum_{i=1}^{n} p_i = np_j$ 

**VARIANCE:** 

$$V[k_{j}] = V \left[ \sum_{i=1}^{n} Y_{ij} \right] = \sum_{i=1}^{n} V[Y_{ij}]$$

$$= \sum_{i=1}^{n} p_{j} (1 - p_{j}) = np_{j} (1 - p_{j})$$

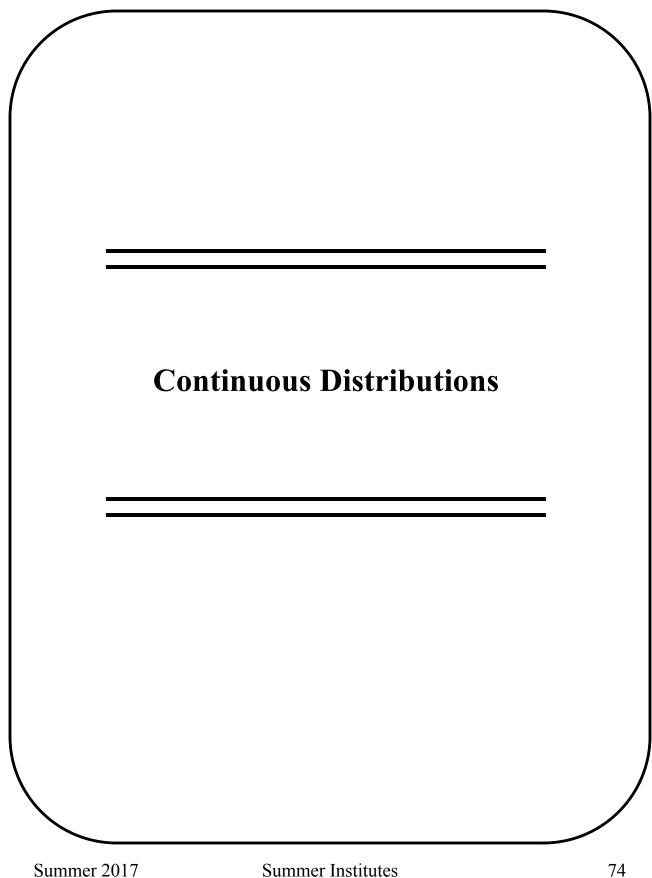
**COVARIANCE:** 

$$Cov[k_j, k_{j'}] = -np_j p_{j'}$$

## **Multinomial Distribution Summary**

### **Multinomial**

- 1. Discrete, bounded
- 2. Parameters  $n, p_1, p_2, ..., p_J$
- 3. Sum of *n* independent outcomes
- 4. Extends binomial distribution
- 5. Polytomous regression, contingency tables



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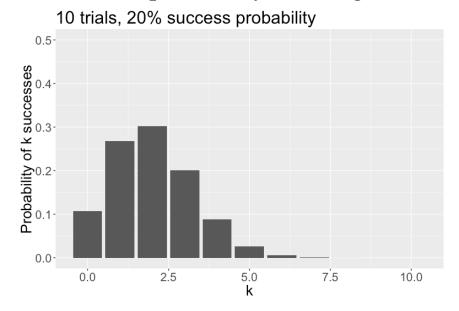
#### **Continuous Distributions**

For measurements like height and weight which can be measured with arbitrary precision, it does not make sense to talk about the probability of any single value. Instead we talk about the probability for an **interval**.

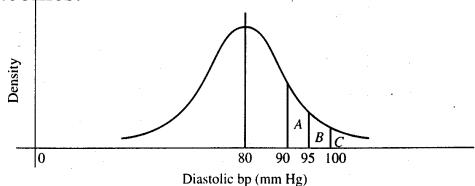
$$P[weight = 70.000kg] \approx 0$$
  
 $P[69.0kg \le weight \le 71.0kg] = 0.08$ 

For discrete random variables we had a probability mass function to give us the probability of each possible value. For continuous random variables we use a **probability density function** to tell us about the probability of obtaining a value within some interval.

With discrete probability distributions, we can determine the probability of a single outcome, e.g.:



With continuous probability distributions, we can determine the probability across a range of outcomes:



For any interval, the **area** under the curve represents the probability of obtaining a value in that interval.

### **Probability density function**

- 1. A function, typically denoted f(x), that gives probabilities based on the **area** under the curve.
- 2.  $f(x) \ge 0$
- 3. Total area under the function f(x) is 1.0.

$$\int f(x)dx = 1.0$$

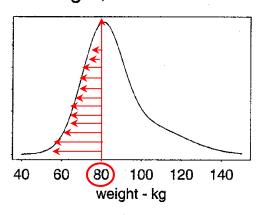
#### **Cumulative distribution function**

The <u>cumulative distribution function</u>, F(t), tells us the total probability less than some value t.

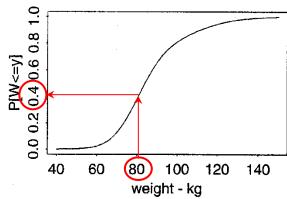
$$F(t) = P(X \le t)$$

This is analogous to the cumulative relative frequency.

### Weight, males 30-40



### Cumulative Dist Fn



$$Prob[wgt < 80] = 0.40$$
*Area under the curve*

### **Normal Distribution**

- A common probability model for continuous data
- Bell-shaped curve
- $\Rightarrow$  takes values between  $-\infty$  and  $+\infty$
- ⇒ symmetric about mean
- ⇒ mean=median=mode
- Examples (common but questionable!)birthweightsblood pressure

CD4 cell counts (perhaps transformed)

The normal distribution is more useful as a derived distribution, as we will see when we talk about the central limit theorem...

#### **Normal Distribution**

Specifying the mean and variance of a normal distribution completely determines the probability distribution function and, therefore, all probabilities.

The normal probability density function is:

$$f(x) = \frac{1}{\sigma\sqrt{2\pi}} \exp\left(-\frac{1}{2} \frac{(x-\mu)^2}{\sigma^2}\right)$$

where

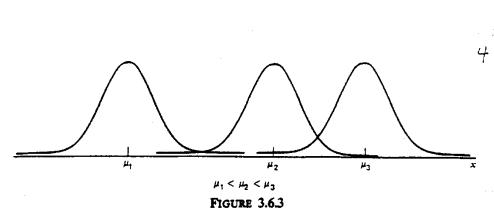
$$\pi \approx 3.14$$
 (a constant)

Notice that the normal distribution has two parameters:

 $\mu$  = the mean of X

 $\sigma$  = the standard deviation of X

We write  $X \sim N(\mu, \sigma^2)$ . The **standard normal** distribution is a special case where  $\mu = 0$  and  $\sigma = 1$ .



Three Normal Distributions with Different Means

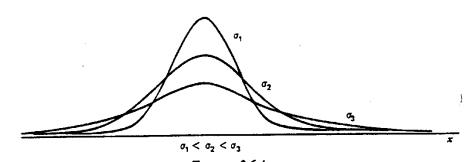
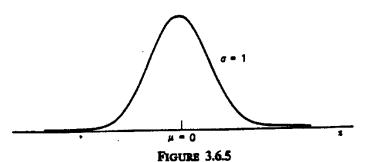


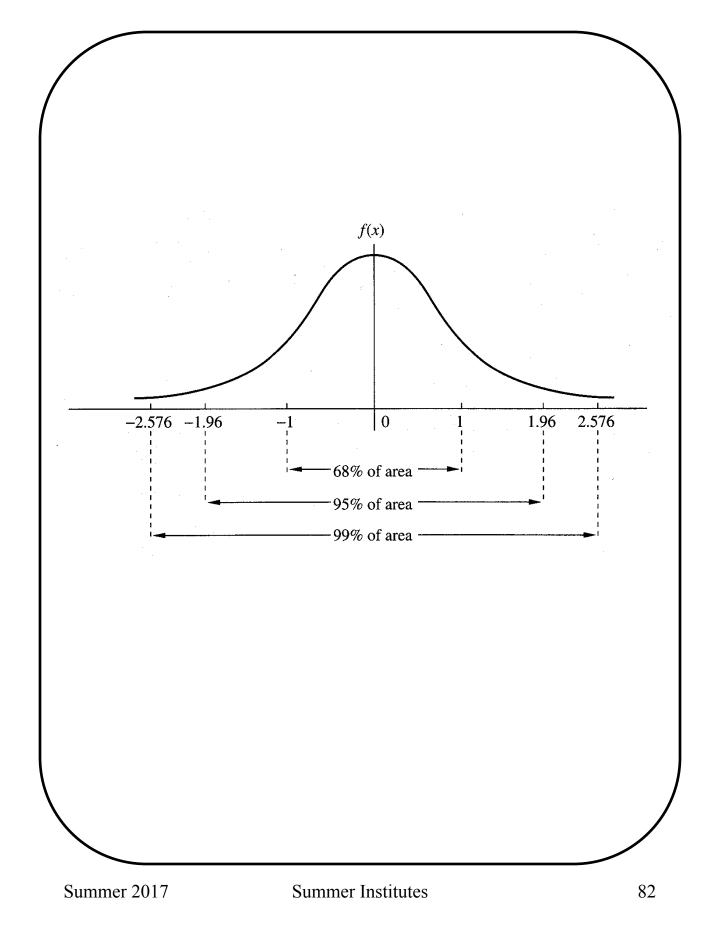
FIGURE 3.6.4

Three Normal Distributions with Different Standard Deviations



The Unit Normal Distribution

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## **Normal Distribution - Calculating Probabilities**

Example: Rosner 5.20

Serum cholesterol is approximately normally distributed with mean 219 mg/mL and standard deviation 50 mg/mL. If the clinically desirable range is < 200 mg/mL, then what proportion of the population falls in this range?

X = serum cholesterol in an individual.

$$\mu =$$

$$\sigma =$$

$$P[x < 200] = \int_{-\infty}^{200} \frac{1}{50\sqrt{2\pi}} \exp\left(-\frac{1}{2} \frac{(x - 219)^2}{50^2}\right) dx$$

negative values for cholesterol - huh?

## **Standard Normal Distribution - Calculating Probabilities**

First, let's consider the **standard normal** - N(0,1). We will usually use Z to denote a random variable with a standard normal distribution. The density of Z is

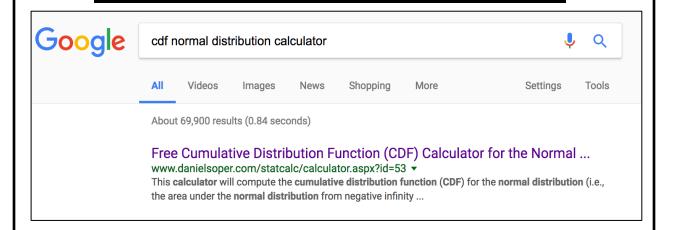
$$f(z) = \frac{1}{\sqrt{2\pi}} \exp\left(-\frac{1}{2}z^2\right)$$

and the **cumulative distribution** of Z is:

$$P(Z \le x) = \Phi(x) = \int_{-\infty}^{x} \frac{1}{\sqrt{2\pi}} \exp\left(-\frac{1}{2}z^{2}\right) dz$$

Any computing software can give you the values of f(z) and  $\Phi(z)$ 

# Standard Normal Distribution - Calculating Probabilities



## Cumulative Distribution Function (CDF) Calculator for the Normal Distribution

This calculator will compute the cumulative distribution function (CDF) for the normal distribution (i.e., the area under the normal distribution from negative infinity to x), given the upper limit of integration x, the mean, and the standard deviation.

Please enter the necessary parameter values, and then click 'Calculate'.

Mean: 0

Standard deviation: 1

x: 0.5

Calculate!

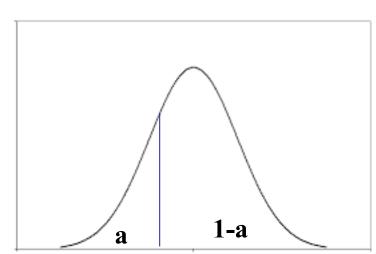
**Cumulative distribution function: 0.69146246** 

$$Pr(Z \le 0.5) = 0.69146$$

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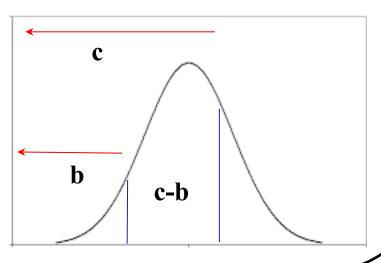
## Facts about probability distributions

$$P[Z \le z] = a$$
  
 $\Rightarrow P[Z > z] = 1-a$ 



$$P[Z \le x] = b, P[Z \le y] = c$$

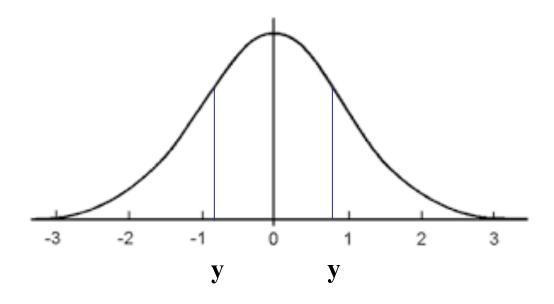
$$\Rightarrow$$
 Pr[x < Z \le y] = c-b



## Facts about the standard normal distribution

Because the N(0,1) distribution is symmetric around 0,

$$Pr[Z \le -y] = Pr[Z \ge y]$$



### **Quiz: 3 minutes**

Google "cdf normal distribution calculator" and find the following if  $Z \sim N(0,1)$ 

$$P[Z \le 1.65] =$$
 $P[Z \ge 0.5] =$ 
 $P[-1.96 \le Z \le 1.96] =$ 
 $P[-0.5 \le Z \le 2.0] =$ 

## **Solutions to Quiz**

$$P[Z \le 1.65] = 0.9505$$

$$P[Z \ge 0.5] = 1 - 0.6915 = 0.3085$$

$$P[-1.96 \le Z \le 1.96] = 0.975 - 0.025 = 0.95$$

$$P[-0.5 \le Z \le 2.0] = 0.9772 - 0.3085 = 0.6687$$

## **Converting to Standard Normal**

This solves the problem for the N(0,1) case. Do we need a special table for every  $(\mu,\sigma)$ ? No!

Define:  $X = \mu + \sigma Z$  where  $Z \sim N(0,1)$ 

$$1. E(X) = \mu + \sigma E(Z) = \mu$$

$$2. V(X) = \sigma^2 V(Z) = \sigma^2.$$

3. X is normally distributed!

Linear functions of normal RV's are also normal.

If 
$$X \sim N (\mu, \sigma^2)$$
 and  $Y = aX + b$   
then 
$$Y \sim N(a\mu + b, a^2\sigma^2)$$

## **Converting to Standard Normal**

How can we convert a  $N(\mu, \sigma^2)$  to a standard normal?

#### **Standardize:**

$$Z = \frac{X - \mu}{\sigma}$$

What is the mean and variance of Z?

1. 
$$E(Z) = (1/\sigma)E(X - \mu) = 0$$

2. 
$$V(Z) = (1/\sigma^2)V(X) = 1$$

## Normal Distribution - Calculating Probabilities

Return to cholesterol example (Rosner 5.20)

Serum cholesterol is approximately normally distributed with mean 219 mg/mL and standard deviation 50 mg/mL. If the clinically desirable range is < 200 mg/mL, then what proportion of the population falls in this range?

$$P(X < 200) = P\left(\frac{X - \mu}{\sigma} < \frac{200 - 219}{50}\right)$$

$$= P\left(Z < \frac{200 - 219}{50}\right)$$

$$= P(Z < -0.38)$$

$$= P(Z > 0.38)$$

$$= 0.3520.$$

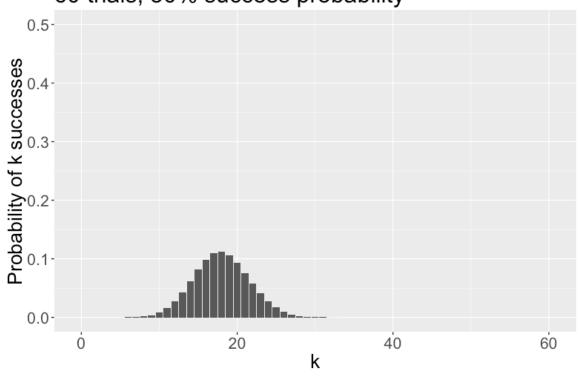
### **Example**

Suppose the prevalence of HPV in women 18 - 22 years old is 0.30. What is the probability that in a sample of 60 women from this population 9 or fewer would be infected?

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### **Binomial**

- When **np(1-p)** is "large" the normal may be used to approximate the binomial.
- $X \sim bin(n,p)$  E(X) = npV(X) = np(1-p)
- X is approximately N(np,np(1-p))

### **Example**

Suppose the prevalence of HPV in women 18 - 22 years old is 0.30. What is the probability that in a sample of 60 women from this population that 9 or less would be infected?

### Solution

X = number infected out of 60

 $X \sim Binomial(n=60, p=0.3)$ 

X close to Normal distribution with mean 60\*0.3=18 and variance 0.6\*0.3\*(1-0.3)=12.6

Cumulative Distribution Function (CDF) Calculator for the Normal Distribution

This calculator will compute the cumulative distribution function (CDF) for the normal distribution (i.e., the area under the normal distribution from negative infinity to x), given the upper limit of integration x, the mean, and the standard deviation.

Please enter the necessary parameter values, and then click 'Calculate'.

