

Survival Analysis

Session 1: Probabilities of Survival

Jonas Schöley

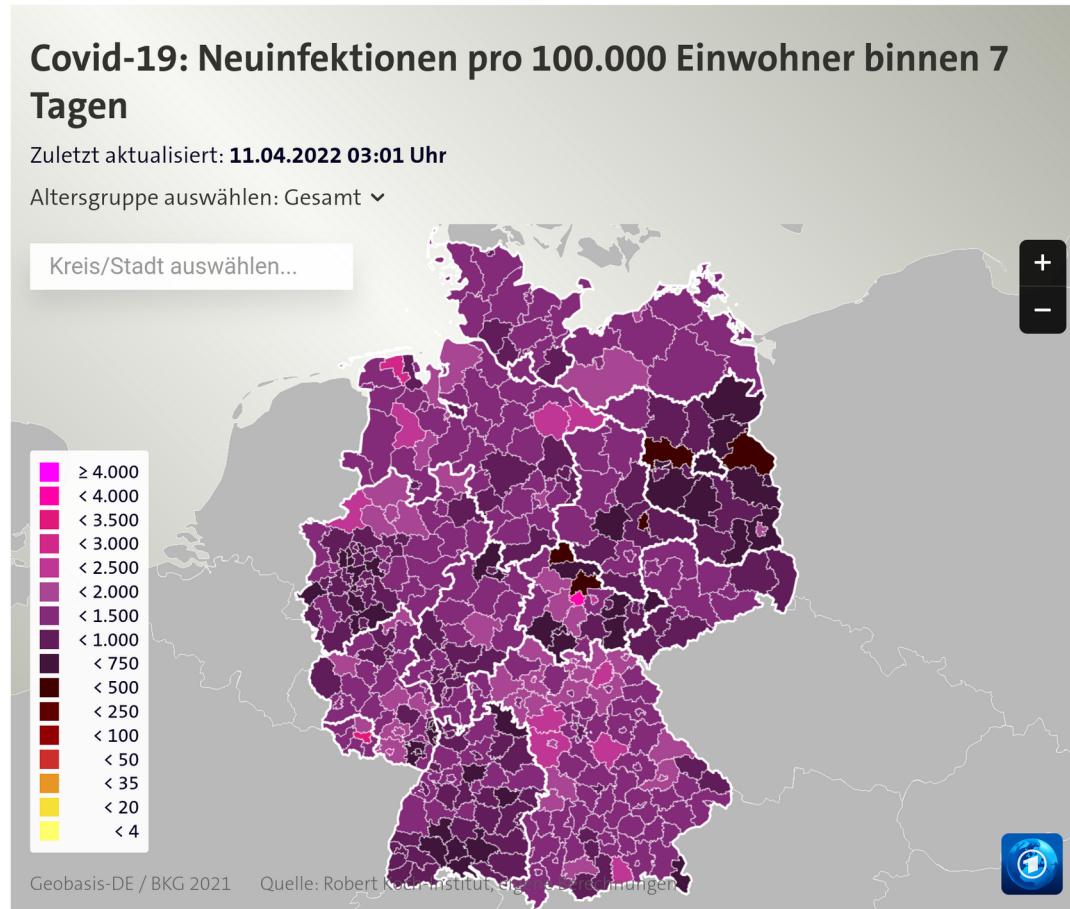
 @jschoeley

 0000-0002-3340-8518

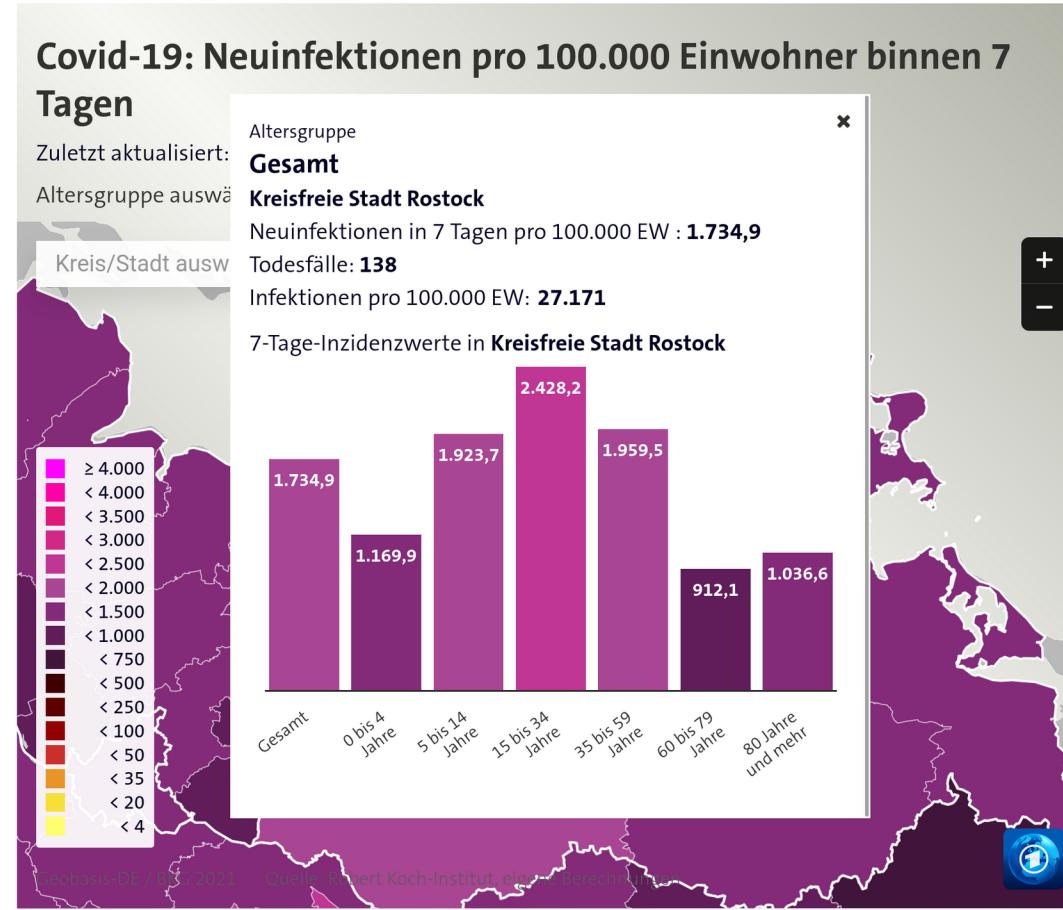
 j.schoeley@uni-rostock.de

CC-BY Jonas Schöley 2022

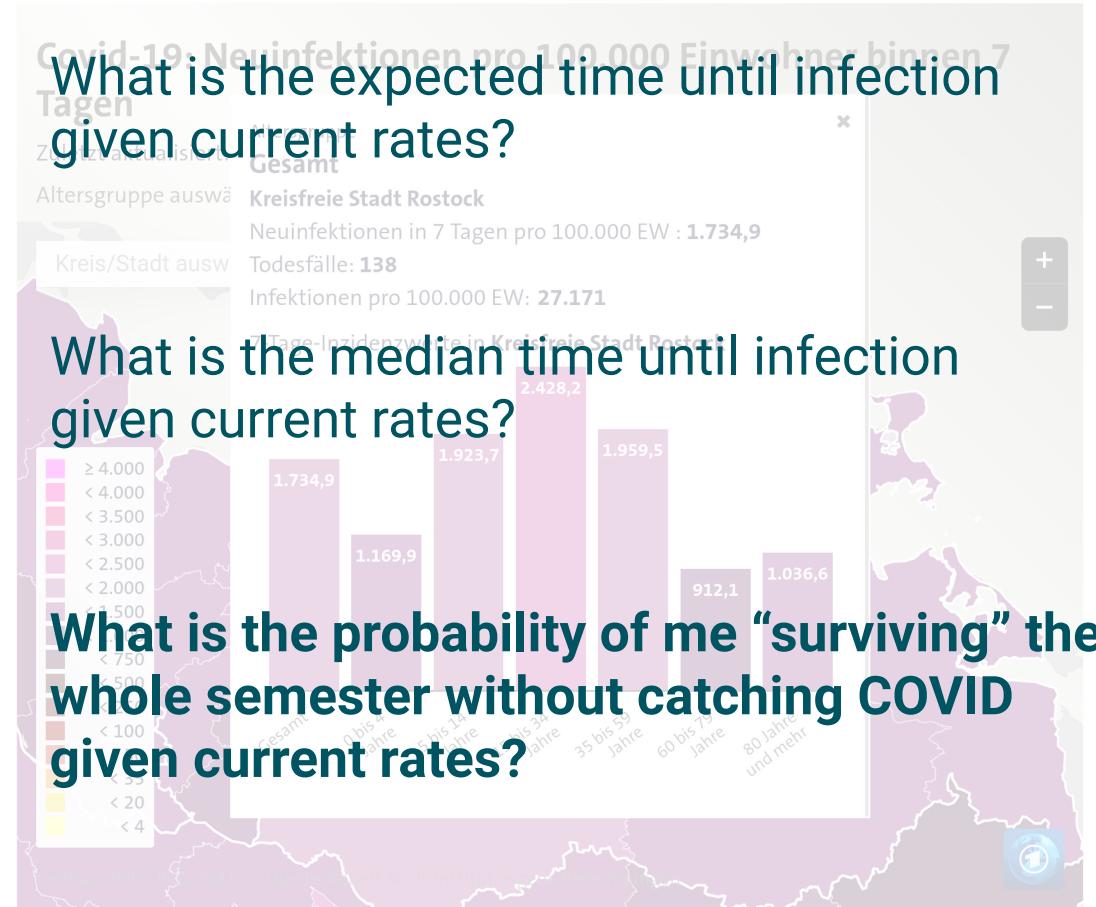
How long until infection?



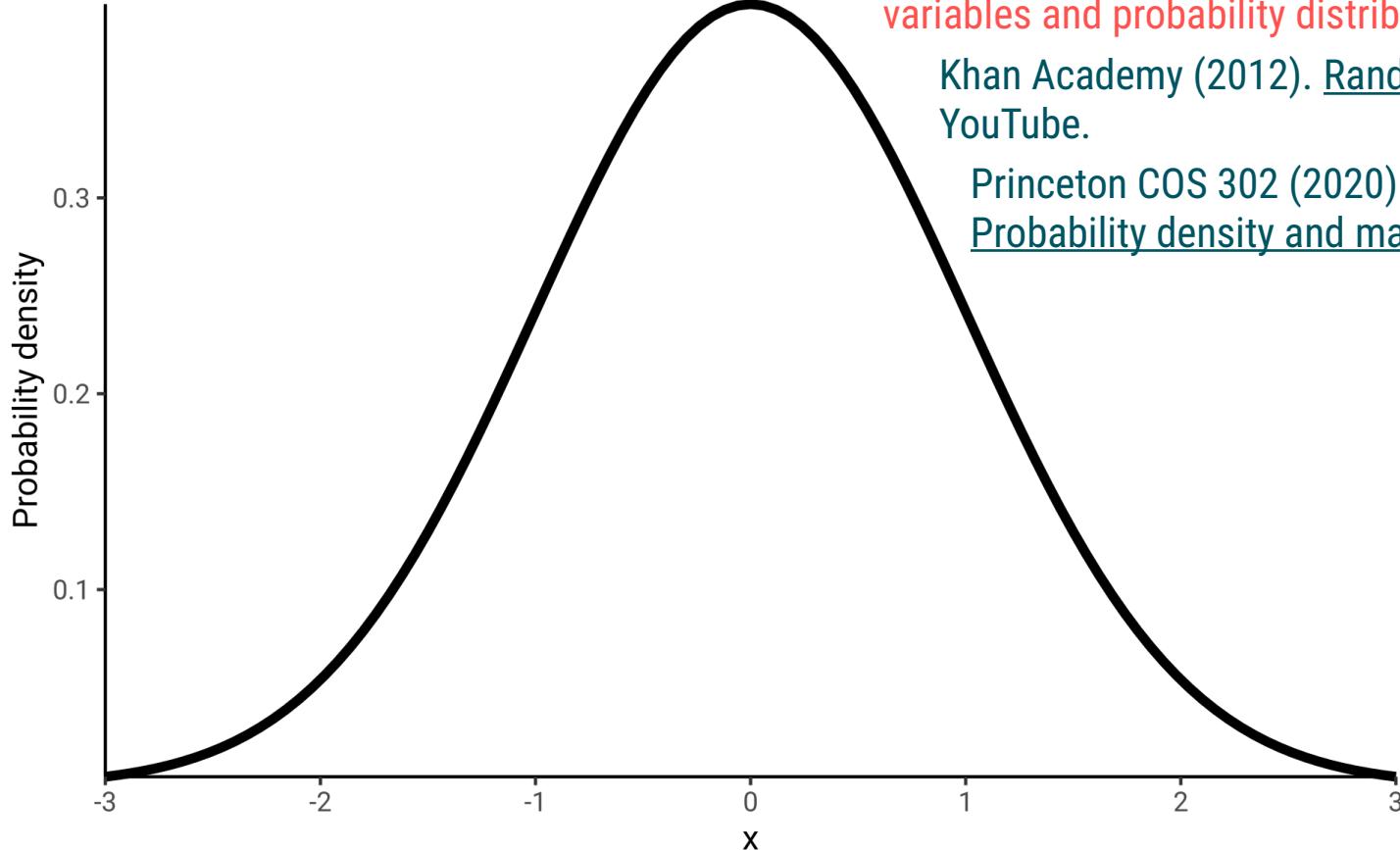
How long until infection?



How long until infection?



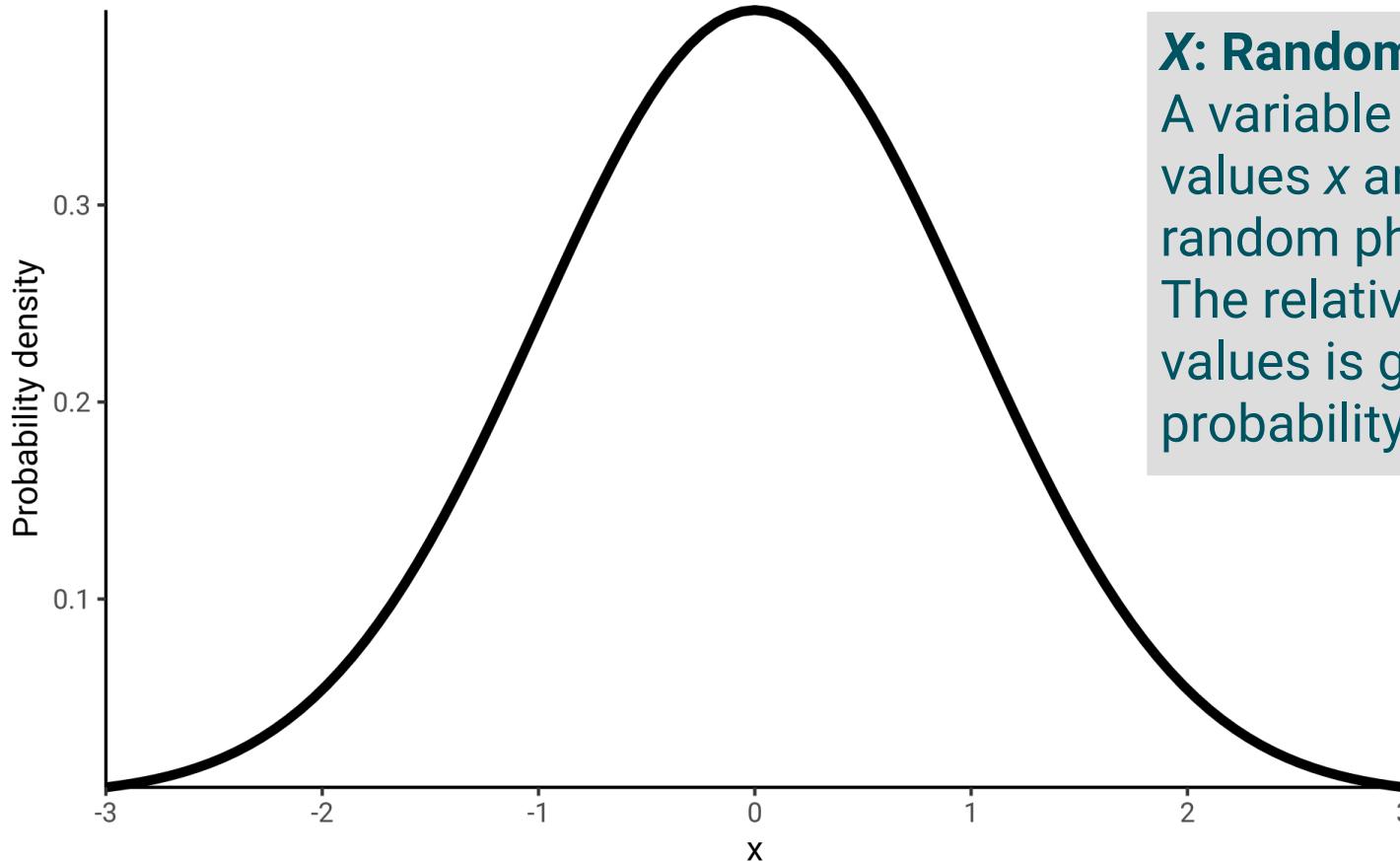
Recap: Random Variables & Probability Distributions



For refreshing your understanding of random variables and probability distributions watch Khan Academy (2012). [Random variables](#). YouTube.

Princeton COS 302 (2020). [Probability density and mass functions](#). YouTube.

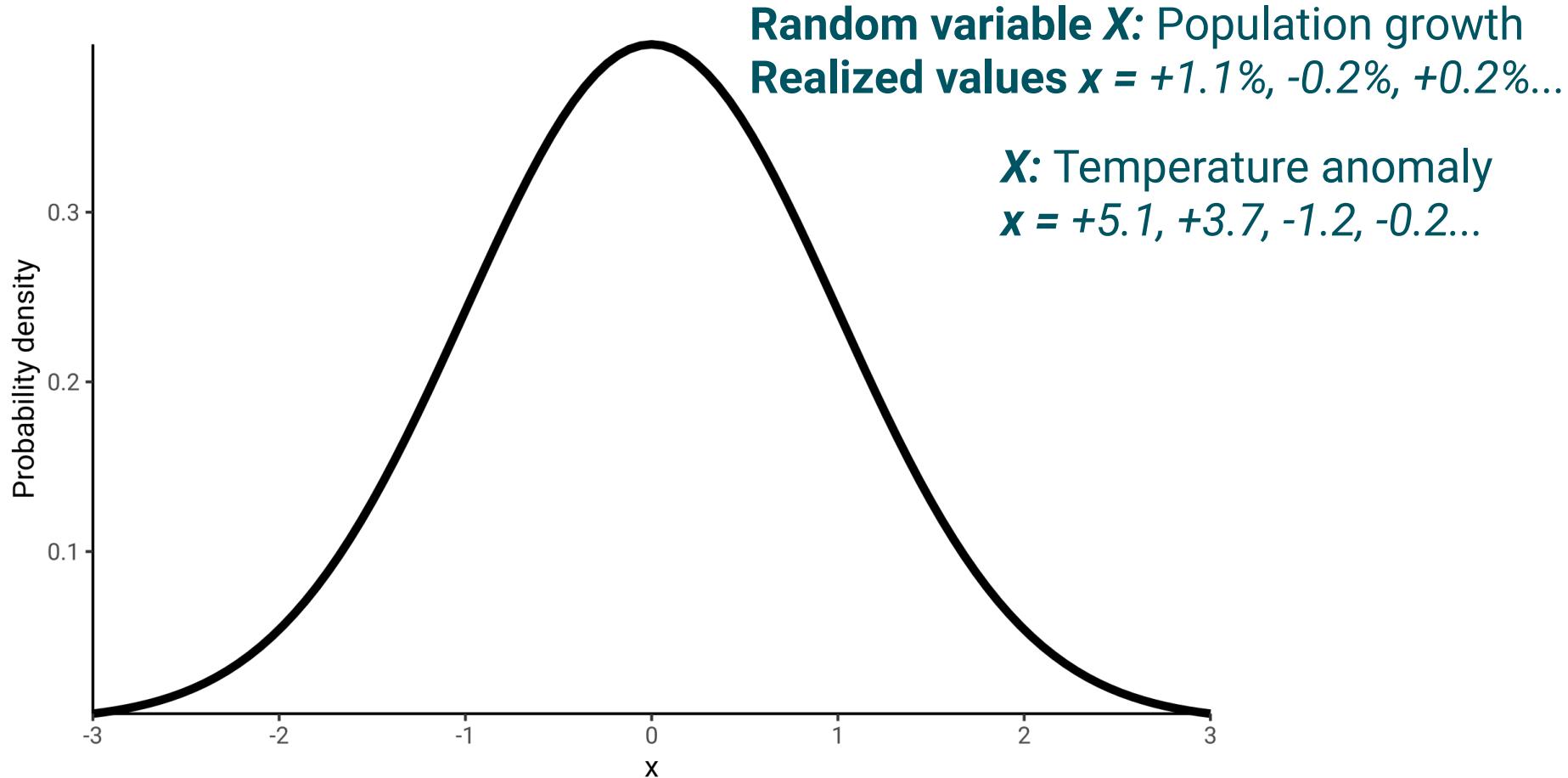
Recap: Random Variables & Probability Distributions



X: Random Variable

A variable whose possible values x are outcomes of a random phenomenon. The relative likelihood of values is given by the probability density $f_x(x)$.

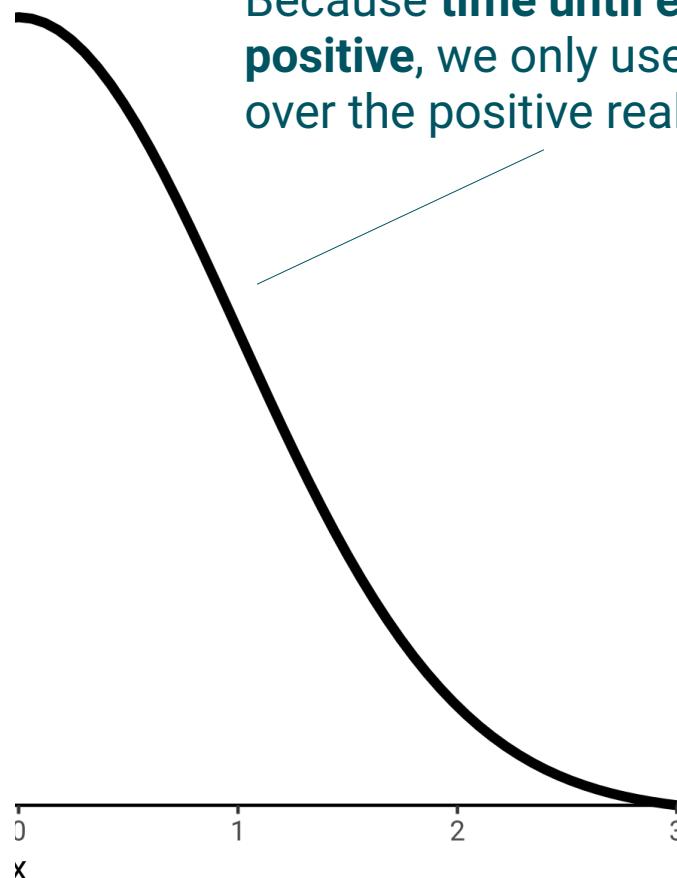
Recap: Random Variables & Probability Distributions



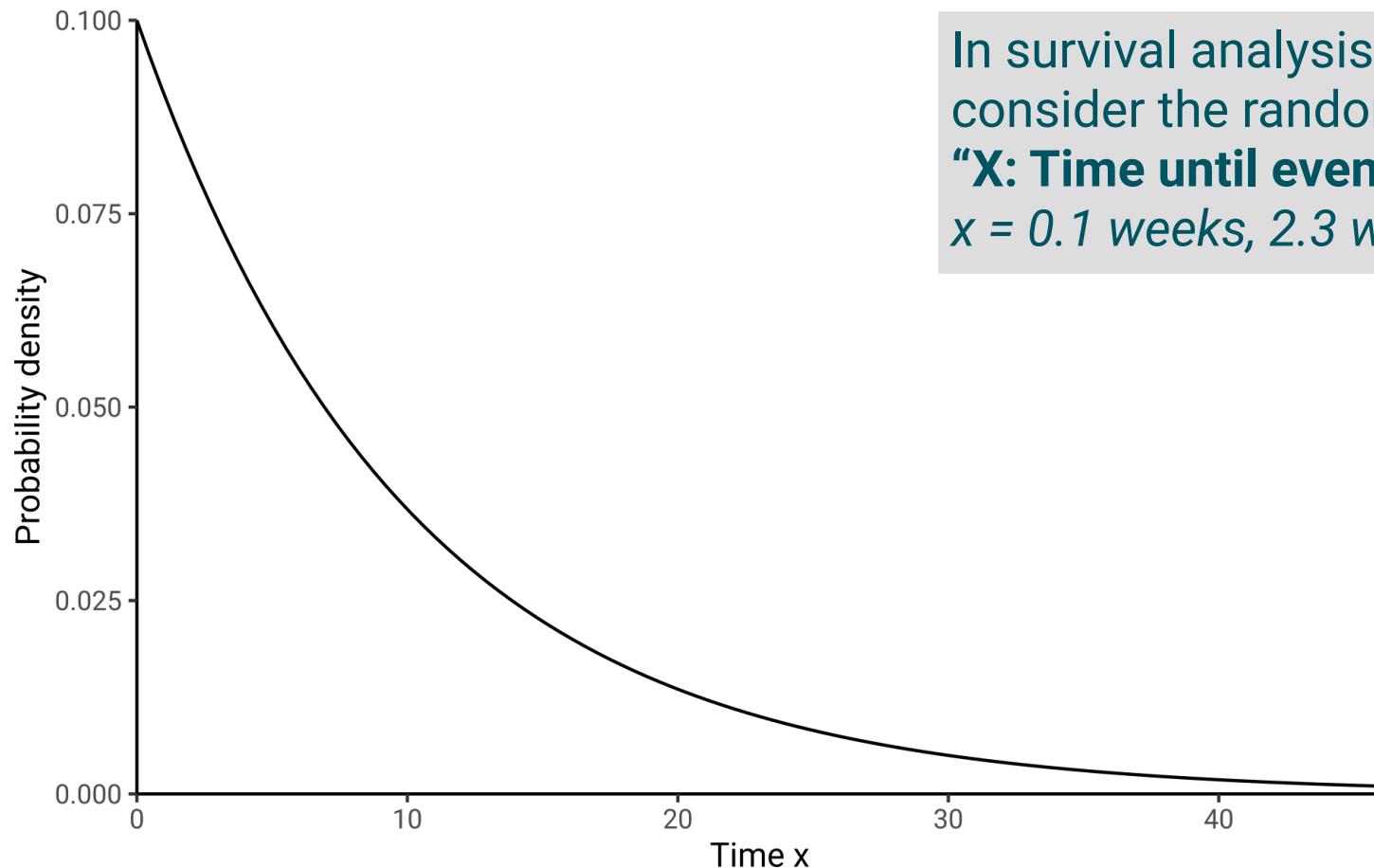
Survival Distributions

In survival analysis we consider the random variable
X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

Because time until event is always positive, we only use distributions over the positive real numbers.



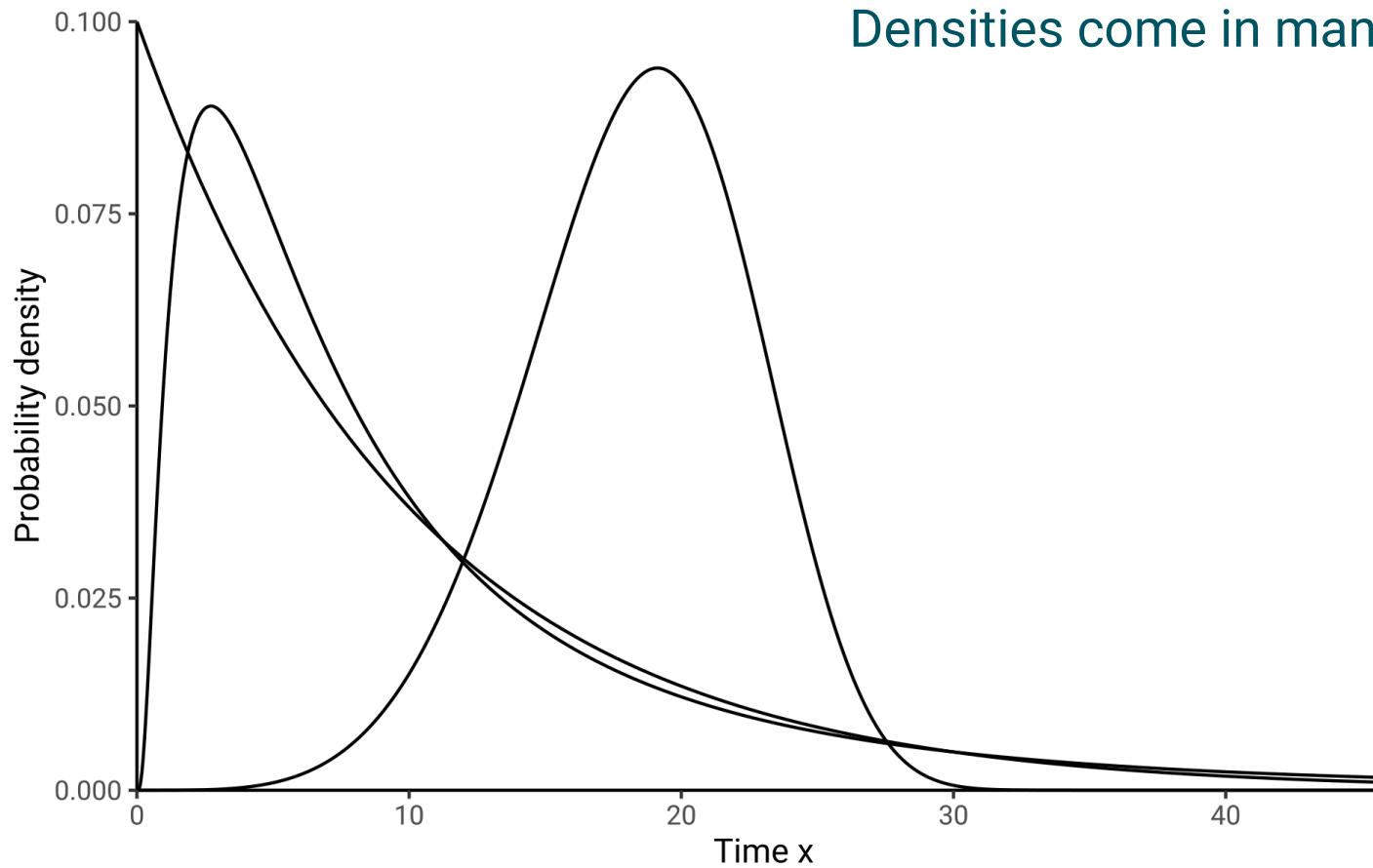
$f(x)$ Density Function



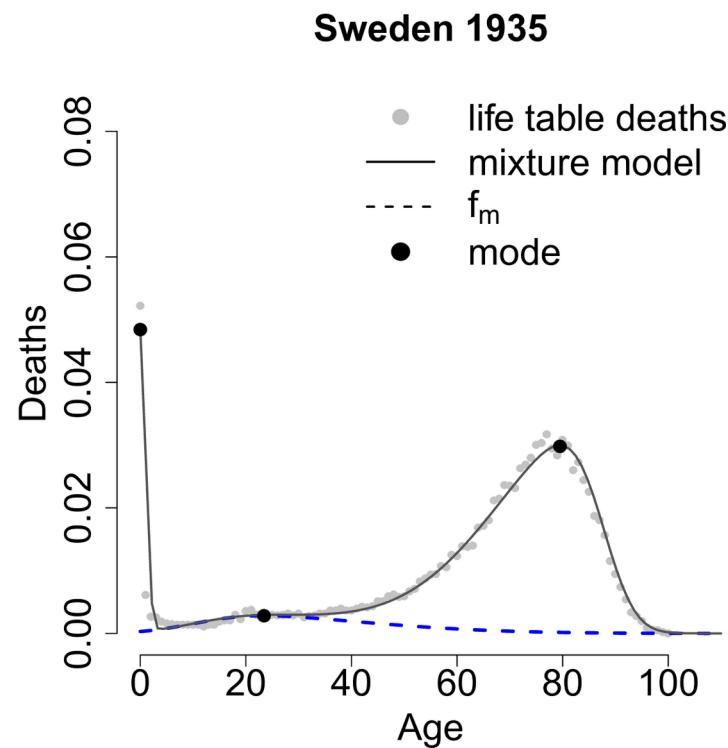
In survival analysis we consider the random variable
“X: Time until event”
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

$f(x)$ Density Function

Densities come in many shapes



$f(x)$ Density Function



Sweden 2011
Densities come in many shapes

Here's the distribution of the random variable "years until death" in Sweden.

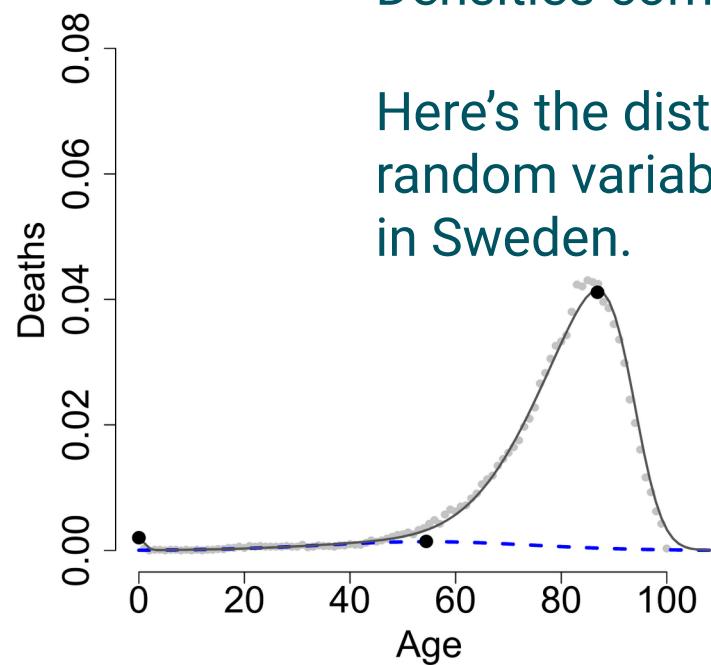
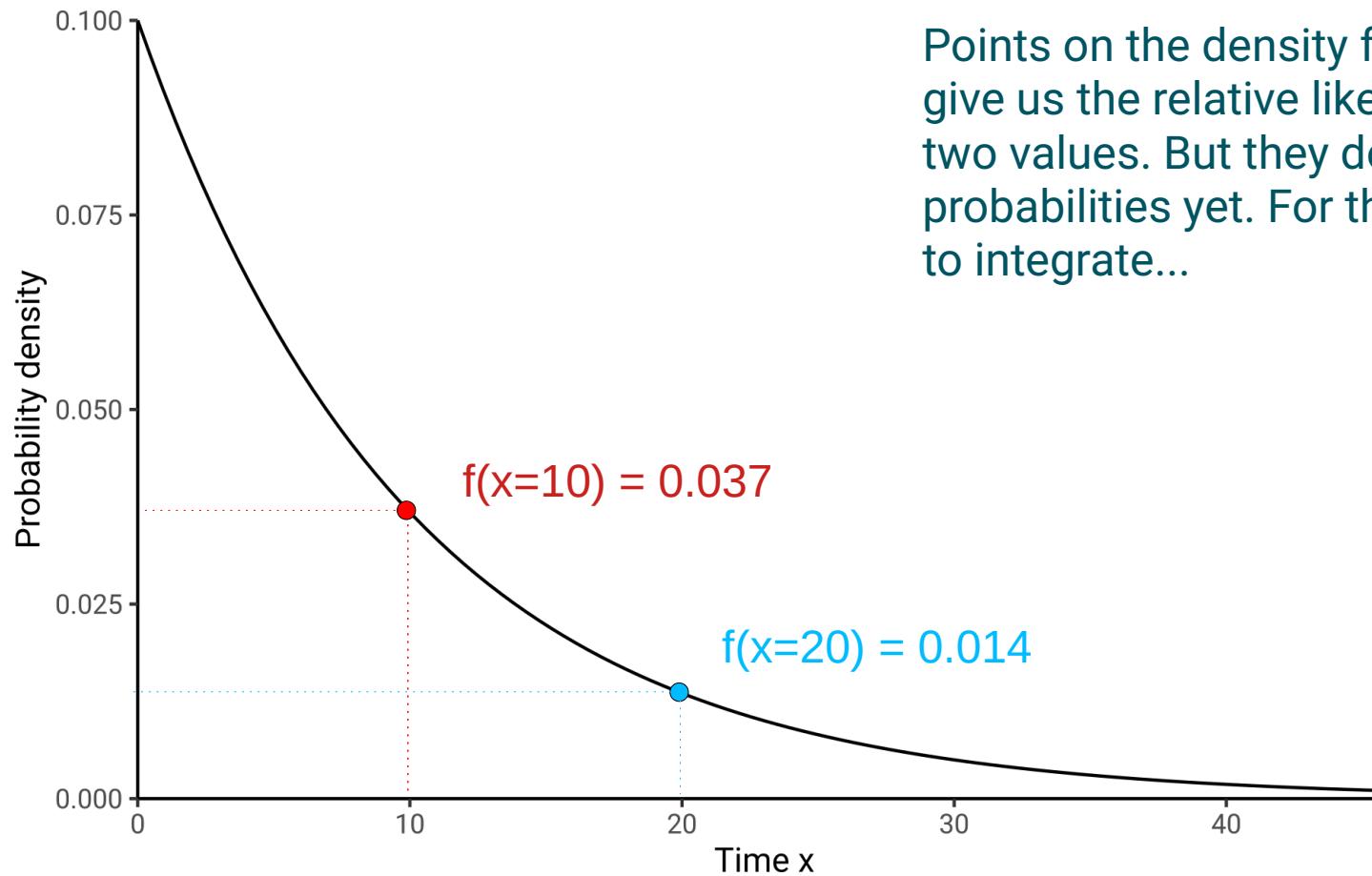


Fig. 2 Model fit on life table deaths for Sweden in 1935 and 2011. The solid line shows the overall mixture model. The dotted line highlights the fit of the Skew Normal employed to estimate accidental and premature mortality. The big dots point out the three modal ages of the distribution

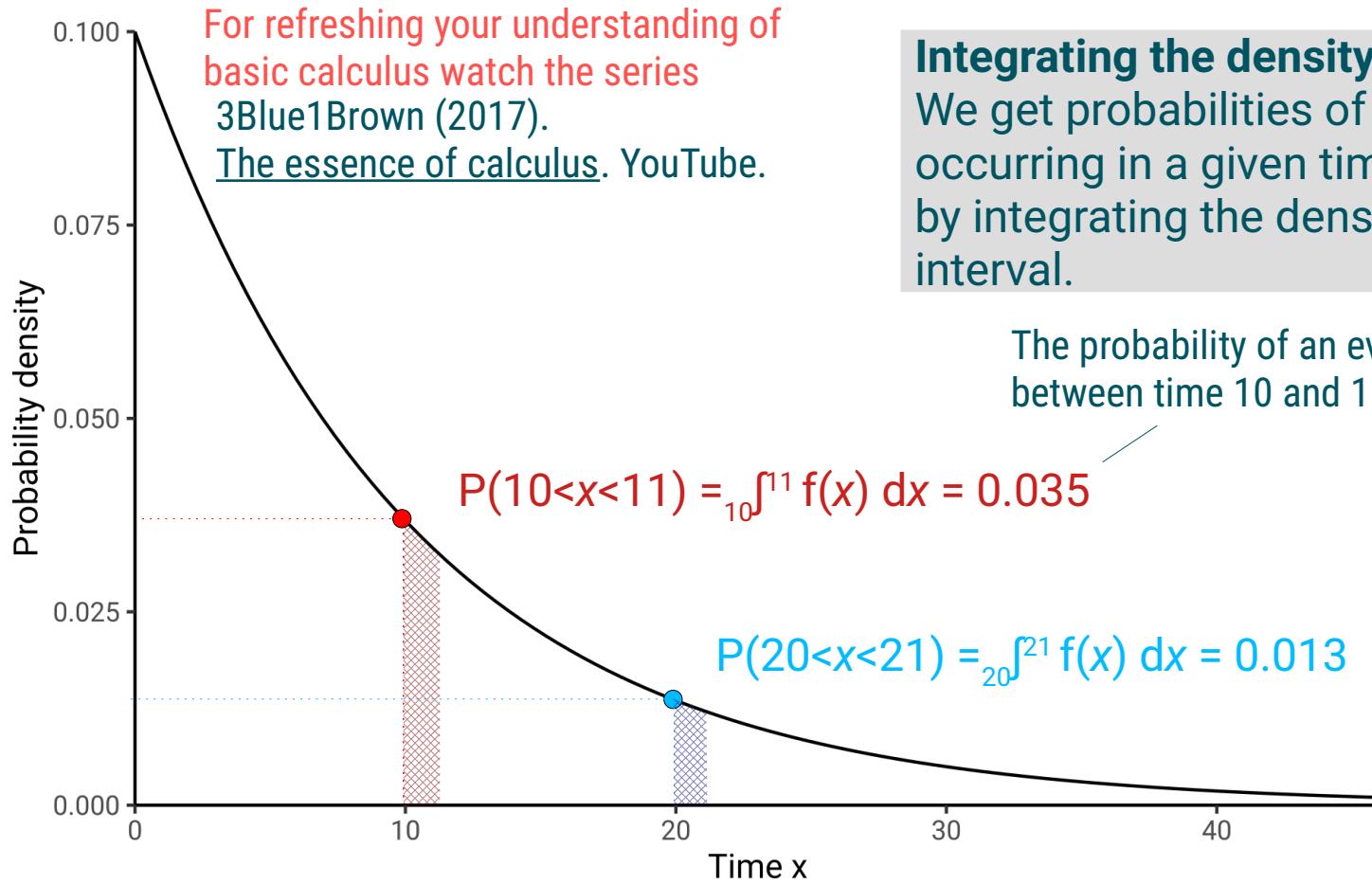
Zanotto et al. (2021). A Mixture-Function Mortality Model.

$f(x)$ Density Function

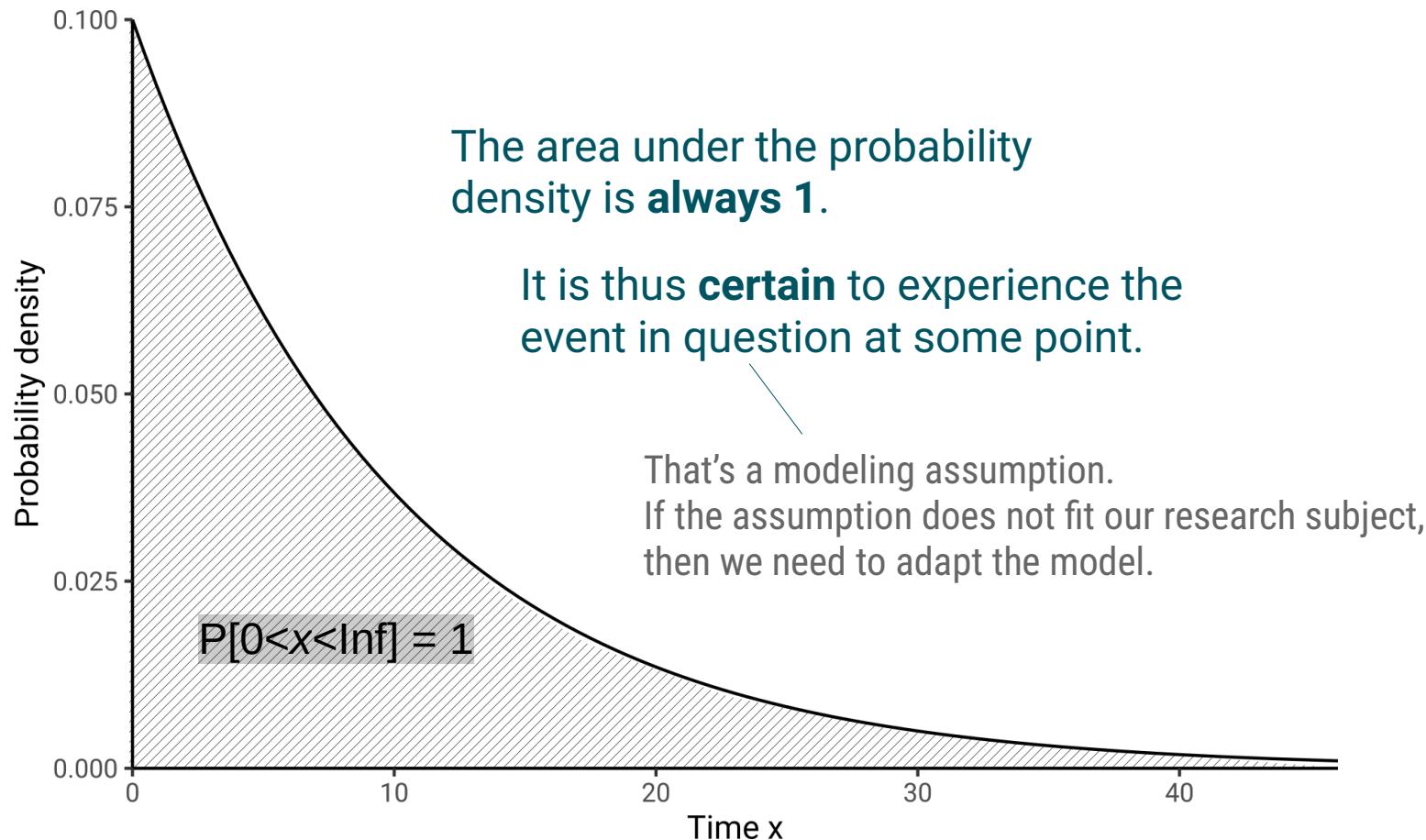


Points on the density function give us the relative likelihood of two values. But they don't give us probabilities yet. For that we need to integrate...

f(x) Density Function



$f(x)$ Density Function

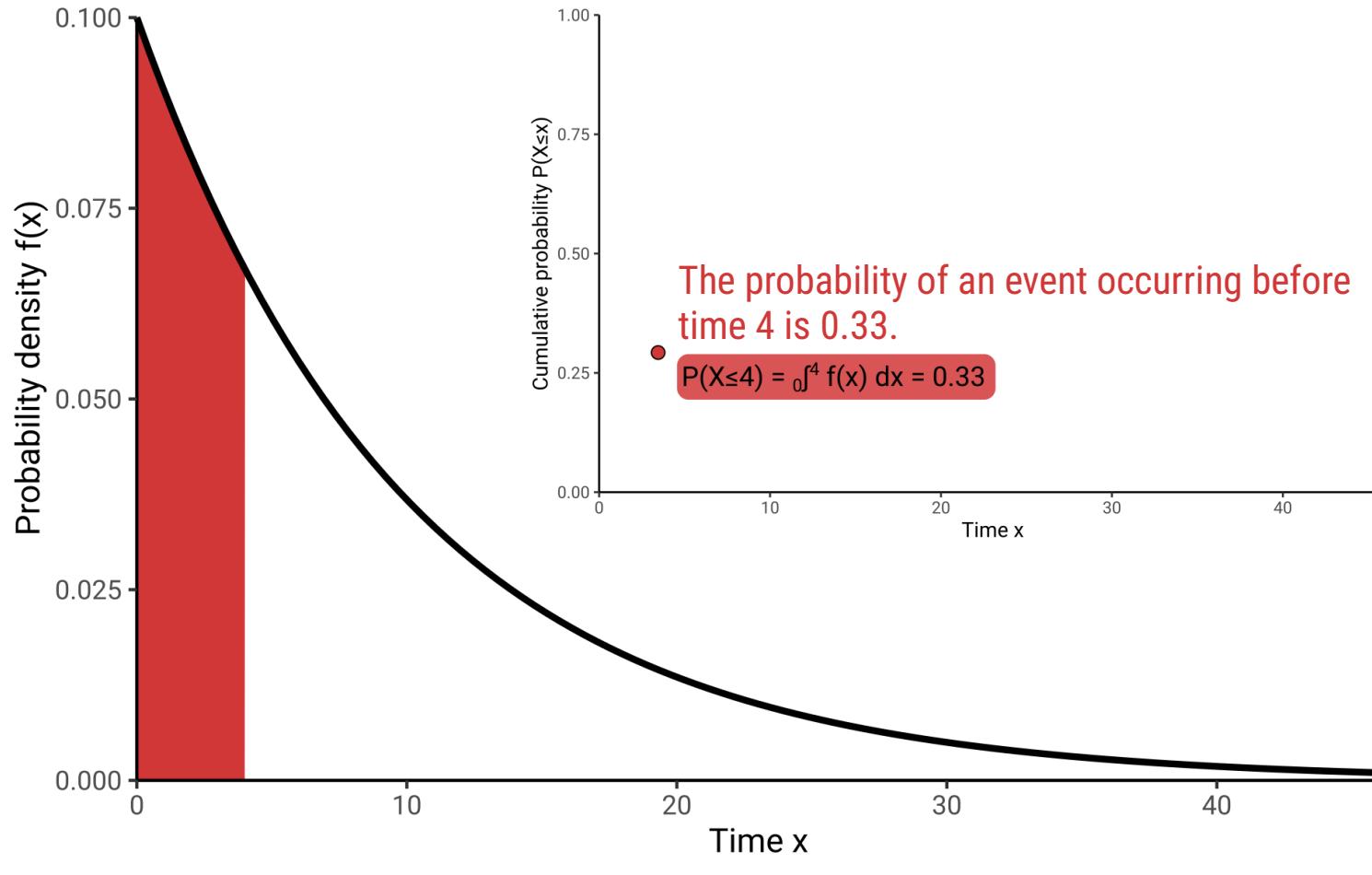


$F(x)$ Distribution Function

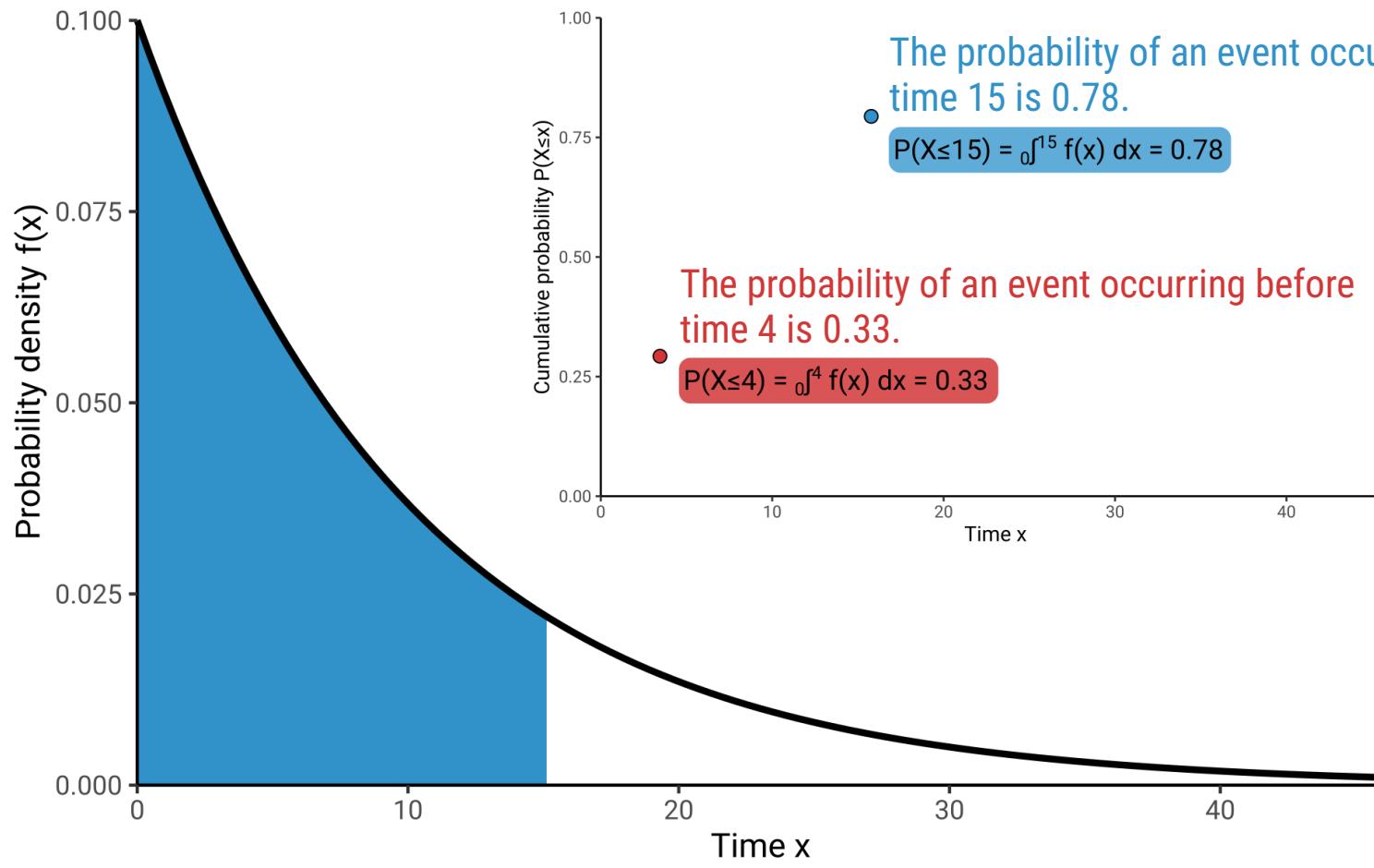
But what about the probability of
an event occurring **before** time x ?

Distribution Function!

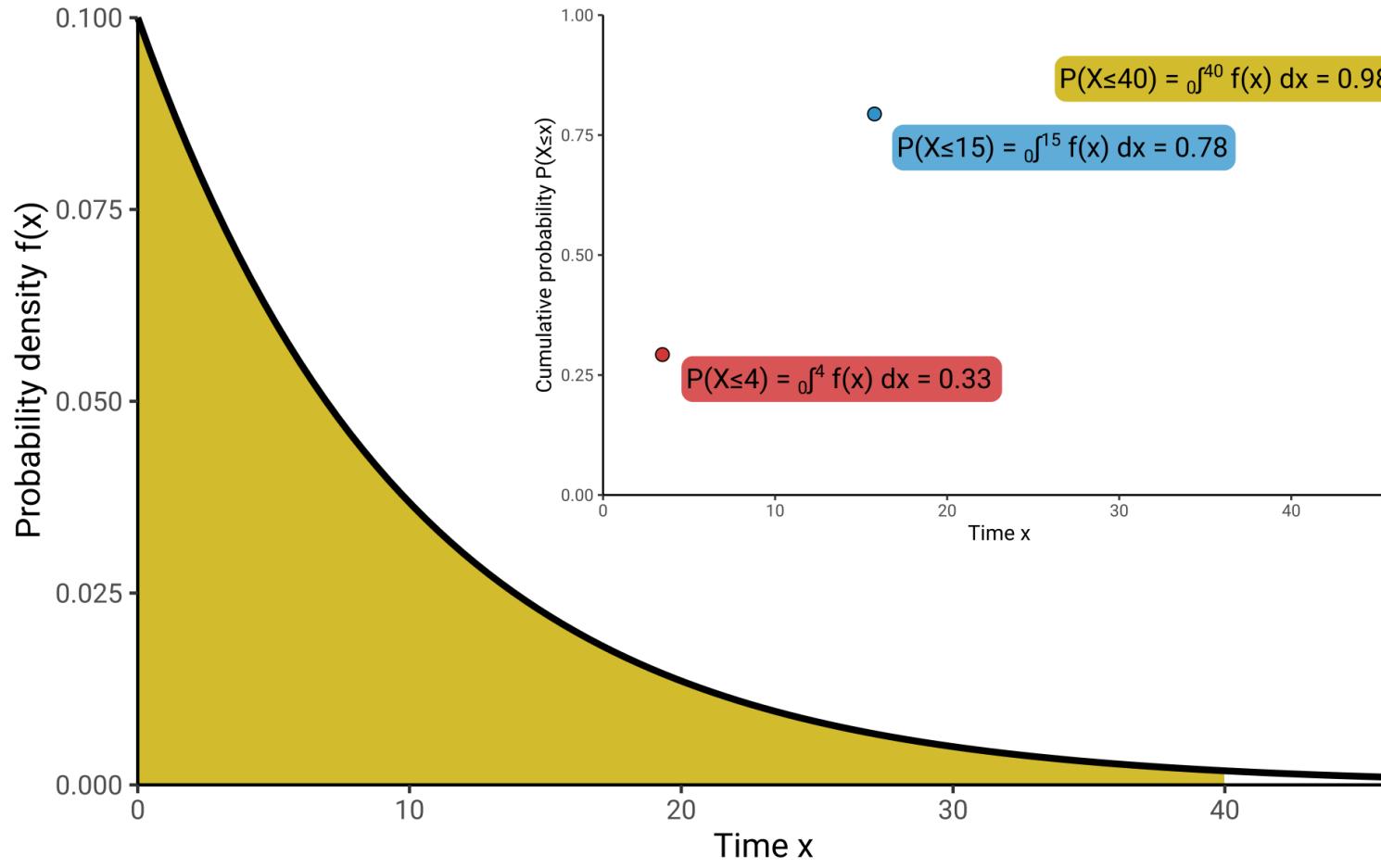
$F(x)$ Distribution Function



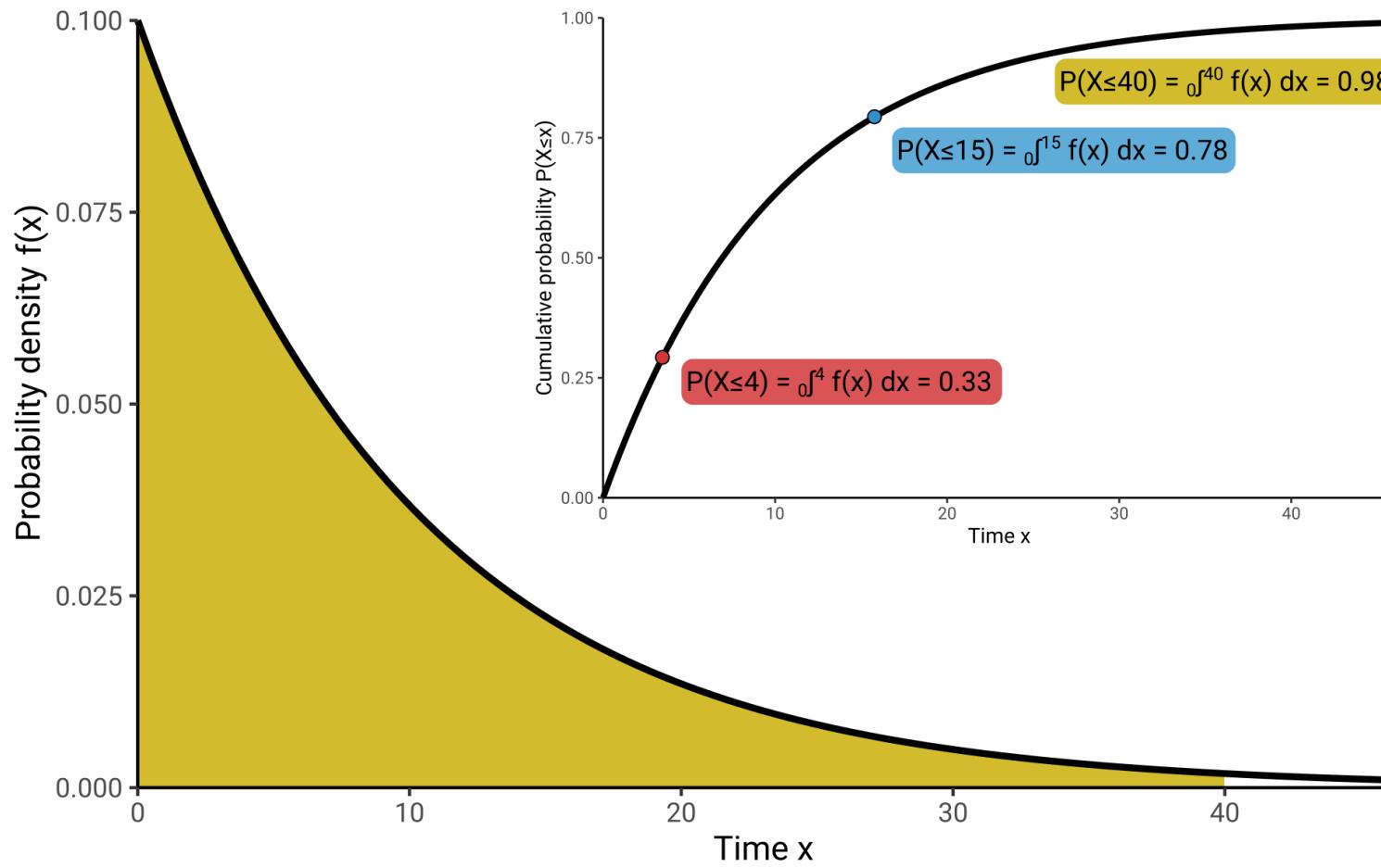
$F(x)$ Distribution Function



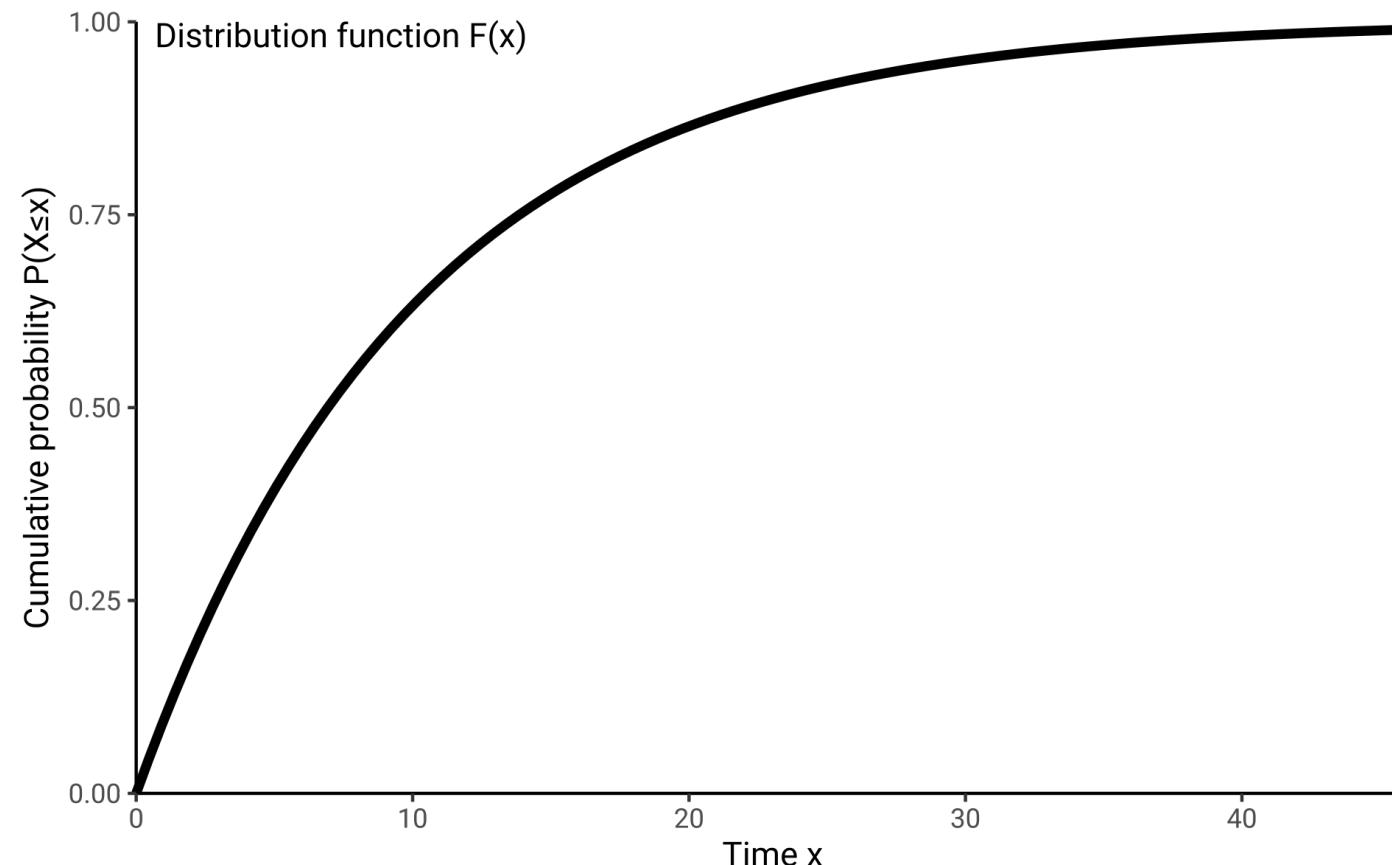
$F(x)$ Distribution Function



$F(x)$ Distribution Function



$F(x)$ Distribution Function

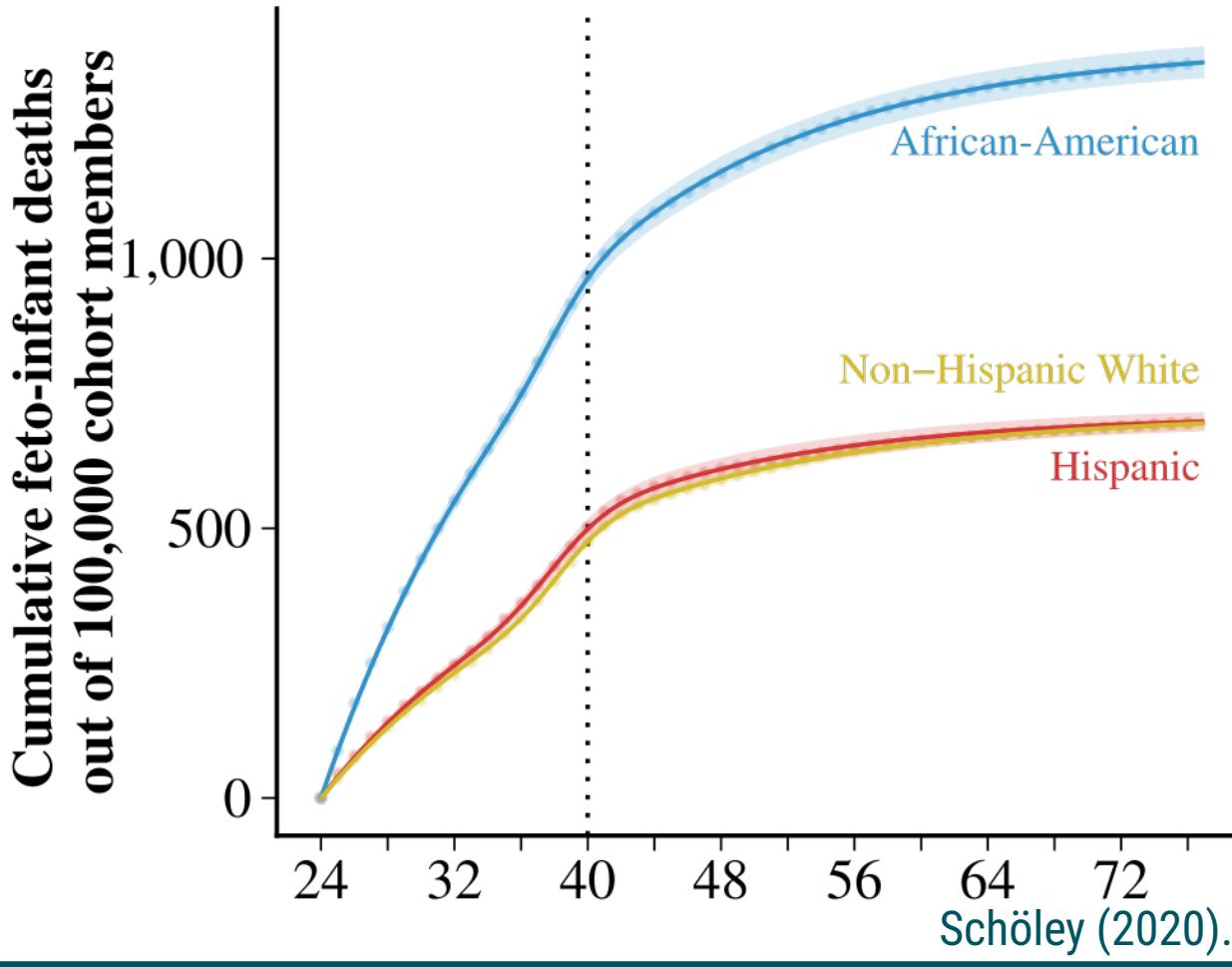


X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

F(x): Distribution function
aka *Cumulative function*
The probability of experiencing the event until time x .

$$F(x) = P(X \leq x) = \int_0^x f(x) dx$$

$F(x)$ Distribution Function



X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

F(x): Distribution function
aka *Cumulative function*
The probability of experiencing the event until time x.

$$F(x) = P(X \leq x) = \int_0^x f(x) dx$$

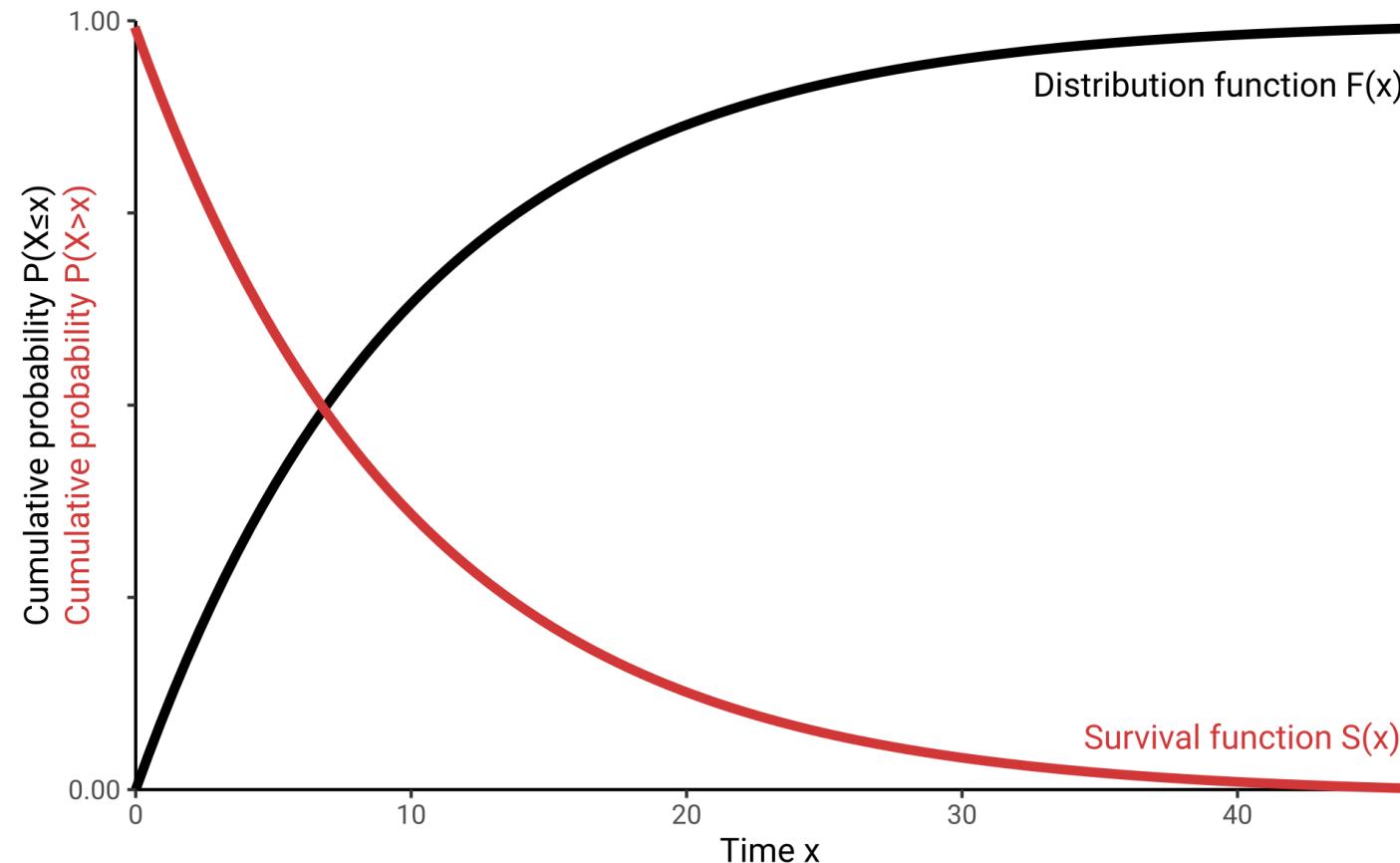
Schöley (2020). The dynamics of ontogenescence.

$S(x)$ Survival Function

But what about the probability of
an event occurring **after** time x ?

Survival Function!

$S(x)$ Survival Function

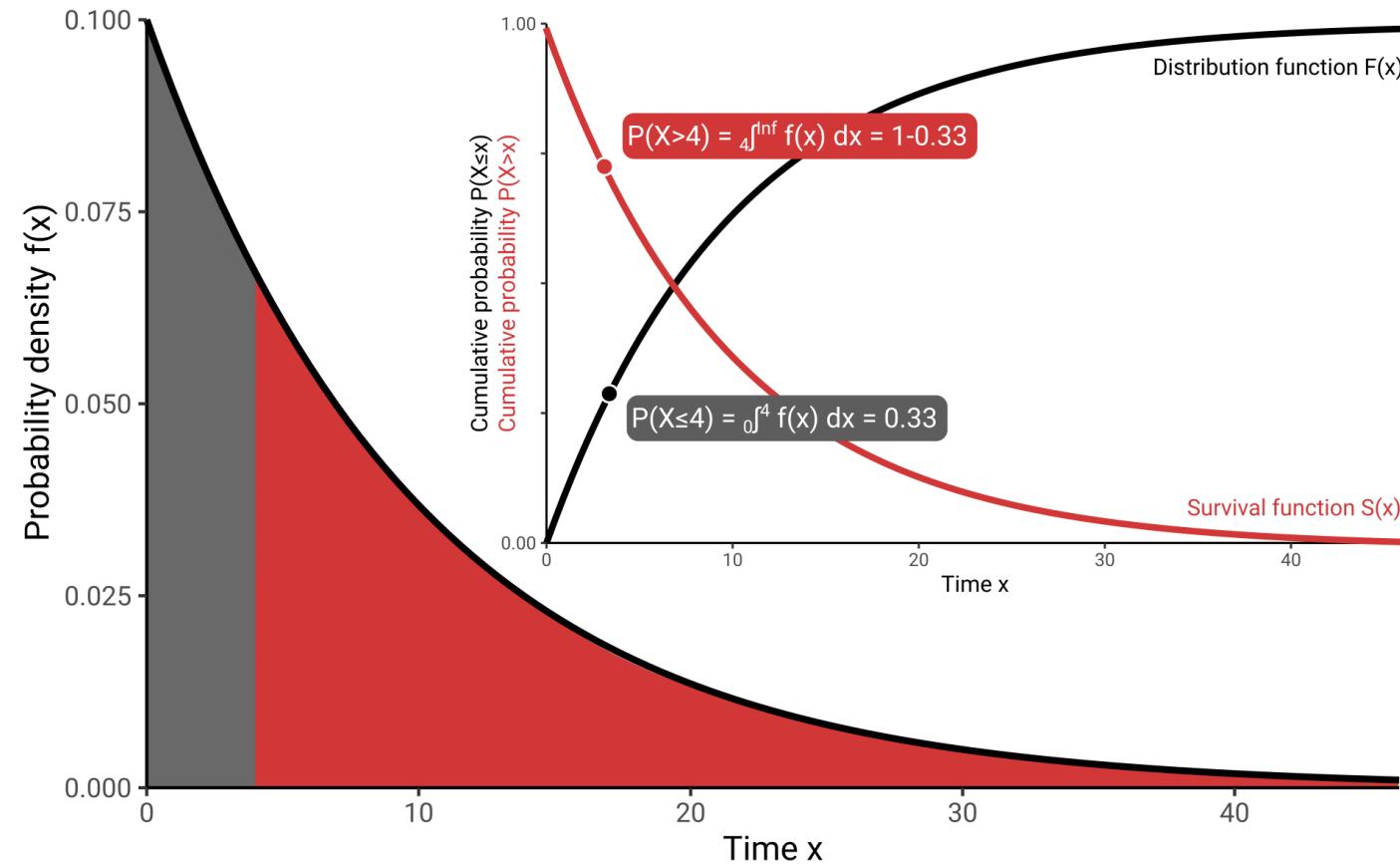


X: Time until event
 $x = 0.1$ weeks, 2.3 weeks...

$S(x)$: Survival function
The probability of *not* experiencing the event until time x .

$$\begin{aligned}S(x) &= P(X > x) = \int_x^{\infty} f(x) dx \\&= 1 - F(x) = 1 - \int_0^x f(x) dx\end{aligned}$$

$S(x)$ Survival Function



X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

$S(x)$: Survival function
The probability of *not* experiencing the event until time x .

$$\begin{aligned} S(x) &= P(X > x) = \int_x^{\infty} f(x) dx \\ &= 1 - F(x) = 1 - \int_0^x f(x) dx \end{aligned}$$

$S(x)$ Survival Function

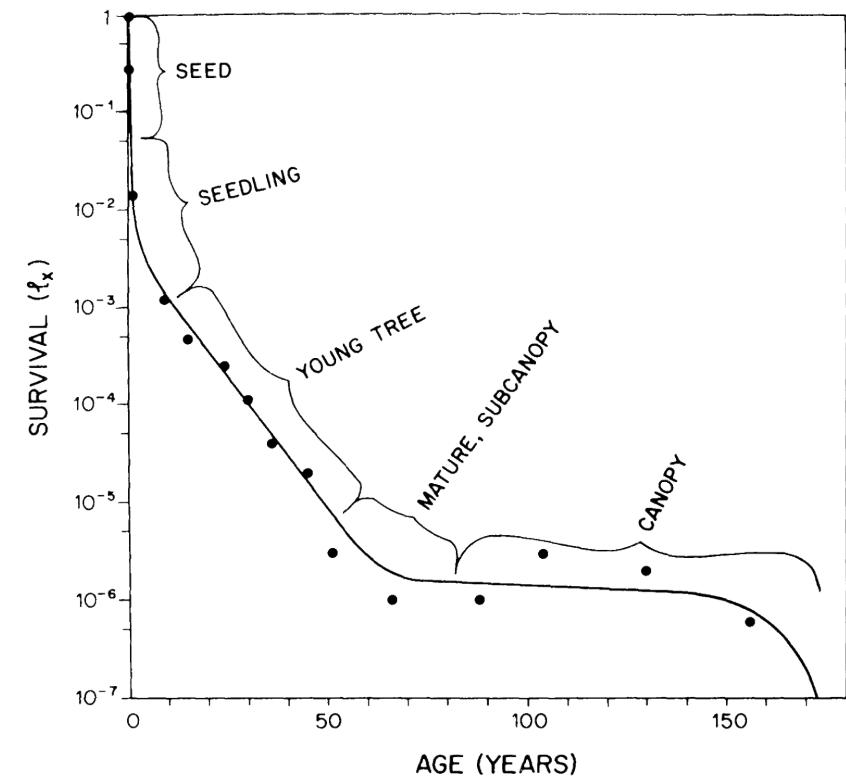


FIGURE 1. Survivorship curve for *Euterpe globosa*, over 7 orders of magnitude. All values of ℓ_x except that at 9 years are completely independent of each other, thus providing an internal check on accuracy.

X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

S(x): Survival function
The probability of *not* experiencing the event until time x.

$$\begin{aligned} S(x) &= P(X>x) = \int_x^{\infty} f(x) dx \\ &= 1 - F(x) = 1 - \int_0^x f(x) dx \end{aligned}$$

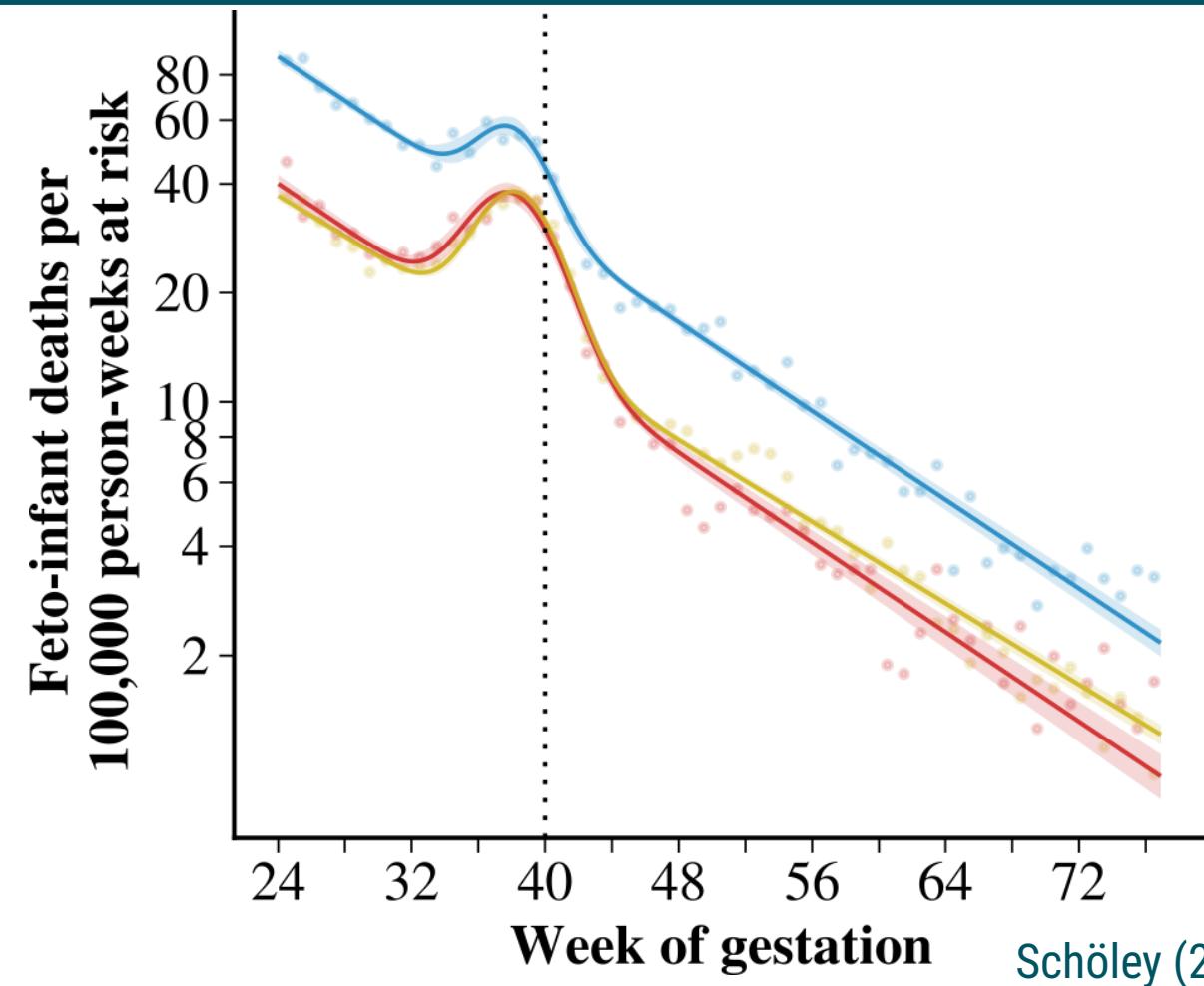
Valen (1975). Life, Death, and Energy of a Tree.

$h(x)$ Hazard Function

But what about the risk of an event occurring around time x , given that it did not occur before?

Hazard Function!

$h(x)$ Hazard Function



X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks} \dots$

h(x): Hazard function
The instantaneous rate of new events at time x among those who did not experience the event yet.
$$h(x) = f(x)/S(x)$$
$$= \lim_{h \rightarrow 0} P(x \leq X < x+h | X \geq x)/h$$

$h(x)$ Hazard Function

We have the *rate* of events

2428 infections per 100,000 persons per 7 days

2428 infections per 100,000 persons-weeks

0.02428 infections per 1 person-week

We want the *survival* probability

What is the probability of me “surviving” the whole semester without catching COVID given current rates?

Thus, we need to express $S(x)$ in terms of $h(x)$

X: Time until event

$x = 0.1$ weeks, 2.3 weeks...

$h(x)$: Hazard function

The instantaneous rate of new events at time x among those who did not experience the event yet.

$$h(x) = f(x)/S(x)$$

$$= \lim_{h \rightarrow 0} P(x \leq X < x+h | X \geq x) / h$$

$h(x)$ Hazard Function

Thus, we need to express $S(x)$ in terms of $h(x)$

$$\begin{aligned} h(x) &= f(x)/S(x) \\ &= F'(x)/S(x) \\ &= [1-S(x)]'/S(x) \\ &= -S'(x)/S(x) \end{aligned}$$

Remember, $F(x) = \int_0^x f(x) dx$.
Thus, $f(x) = d/dx F(x) = F'(x)$.

X: Time until event

$x = 0.1$ weeks, 2.3 weeks...

h(x): Hazard function

The instantaneous rate of new events at time x among those who did not experience the event yet.

$$\begin{aligned} h(x) &= f(x)/S(x) \\ &= \lim_{h \rightarrow 0} P(x \leq X < x+h | X \geq x) / h \end{aligned}$$

$H(x)$ Cumulative Hazard Function

Thus, we need to express $S(x)$ in terms of $h(x)$

$$\begin{aligned} h(x) &= f(x)/S(x) \\ &= F'(x)/S(x) \\ &= [1-S(x)]'/S(x) \\ &= -S'(x)/S(x) \end{aligned}$$

Remember, $F(x) = \int_0^x f(x) dx$.
Thus, $f(x) = d/dx F(x) = F'(x)$.

$$\int_0^x h(x) = \int_0^x -S'(x)/S(x) dx$$

Rule for logarithmic derivatives: $\int g'(x)/g(x) dx = \log |g(x)| + C$
 $= -\log S(x)$

$$-\int_0^x h(x) = \log S(x)$$

$$\exp[-\int_0^x h(x)] = S(x)$$

X: Time until event

$x = 0.1$ weeks, 2.3 weeks...

H(x): Cumulative Hazard
The integral of $h(x)$.

$$H(x) = \int_0^x h(x) dx = -\log S(x)$$

The Exponential Distribution

We have the *rate of events*

2428 infections per 100,000 persons per 7 days

2428 infections per 100,000 persons-weeks

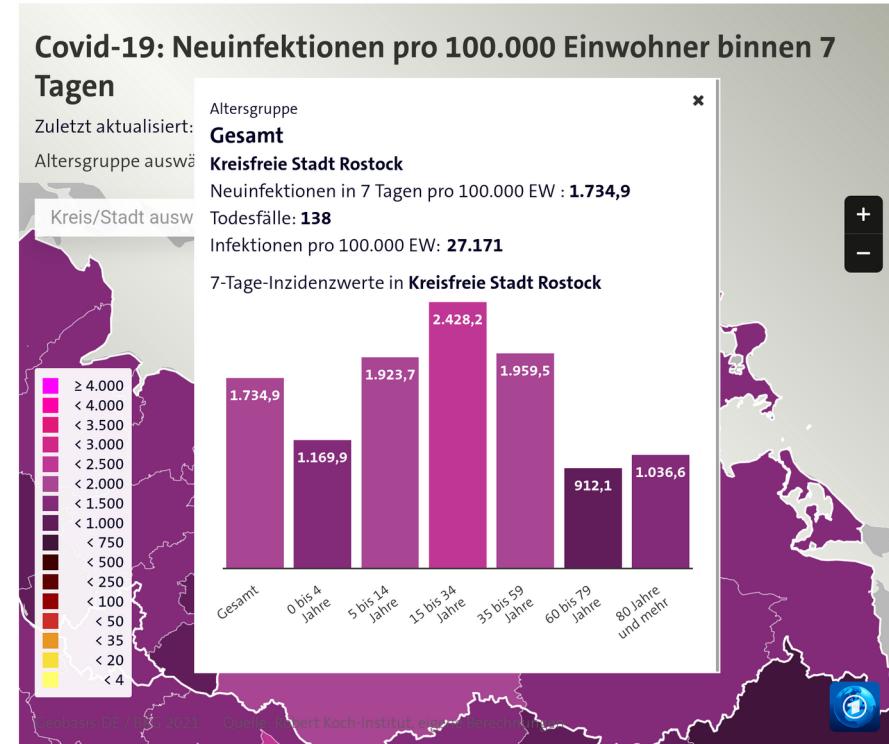
0.02428 infections per 1 person-week

We want the *survival probability*

What is the probability of me “surviving” the whole semester without catching COVID given current rates?

Thus, we need to express $S(x)$ in terms of $h(x)$

$$\exp[-\int_0^x h(x)dx] = S(x)$$



The Exponential Distribution

We have the *rate of events*

2428 infections per 100,000 persons per 7 days

2428 infections per 100,000 persons-weeks

0.02428 infections per 1 person-week

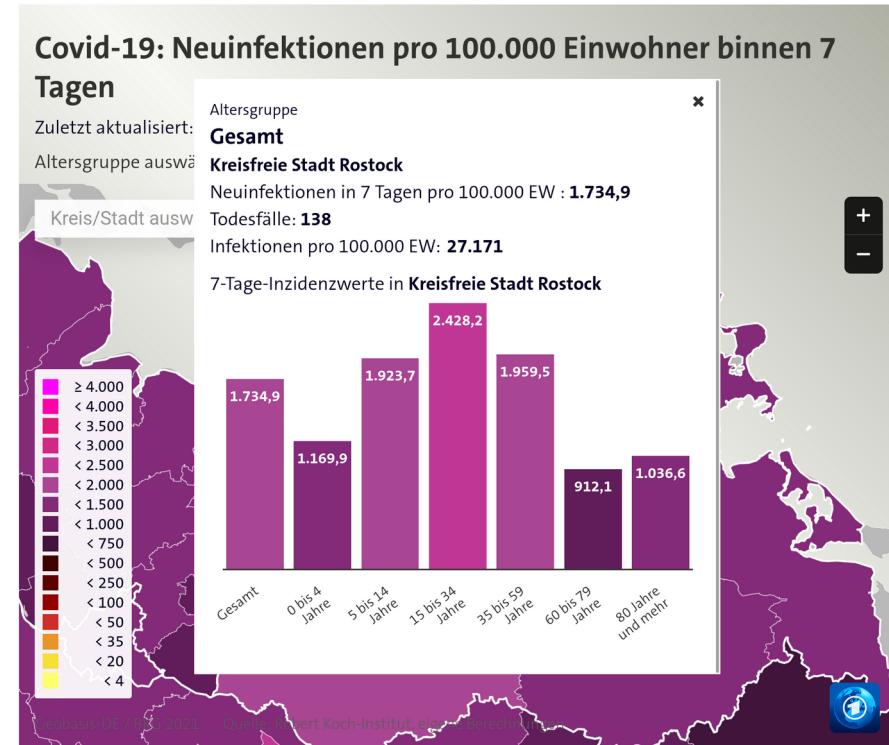
We want the *survival probability*

What is the probability of me “surviving” the whole semester without catching COVID given current rates?

Thus, we need to express $S(x)$ in terms of $h(x)$

$$\exp[-\int_0^x h(x)dx] = S(x)$$

But what shape should $h(x)$ have?



The Exponential Distribution

But what **shape** should $h(x)$ have?

We have the *rate* of events

2428 infections per 100,000 persons per 7 days

2428 infections per 100,000 persons-weeks

0.02428 infections per 1 person-week

The Exponential Distribution

But what **shape** should $h(x)$ have?

We have the *rate* of events

2428 infections per 100,000 persons per 7 days

2428 infections per 100,000 persons-weeks

0.02428 infections per 1 person-week

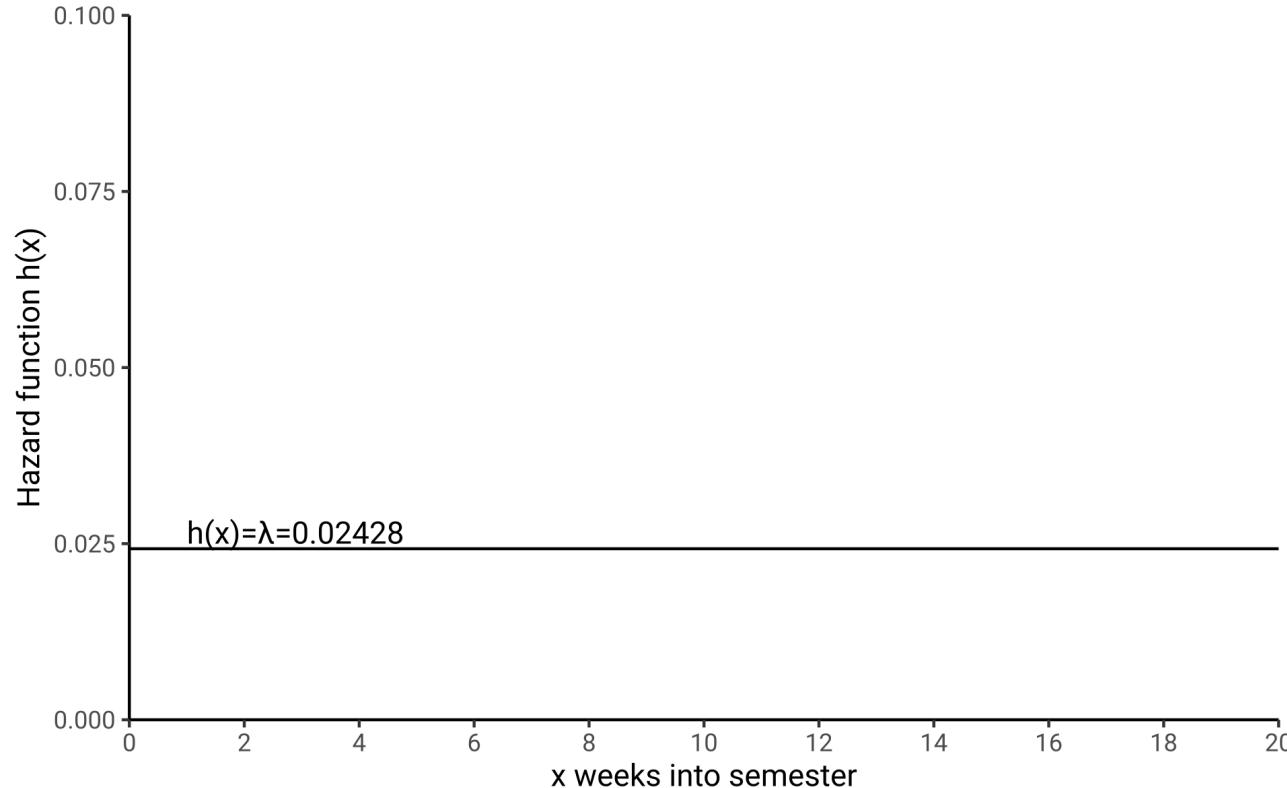
Assuming the rate stays constant over time, **$h(x)$ is a constant!**

$$h(x) = \lambda = 0.02428$$

The Exponential Distribution

Assuming the rate stays constant over time, $h(x)$ is a constant!

$$h(x) = \lambda = 0.02428$$



The Exponential Distribution

Assuming the rate stays constant over time, $h(x)$ is a constant!

$$h(x) = \lambda = 0.02428$$

Applying the survival identity from earlier...

$$\exp[-\int_0^x h(x) dx] = S(x)$$

...yields the survival function of the **Exponential Distribution**

$$\begin{aligned} S(x) &= \exp(-\int_0^x \lambda dx) = \exp(-\lambda x) \\ &= \exp(-0.02428x) \end{aligned}$$

Exponential distribution

If the hazard does not change over time, the time until event is exponentially distributed.

$$h(x) = \lambda$$

$$S(x) = \exp(-\lambda x)$$

The Exponential Distribution

Assuming the rate stays constant over time, $h(x)$ is a constant!

$$h(x) = \lambda = 0.02428$$

Applying the survival identity from earlier...

$$\exp[-\int_0^x h(x) dx] = S(x)$$

...yields the survival function of the **Exponential Distribution**

$$\begin{aligned} S(x) &= \exp(-\int_0^x \lambda dx) = \exp(-\lambda x) \\ &= \exp(-0.02428x) \end{aligned}$$

What is the probability of me “surviving” the whole semester without catching COVID given current rates?

Exponential distribution

If the hazard does not change over time, the time until event is exponentially distributed.

$$h(x) = \lambda$$

$$S(x) = \exp(-\lambda x)$$

Survival Identities

In survival analysis we consider the random variable
"X: Time until event"
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

Survival Identities

In survival analysis we consider the random variable
X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

Survival Identities

In survival analysis we consider the random variable
X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

f(x): Density function
The relative likelihood of experiencing the event around time x.

Survival Identities

In survival analysis we consider the random variable
X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

f(x): Density function
The relative likelihood of experiencing the event around time x.

$$F(x) = \int_0^x f(x) dx$$

F(x): Distribution function
aka Cumulative function
The probability of experiencing the event until time x.
F(x) = P(X ≤ x)

Survival Identities

In survival analysis we consider the random variable
X: Time until event
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

f(x): Density function

The relative likelihood of experiencing the event around time x.

$$F(x) = \int_0^x f(x) dx$$

$$S(x) = \int_x^\infty f(x) dx$$

F(x): Distribution function

aka *Cumulative function*

The probability of experiencing the event until time x.

$$F(x) = P(X \leq x)$$

$$S(x) = 1 - F(x)$$

S(x): Survival function

The probability of *not* experiencing the event until time x.

$$S(x) = P(X > x)$$

Survival Identities

In survival analysis we consider the random variable
"X: Time until event"
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

f(x): Density function

The relative likelihood of experiencing the event around time x.

$$F(x) = \int_0^x f(x) dx$$

F(x): Distribution function

aka *Cumulative function*

The probability of experiencing the event until time x.

$$F(x) = P(X \leq x)$$

S(x): Survival function

The probability of *not* experiencing the event until time x.

$$S(x) = P(X > x)$$

h(x): Hazard function

The instantaneous rate of new events at time x among those who did not experience the event yet.

$$h(x) = \lim_{h \rightarrow 0} \frac{P(x \leq X < x+h | X \geq x)}{h}$$

$$S(x) = 1 - F(x)$$

$$S(x) = \int_x^{\infty} f(x) dx \quad h(x) = -S'(x)/S(x)$$

Survival Identities

In survival analysis we consider the random variable
"X: Time until event"
 $x = 0.1 \text{ weeks}, 2.3 \text{ weeks}...$

We express our knowledge about the distribution of X in any of these functions. Knowing any single function we can derive all other via the **Survival Identities**.

f(x): Density function

The relative likelihood of experiencing the event around time x.

$$F(x) = \int_0^x f(x) dx$$

F(x): Distribution function

aka *Cumulative function*

The probability of experiencing the event until time x.

$$F(x) = P(X \leq x)$$

S(x): Survival function

The probability of *not* experiencing the event until time x.

$$S(x) = P(X > x)$$

$$S(x) = \int_x^\infty f(x) dx \quad h(x) = -S'(x)/S(x)$$

$$S(x) = 1 - F(x)$$

h(x): Hazard function

The instantaneous rate of new events at time x among those who did not experience the event yet.

$$h(x) = \lim_{h \rightarrow 0} \frac{P(x \leq X < x+h | X \geq x)}{h}$$

$$H(x) = \int_0^x h(x) dx$$

$$H(x) = -\log S(x)$$

H(x): Cumulative Hazard
The integral of h(x).

Survival Identities

There are many more identities...

Table 2/1: Relations among the six functions describing stochastic lifetime

to from \ \diagdown	$f(x)$	$F(x)$	$R(x)$	$h(x)$	$H(x)$	$\mu(x)$
$f(x)$	—	$\int_0^x f(z) dz$	$\int_x^\infty f(z) dz$	$\frac{f(x)}{\int_x^\infty f(z) dz}$	$-\ln\left\{\int_x^\infty f(z) dz\right\}$	$\frac{\int_0^x z f(x+z) dz}{\int_x^\infty f(z) dz}$
$F(x)$	$F'(x)$	—	$1 - F(x)$	$\frac{F'(x)}{1 - F(x)}$	$-\ln\{1 - F(x)\}$	$\frac{\int_x^\infty [1 - F(z)] dz}{1 - F(x)}$
$R(x)$	$-R'(x)$	$1 - R(x)$	—	$\frac{-R'(x)}{R(x)}$	$-\ln[R(x)]$	$\frac{\int_x^\infty R(z) dz}{R(x)}$
$h(x)$	$h(x) \exp\left\{-\int_0^x h(z) dz\right\}$	$1 - \exp\left\{-\int_0^x h(z) dz\right\}$	$\exp\left\{-\int_0^x h(z) dz\right\}$	—	$\int_0^x h(z) dz$ $\frac{\int_x^\infty \exp\left\{-\int_0^z h(v) dv\right\} dz}{\exp\left\{-\int_0^x h(z) dz\right\}}$	
$H(x)$	$-\frac{d\{\exp[-H(x)]\}}{dx}$	$1 - \exp\{-H(x)\}$	$\exp\{-H(x)\}$	$H'(x)$	—	$\frac{\int_x^\infty \exp\{-H(z)\} dz}{\exp\{-H(x)\}}$
$\mu(x)$	$\frac{1 + \mu'(x)}{\mu^2(x)} \times \mu(0) \times$ $\times \exp\left\{-\int_0^x \frac{1}{\mu(z)} dz\right\}$	$1 - \frac{\mu(0)}{\mu(x)} \times$ $\times \exp\left\{-\int_0^x \frac{1}{\mu(z)} dz\right\}$	$\frac{\mu(0)}{\mu(x)} \times$ $\times \exp\left\{-\int_0^x \frac{1}{\mu(z)} dz\right\}$	$\frac{1}{\mu(x)} \{1 + \mu'(x)\}$	$\ln\left\{\frac{\mu(x)}{\mu(0)}\right\} +$ $+ \int_0^x \frac{1}{\mu(z)} dz$	—

Rinne (2008). The Weibull Distribution.

Recap

For survival distributions and identities read

Klein & Moeschberger (2003). Survival Analysis. Sections 2.1–2.4.

For refreshing your understanding of basic calculus watch the series

3Blue1Brown (2017). The essence of calculus. YouTube.

For refreshing your understanding of random variables and probability distributions watch

Khan Academy (2012). Random variables. YouTube.

Princeton COS 302 (2020). Probability density and mass functions. YouTube.

Homework

Choose a time-to-event setting that interests you and look up a constant rate related to that setting. What is the time scale for your setting? When does the time-to-event start? When have half of the population experienced the event given the chosen rate?

Example: Today we looked at the time until I catch COVID. I choose the rate 2,428 infections per 100,000 persons per 7 days from the local COVID incidences and assumed this rate to be constant. The timescale was “weeks into the semester” and it starts at the first week of the semester. I used the survival function of the exponential distribution to calculate the time until the probability of catching COVID reached 50%.
 $S(x) = \exp(-\int_0^x \lambda dx) = \exp(-\lambda x)$

Materials for this lecture

github.com/jschoeley/survival_analysis-ur-ss22

Jonas Schöley

 @jschoeley

 0000-0002-3340-8518

 j.schoeley@uni-rostock.de

CC-BY Jonas Schöley 2022