The Distributions Package
for SBML Level 3

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Version 0.22 (Draft)

Feburary, 2019

Contents

Revision History

Version	Date	Author	Comments
0.1 (Draft)	15 Oct 2011	Stuart Moodie	First draft
0.2 (Draft)	16 Oct 2011	Stuart Moodie	Added introductory text and background info. Other minor changes etc.
0.3 (Draft)	16 Oct 2011	Stuart Moodie	Filled empty invocation semantics section.
0.4 (Draft)	4 Jan 2012	Stuart Moodie	Incorporated comments from NlN, MS and SK. Some minor revisions and corrections.
0.5 (Draft)	6 Jan 2012	Stuart Moodie	Incorporated addition comments on aim of package from NlN.
0.6 (Draft)	19 Jul 2012	Stuart Moodie	Incorporated revisions discussed and agreed at HARMONY 2012.
0.7 (Draft)	6 Aug 2012	Stuart Moodie	Incorporated review comments from Maciej Swat and Sarah Keating.
0.8 (Draft)	21 Dec 2012	Stuart Moodie	Incorporated changes suggested at combine and subsequently through list discussions.
0.9 (Draft)	9 Jan 2013	Stuart Moodie	Incorporated corrections and comments from Maciej Swat and Sarah Keating.
0.10 (Draft)	10 Jan 2013	Stuart Moodie	Modified based on comments from MS.
0.11 (Draft)	17 May 2013	Lucian Smith	Modified based on Stuart's proposals and PWG discussion.
0.12 (Draft)	June 2013	Lucian Smith and Stuart Moodie	Modified based on HARMONY 2013 discussion.
0.13 (Draft)	July 2013	Lucian Smith and Stuart Moodie	Modified based PWG discussion, particularly with respect to UncertML.
0.14 (Draft)	March 2015	Lucian Smith	Modified to match UncertML 3.0.
0.15 (Draft)	March 2015	Lucian Smith and Sarah Keating	Modified to match UncertML 3.0 for real this time.
0.16 (Draft)	March 2015	Lucian Smith	Added information about UncertML 3.0 distributions, and the distributions custom annotations.
0.17 (Draft)	June 2017	Lucian Smith	Extensive update to reflect demise of UncertML 3.0, and appearance of ProbOnto.
0.18 (Draft)	June 2017	Lucian Smith	Fixes to reflect feedback on version 0.17.
0.19 (Draft)	June 2018	Lucian Smith	Resolved id/name issues with SBML core l3v1 vs. l3v2.
0.20 (Draft)	December 2018	Lucian Smith	Updates to allow distributions as new MathML csymbols.
0.21 (Draft)	January 2018	Lucian Smith	Revisions based on suggestions from sbml-distrib, including extensive edits from Matthias. Also removed the extended function definitions entirely.
0.22 (Draft)	February 2018	Lucian Smith	Addition of `sampleSize`, mean values of distributions for fallback.

1 Introduction and motivation

1.1 What is it?

The Distributions package (also known as *distrib*) provides an extension to SBML Level 3 that enables a model to encode and sample from both discrete and continuous probability distributions, and provides the ability to annotate elements with information about the distribution their values were drawn from. Applications of the package include for instance descriptions of population based models, an important subset of which are pharmacokinetic/pharma-codynamic (PK/PD) models[1], which are used to model the action of drugs.

1.2 Scope

The Distributions package adds support to SBML for sampling from a probability distribution. In particular the following are in scope:

- Sampling from a continuous distribution.

- Sampling from a discrete distribution.

- Sampling from a user-defined discrete probability density function.

- Specification of descriptive statistics (mean, standard deviation, standard error, etc.).

At one point the following were considered for inclusion in this package but are now **out of scope**:

- Definitions of ranges (the original name of the package was 'Distributions and Ranges').

- Sampling from user-defined probability density function.

- Stochastic differential equations.

- Other functions used to characterise a probability distribution, such as cumulative distribution functions (CDF) or survival functions, etc.

1.3 This Document

This proposal describes the consensus view of workshop participants and subscribers to the sbml-distrib mailing list. Although it was written by the listed authors it does not soley reflect their views nor is it their proposal. Rather, it is their understanding of the consensus view of what the Distributions package should do and how it should do it. The contributors listed have made significant contributions to the development and writing of this specification and are credited accordingly, but a more comprehensive attribution is provided in the acknowledgements (Section 7 on page 38).

Finally, the authors would encourage the reader to consider them and contribute their ideas or comments — indeed any feedback about this proposal — to the *distrib* discussion list[2].

Once the proposal is finalised this will be the first step towards the formal adoption of *distrib* as a package in SBML Level 3. After this, two implementations based on this proposal are required and then the SBML editors must agree that the implementations and specification are complete. The proposal will then provide the basis for a future package specification document. More details of the SBML package adoption process can be found at: http://sbml.org/Documents/SBML_Development_Process.

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[1] for more information see: http://www.pharmpk.com/.
[2] sbml-distrib@lists.sourceforge.net

1.4 Conventions used in this document

There are some parts of this proposal where there is no clear consensus on the correct solution or only recent agreement or agreement by a group which may not be representative of the SBML community as a whole. These cases are indicated by the question mark in the left margin (illustrated). The reader should pay particular attention to these points and ideally provide feedback, especially if they disagree with what is proposed. Similarly there will be points — especially as the proposal is consolidated — which are agreed, but which the reader should take note of and perhaps read again. These points are emphasised by the hand pointer in the left margin (illustrated).

2 Background

2.1 Problems with current SBML approaches

SBML Level 3 Core has no direct support for encoding values sampled from distributions within a model. Currently there is no workaround within the core language itself, although it is possible to define such information using annotations. Frank Bergmann had proposed such an semi-formalised extension for use with SBML L2 and L3 (See Section 4.1 on page 29).

2.2 Past work on this problem or similar topics

2.2.1 The Newcastle Proposal

In 2005 there was a proposal from Colin Gillespie and others [3] to introduce support for probability distributions in the SBML core specification. This was based on their need to use such distributions to represent the models they were creating as part of the BASIS project (http://www.basis.ncl.ac.uk).

They proposed that distributions could be referred to in SBML using the **csymbol** element in the MathML subset used by the SBML Core specification. An example is below:

```
<math xmlns=''http://www.w3.org/1998/Math/MathML''>
  <apply>
    <csymbol encoding=''text''
        definitionURL=''http://www.sbml.org/sbml/symbols/uniformRandom''>
      uniformRandom
    </csymbol>
    <ci>mu</ci>
    <ci>sigma</ci>
  </apply>
</math>
```

This required that a library of definitions be maintained as part of the SBML standard and in their proposal they defined an initial small set of commonly used distributions. The proposal was never implemented.

2.2.2 Seattle 2010

The "distrib" package was discussed at the Seattle SBML Hackathon[4] and this section is an almost verbatim reproduction of Darren Wilkinson's report on the meeting[5]. There Darren presented an overview of the problem[67], building on the old proposal from the Newcastle group (see above: Section 2.2.1). There was broad support at the meeting for development of such a package, and for the proposed feature set. Discussion following the presentation led to a consensus on the following points:

- There is an urgent need for such a package.

- It is important to make a distinction between a description of uncertainty regarding a model parameter and the mechanistic process of selecting a random number from a probability distribution, for applications such as parameter scans and experimental design

- It is probably worth including the definition of PMFs, PDFs and CDFs in the package

- It is worth including the definition of random distributions using particle representations within such a package, though some work still needs to be done on the precise representation

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[3] http://sbml.org/Community/Wiki/SBML_Leve\T1\l_3_Proposals/Distributions_and_Ranges
[4] http://sbml.org/Events/Hackathons/The_2010_SBML-BioModels.net_Hackathon
[5] http://sbml.org/Forums/index.php?t=tree&goto=6141&rid=0
[6] Slides: http://sbml.org/images/3/3b/Djw-sbml-hackathon-2010-05-04.pdf
[7] Audio: http://sbml.org/images/6/67/Wilkinson-distributions-2010-05-04.mov

- It could be worth exploring the use of xinclude to point at particle representations held in a separate file

- Random numbers must not be used in rate laws or anywhere else that is continuously evaluated, as then simulation behaviour is not defined

- Although there is a need for a package for describing extrinsic noise via stochastic differential equations in SBML, such mechanisms should not be included in this package due to the considerable implications for simulator developers

- We probably don't want to layer on top of UncertML (www.uncertml.org), as this spec is fairly heavy-weight, and somewhat tangential to our requirements

- A random number seed is not part of a model and should not be included in the package

- The definition of truncated distributions and the specification of hard upper and lower bounds on random quantities should be considered.

It was suggested that new constructs should be introduced into SBML by the package embedded as user-defined functions using the following syntax:

```
<listOfFunctionDefinitions>
  <functionDefinition id="myNormRand">
    <distrib:####>
      #### distrib binding information here ####
    </distrib:####>
    <math>
      <lambda>
        <bvar>
          <ci>mu</ci>
          <ci>sigma</ci>
        </bvar>
        <ci>mu</ci>
      </lambda>
    </math>
  </functionDefinition>
</listOfFunctionDefinitions>
```

which allows the use of a "default value" by simulators which do not understand the package (but simulators which do will ignore the <math> element). The package would nevertheless be "required", as it will not be simulated correctly by software which does not understand the package.

Informal discussions following the break-out covered topics such as:

- how to work with vector random quantities in the absence of the vector element in the MathML subset used by SBML

- how care must be taken with the semantics of random variables and the need to both:
  - reference multiple independent random quantities at a given time
  - make multiple references to the same random quantity at a given time.

2.2.3 Hinxton 2011

Detailed discussion was continued at the Statistical Models Workshop in Hinxton in June 2011[8]. There those interested in representing Statistical Models in SBML came together to work out the details of how this package would work in detail. Dan Cornford from the UncertML project[9] attended the meeting and described how that resource could be used to describe uncertainty and in particular probability distributions. Perhaps the most

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[8]http://sbml.org/Events/Other_Events/statistical_models_workshop_2011
[9]http://www.uncertml.org/

significant decision at this meeting was to adopt the UncertML resource as a controlled vocabulary that is referenced by the Distributions package.

Much has changed since this meeting, but the output from this meeting was the basis for the first version of this proposal.

2.2.4 HARMONY 2012: Maastricht

Two sessions were dedicated to discussion of Distributions at HARMONY based around the proposals described in version 0.5 of this document. In addition there was discussion about the Arrays proposal which was very helpful in solving the problem of multivariate distributions in Distributions. The following were the agreed outcomes of the meeting:

- The original proposal included UncertML markup directly in the function definition. This proved unwieldy and confusing and has been replaced by a more elegant solution that eliminates the UncertML markup and integrates well with the fallback function (see details below).

- Multivariate distributions can be supported using the Arrays package to define a covariance matrix.

- User defined continuous distributions would define a PDF in MathML.

- Usage semantics were clarified so that invocation of a function definition implied a value was sampled from the specified distribution.

- It was agreed from which sections of an SBML model a distribution could be invoked.

- Statistical descriptors of variables (for example mean and standard deviation) would be separated from Distributions and either provided in a new package or in a later version of SBML L3 core.

2.2.5 COMBINE 2012: Toronto

The August proposal was reviewed and an improvement was agreed to the user-defined PMF part of the proposal. In particular is was agreed that the categories should be defined by *distrib* classes rather than by passing in the information as an array. Questions were also raised about whether UncertML was suitably well defined to be used as an external definition for probability distributions. This was resolved subsequent to the meeting with a teleconference to Dan Cornford and colleagues. These changes are incorporated here. Finally, there was considerable debate about whether to keep the dependence of *distrib* on the Arrays package in order to support multi-variate distributions. The outcome was an agreement that we would review this at the end of 2012, based on the results of an investigation into how feasible it would be to implement Arrays as a package.

2.2.6 2013 Package Working Group discussions

Early 2013 saw a good amount of discussion on the *distrib* Package Working Group mailing list, spurred by proposals by Stuart Moodie[10]. While not all of his suggestions ended up being fully accepted by the group, several changes were accepted, including:

- To use UncertML as actual XML, instead of as a set of reference definitions.

- To use UncertML to encode descriptive statistics of SBML elements such as mean, standard deviation, standard error, etc.) bringing this capability back in scope for this package.

2.2.7 HARMONY 2013: Connecticut

At HARMONY at UConn in Connecticut, further discussions revealed the importance of distinguishing the ability to describe an element as a distributed variable vs. a function call within the model performing a draw from a distribution.

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[10]http://thestupott.wordpress.com/2013/03/12/an-improved-distrib-proposal/

We also decided to discard the encoding of explicit PDFs for now, as support for it is remarkably complicated, and there no demand for it. The current design could be extended to support this feature so if there is demand for it in the future support for explicit PDFs could be reintroduced.

2.2.8 Early 2017 and HARMONY: Seattle

In early 2017, it became clear that UncertML was no longer being worked on; the web page had lapsed, and its authors had moved on to other things. At the same time, the ProbOnto ontology (Swat et al. 2016; http://probonto.org/) was developed that included all the distributions from UncertML as well as a huge number of other distributions. On the mailing list, it was discussed whether to create essentially our own version of UncertML, or to implement a generic 'reference' format that used ProbOnto. The v0.17 draft specification was developed as a compromise 'hybrid' system that did parts of both, so that basic distributions would be hard-coded, but the ability to reference any ProbOnto ontology would also be present. The hope is that with working examples of both approaches, either the hybrid approach will be approved, or if one is preferred, the other approach may be removed. This version of the specification was created for presentation at HARMONY 2017 in Seattle.

2.2.9 Early 2018 and HARMONY: Oxford

At HARMONY in Oxford, for the first time since the change from UncertML, a libsbml implementation of the specification was available. This let people experiment with the package, and conclude that a simpler method of defining calls to distributions was desired. It was proposed to define new MathML 'csymbol' definitions for the common distributions. It was unclear whether the other methods for everything would be useful or not; for now, we are leaving them in, and if people develop support for them, they can stay.

3 Proposed syntax and semantics

3.1 Overview

Following the precedent set by the SBML Level 3 Core specification document, we use UML 1.0 (Unified Modeling Language; Eriksson and Penker 1998; Oestereich 1999) class diagram notation to define the constructs provided by this package. We also use color in the diagrams to carry additional information for the benefit of those viewing the document on media that can display color. The following are the colors we use and what they represent:

- ■ *Black*: Items colored black in the UML diagrams are components taken unchanged from their definition in the SBML Level 3 Core specification document.

- ■ *Green*: Items colored green are components that exist in SBML Level 3 Core, but are extended by this package. Class boxes are also drawn with dashed lines to further distinguish them.

- ■ *Blue*: Items colored blue are new components introduced in this package specification. They have no equivalent in the SBML Level 3 Core specification.

- ■ *Red lines*: Classes with red lines in the corner are fully defined in a different figure.

We also use the following typographical conventions to distinguish the names of objects and data types from other entities; these conventions are identical to the conventions used in the SBML Level 3 Core specification document:

AbstractClass: Abstract classes are never instantiated directly, but rather serve as parents of other classes. Their names begin with a capital letter and they are printed in a slanted, bold, sans-serif typeface. In electronic document formats, the class names defined within this document are also hyperlinked to their definitions; clicking on these items will, given appropriate software, switch the view to the section in this document containing the definition of that class. (However, for classes that are unchanged from their definitions in SBML Level 3 Core, the class names are not hyperlinked because they are not defined within this document.)

Class: Names of ordinary (concrete) classes begin with a capital letter and are printed in an upright, bold, sans-serif typeface. In electronic document formats, the class names are also hyperlinked to their definitions in this specification document. (However, as in the previous case, class names are not hyperlinked if they are for classes that are unchanged from their definitions in the SBML Level 3 Core specification.)

`SomeThing`, `otherThing`: Attributes of classes, data type names, literal XML, and tokens *other* than SBML class names, are printed in an upright typewriter typeface. Primitive types defined by SBML begin with a capital letter; SBML also makes use of primitive types defined by XML Schema 1.0 (Biron and Malhotra, 2000; Fallside, 2000; Thompson et al., 2000), but unfortunately, XML Schema does not follow any capitalization convention and primitive types drawn from the XML Schema language may or may not start with a capital letter.

`[elementName]`: In some cases, an element may contain a child of any class inheriting from an abstract base class. In this case, the name of the element is indicated by giving the abstract base class name in brackets, meaning that the actual name of the element depends on whichever subclass is used, with capitalization following the capitalization of the name in brackets.

For other matters involving the use of UML and XML, we follow the conventions used in the SBML Level 3 Core specification document.

3.2 Namespace URI and other declarations necessary for using this package

Every SBML Level 3 package is identified uniquely by an XML namespace URI. For an SBML document to be able to use a given Level 3 package, it must declare the use of that package by referencing its URI. This version of the Distributions package has two URIs, depending on which version of core SBML is being used:

> "`http://www.sbml.org/sbml/level3/version1/distrib/version1`"   1

> "`http://www.sbml.org/sbml/level3/version2/distrib/version1`"   2

Note that the Distributions package may be used with both SBML Level 3 Version 1 and SBML Level 3 Version 2   3
documents, with the only semantic change between the two present in the **DistribBase** class. Note that if used with   4
SBML Level 3 Version 1 the corresponding Distributions namespace for SBML Level 3 Version 1 must be used and   5
similarly if SBML Level 3 Version 2 is being used, the Distributions namespace must be the one for SBML Level 3   6
Version 2.   7

In addition, SBML documents using a given package must indicate whether the package may be used to change the   8
mathematical meaning of SBML Level 3 Core elements. This is done using the attribute `required` on the `<sbml>`   9
element in the SBML document. For the Distributions package, the value of this attribute must be "`true`", as it   10
defined new csymbols that may be used in any MathML. Note that the value of this attribute must *always* be set to   11
"`true`", even if the particular model does not contain any of these csymbols.   12

The following fragment illustrates the beginning of a typical SBML model using SBML Level 3 Version 1 and this   13
version of the Distributions package: 14

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version1/core" level="3" version="1"
      xmlns:distrib="http://www.sbml.org/sbml/level3/version1/distrib/version1"
      distrib:required="true">
```

The following fragment illustrates the beginning of a typical SBML model using SBML Level 3 Version 2 and this   21
version of the Distributions package: 22

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version2/core" level="3" version="2"
      xmlns:distrib="http://www.sbml.org/sbml/level3/version2/distrib/version1"
      distrib:required="true">
```

There is no difference between these two namespaces, and all package semantics are identical.   29

XML Namespace use   30

For element names, XML has clear rules about how to declare and use namespaces. In typical SBML documents, the   31
Distributions namespace will be defined as above, and elements will therefore need to be prefixed with "`distrib:`".   32

In contrast to element names, XML *attribute* names are completely defined by the element in which they appear,   33
and never have a 'default' namespace defined: the element itself declares whether any attributes should be defined   34
with a namespace prefix or not.   35

As is convention for SBML packages, any attribute that appears in a UML diagram in this specification may *either*   36
be defined with no namespace prefix, *or* be defined with the *distrib* namespace as a prefix. (No attributes are   37
defined here as extentions of existing core SBML elements, and thus none of them are required to have the *distrib*   38
namespace as a prefix.)   39

3.3 Primitive data types   40

The Distributions package uses data types described in Section 3.1 of the SBML Level 3 Core specification, and adds   41
the additional primitive types described below.   42

3.3.1 Type `ExternalRef`   43

The type `ExternalRef` is derived from `string` with the additional requirement that it be a valid URI. An `ExternalRef`   44
is used in Distributions to point to ontologies such as ProbOnto (Swat et al., 2016) which contain defined distribu-   45
tions and parameters.   46

3.4 Defining Distributions

3.4.1 The approach

The Distributions package has two very simple purposes. First, it provides a mechanism for sampling a random value from a probability distribution. This implies that it must define the probability distribution and then must sample a random value from that distribution.

Secondly, it provides a mechanism for describing elements with information about their uncertainty. One common use case for this is to provide the standard deviation for a value. Another may be describing a parameter's distribution, so that a better search can be performed in a parameter scan experiment.

The first purpose is achieved by allowing new MathML elements, and the second by extending **SBase**, which in turn uses the **Distribution** and **UncertStatistics** classes, modeled after UncertML. Several distributions and statistics are defined explicitly in this specification, but more can be defined by referencing an external ontology such as ProbOnto through the **ExternalDistribution** and **ExternalParameter** classes.

When a call to a distribution is defined in the extended **Math**, it is sampled when it is invoked. If a particular returned value needs to be used again, that value must be assigned to a parameter first, such as through the use of an **InitialAssignment** or **EventAssignment**. When a distribution is defined elsewhere, that information may be used outside of the model, using whatever methodology is appropriate to answer the question being pursued.

3.5 Extended Math

To allow quick access to a variety of common functions, the Distributions package allows the use of new types of `csymbol` elements anywhere that **Math** is used. These `csymbol`s are functions, and therefore must be the first child of an `apply` element, and their arguments are predefined: you cannot call $normal(mean, variance)$, because the definition of the `normal csymbol` is $normal(mean, stdev)$.

The newly-allowed `csymbol` elements are defined in the following table:

URL	arguments
http://www.sbml.org/sbml/symbols/distrib/normal	$normal(mean, stdev)$ or $normal(mean, stdev, min, max)$
http://www.sbml.org/sbml/symbols/distrib/uniform	$uniform(min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/bernoulli	$bernoulli(prob)$
http://www.sbml.org/sbml/symbols/distrib/normal/binomial	$binomial(nTrials, probabilityOfSuccess)$ or $binomial(nTrials, probabilityOfSuccess, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/cauchy	$cauchy(location, scale)$ or $cauchy(location, scale, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/chi-square	$chisquare(degreesOfFreedom)$ or $chisquare(degreesOfFreedom, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/exponential	$exponential(rate)$ or $exponential(rate, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/gamma	$gamma(shape, scale)$ or $gamma(shape, scale, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/laplace	$laplace(location, scale)$ or $laplace(location, scale, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/log-normal	$lognormal(shape, scale)$ or $lognormal(shape, scale, min, max$
http://www.sbml.org/sbml/symbols/distrib/normal/poisson	$poisson(rate)$ or $poisson(rate, min, max)$
http://www.sbml.org/sbml/symbols/distrib/normal/rayleigh	$rayleigh(scale)$ or $rayleigh(scale, min, max)$

Many of the distributions take exactly two or four arguments (or exactly one or three arguments). For those functions, the optional last two arguments are 'min' and 'max', for when the draw from the distribution is constrained to be between those two values. For all functions, the 'min' boundary is *inclusive*; that is, a value of `min` may be returned by the function (though this may be very unlikely for draws from a continuous distribution). For all continuous distributions, the 'max' boundary is *not inclusive*; that is, a value of `max` will never be returned. The continuous distributions are `normal`, `cauchy`, `chisquare`, `exponential`, `gamma`, `laplace`, `lognormal`, and `rayleigh`. For the discrete distributions, the 'max' boundray is *inclusive*: that is, a value of `max` may indeed be returned. The discrete distributions are `binomial` and `poisson`.

Fallback functions

If an interpreter is unable to calculate one or more of the above extended MathML functions, it might simply fail, or it might choose to return the mean of the given function instead. In either case, it is a good idea to inform the user that the model cannot be interpreted by the software as intended.

The following mean values may be used as a fallback for software that cannot perform draws from a distribution. Note that truncated versions of these functions will have different means. Note also that the `cauchy` distribution has no mean, by definition.

Function	Fallback (mean)
$normal(mean, stdev)$	$mean$
$uniform(min, max)$	$\frac{min+max}{2}$
$bernoulli(prob)$	$prob$
$binomial(nTrials, probabilityOfSuccess)$	$nTrials * probabilityOfSuccess$
$cauchy(location, scale)$	$undefined$
$chisquare(degreesOfFreedom)$	$degreesOfFreedom$
$exponential(rate)$	$rate^{-1}$
$gamma(shape, scale)$	$shape * scale$
$laplace(location, scale)$	$location$
$lognormal(shape, scale)$	$e^{scale+\frac{shape}{2}}$
$poisson(rate)$	$rate$
$rayleigh(scale)$	$scale\sqrt{\frac{pi}{2}}$

3.6 Discrete vs. continuous sampling

MathML csymbols may be used in SBML Level 3 Core in both discrete and continuous contexts: **InitialAssignment**, **EventAssignment**, **Priority**, and **Delay** elements are all discrete, while **Rule**, **KineticLaw**, and **Trigger** elements are all continuous in time. For discrete contexts, the behavior of *distrib*-extended **FunctionDefinition** elements is well-defined: one or more random values are sampled from the distribution each time the function definition is invoked. Each invocation implies one sampling operation. In continuous contexts, however, their behavior is ill-defined. More information than is defined in this package (such as autocorrelation values or full conditional probabilities) would be required to make random sampling tractable in continuous contexts, and is beyond the scope of this version of the package. If some package is defined in the future that adds this information, or if custom annotations are provided that add this information, such models may become simulatable. However, this package does not define how to handle sampling in continuous contexts, and recommends against it: a warning may be produced by any software encountering the use of a *distrib*-extended MathML in a continuous context. Assuming such models are desirable, and the information is not provided in a separate package, this information may be incorporated into a future version of this specification.

Any other package that defines new contexts for MathML will also either be discrete or continuous. Discrete situations (such as those defined in the Qualitative Models package) are, as above, well-defined. Continuous situations (as might arise within the Spatial Processes package, over space instead of over time) will most likely be ill-defined. Those packages must therefore either define for themselves how to handle *distrib*-extended MathML elements, or leave it to some other package/annotation scheme to define how to handle the situation.

3.7 Examples using the extended `csymbol` elements

Several examples are given below that illustrate various uses of the new `csymbol` elements.

3.7.1 Using a normal distribution

In this example, the initial value of 'y' is set as a draw from a normal distribution:

```
...
  <initialAssignment symbol="y">
    <math xmlns="http://www.w3.org/1998/Math/MathML">
      <apply>
        <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/distrib/normal">
          normal
        </csymbol>
        <ci> z </ci>
        <cn> 10 </cn>
      </apply>
    </math>
  </initialAssignment>
...
```

This use would apply a draw from a normal distribution with mean "`z`" and standard deviation "`10`" to the SBML element "`y`".

3.7.2 Defining a truncated normal distribution

When used with four arguments instead of two, the normal distribution is truncated:

```
...
  <initialAssignment symbol="y">
    <math xmlns="http://www.w3.org/1998/Math/MathML">
      <apply>
        <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/distrib/normal">
          normal
        </csymbol>
        <ci> z </ci>
        <cn type="integer"> 10 </cn>
        <apply>
          <minus/>
          <ci> z </ci>
          <cn type="integer"> 2 </cn>
        </apply>
        <apply>
          <plus/>
          <ci> z </ci>
          <cn type="integer"> 2 </cn>
        </apply>
      </apply>
    </math>
  </initialAssignment>
...
```

This use would apply a draw from a normal distribution with mean z, standard deviation 10, lower bound $z - 2$ (inclusive) and upper bound $z + 2$ (not inclusive) to the SBML element "`y`".

3.7.3 Defining conditional events

Simultaneous events in SBML are ordered based on their **Priority** values, with higher values being executed first, and potentially cancelling events that fire after them. In this example, two simultaneous events have priorities set with csymbols defined in *distrib*. The event E0 has a priority of "`uniform(0,1)`", while the event E1 has a priority of "`uniform(0,2)`". This means that 75% of the time, event E1 will have a higher priority than E0, and will fire first, assigning a value of 5 to parameter "`x`". Because this negates the trigger condition for E0, which is set `persistent`="`false`", this means that E0 never fires, and the value of x remains at 5. The remaining 25% of the time, the reverse happens, with E0 setting the value of x to 3 instead.

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version2/core"
      xmlns:distrib="http://www.sbml.org/sbml/level3/version2/distrib/version1"
      level="3" version="2" distrib:required="true">
  <model metaid="__main" id="__main">
    <listOfParameters>
      <parameter metaid="__main.x" id="x" value="0" constant="false"/>
    </listOfParameters>
    <listOfEvents>
      <event id="E0" useValuesFromTriggerTime="true">
        <trigger initialValue="true" persistent="false">
          <math xmlns="http://www.w3.org/1998/Math/MathML">
            <apply>
              <and/>
              <apply>
                <gt/>
                <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/time">
                        time </csymbol>
                <cn type="integer"> 2 </cn>
              </apply>
              <apply>
                <lt/>
                <ci> x </ci>
                <cn type="integer"> 1 </cn>
              </apply>
            </apply>
          </math>
        </trigger>
        <priority>
          <math xmlns="http://www.w3.org/1998/Math/MathML">
            <apply>
              <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/distrib/uniform">
                        uniform </csymbol>
              <cn type="integer"> 0 </cn>
              <cn type="integer"> 1 </cn>
            </apply>
          </math>
        </priority>
        <listOfEventAssignments>
          <eventAssignment variable="x">
            <math xmlns="http://www.w3.org/1998/Math/MathML">
              <cn type="integer"> 3 </cn>
            </math>
          </eventAssignment>
        </listOfEventAssignments>
      </event>
      <event id="E1" useValuesFromTriggerTime="true">
        <trigger initialValue="true" persistent="false">
          <math xmlns="http://www.w3.org/1998/Math/MathML">
            <apply>
              <and/>
              <apply>
                <gt/>
                <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/time">
```

```
            time </csymbol>
        <cn type="integer"> 2 </cn>
      </apply>
      <apply>
        <lt/>
        <ci> x </ci>
        <cn type="integer"> 1 </cn>
      </apply>
    </apply>
  </math>
</trigger>
<priority>
  <math xmlns="http://www.w3.org/1998/Math/MathML">
    <apply>
      <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/distrib/uniform">
              uniform </csymbol>
      <cn type="integer"> 0 </cn>
      <cn type="integer"> 2 </cn>
    </apply>
  </math>
</priority>
<listOfEventAssignments>
  <eventAssignment variable="x">
    <math xmlns="http://www.w3.org/1998/Math/MathML">
      <cn type="integer"> 5 </cn>
    </math>
  </eventAssignment>
</listOfEventAssignments>
    </event>
  </listOfEvents>
  </model>
</sbml>
```


*Figure 1: The definition of the extended **SBase** class to include a new optional **Uncertainty** child, which in turn has optional **UncertStatistics** and **Distribution** children. Intended for use with any element with mathematical meaning, or with a **Math** child.*

3.8 The extended *SBase* class

As can be seen in Figure 1, the SBML base class **SBase** is extended to include an optional **Uncertainty** child, which in turn contains an optional **Distribution** child, and an optional **UncertStatistics** child, either or both of which may be used to include information about the uncertainty of the parent element. In SBML Level 3 Core, one should only extend those **SBase** elements with mathematical meaning (**Compartment**, **Parameter**, **Reaction**, **Species**, and **SpeciesReference**), or those **SBase** elements with **Math** children (**Constraint**, **Delay**, **EventAssignment**, **FunctionDefinition**, **InitialAssignment**, **KineticLaw**, **Priority**, **Rule**, and **Trigger**). The **Uncertainty** child is added to **SBase** instead of to each SBML element so that other packages inherit the ability to extend their own elements in the same fashion: for example, the **FluxBound** class from the Flux Balance Constraints package has mathematical

meaning, and could be given an **Uncertainty** child containing information about the distribution or set of samples from which it was drawn. Similarly, the **FunctionTerm** class from the Qualitative Models package has a **Math** child, and could be extended.

A few SBML elements can interact in interesting ways that can confuse the semantics here. A **Reaction** element and its **KineticLaw** child, for example, both reference the exact same mathematics, so only one should be extended with a child **Uncertainty**. Similarly, the uncertainty of an **InitialAssignment** will be identical to the uncertainty of the element it assigns to, and therefore only one should be extended.

Other elements not listed above should probably not be given an **Uncertainty** child, as it would normally not make sense to talk about the uncertainty of something that doesn't have a corresponding mathematical meaning. However, because packages or annotations can theoretically give new meaning (including mathematical meaning) to elements that previously did not have them, this is not a requirement.

It is important to note that the uncertainty described either by the **Distribution** or the **UncertStatistics** elements are defined as being the uncertainty at the moment the element's mathematical meaning is calculated, and does not describe the uncertainty of how that element changes over time. For a **Species**, **Parameter**, **Compartment**, and **SpeciesReference**, this means that it is the uncertainty of their initial values, and does not describe the uncertainty in how those values evolve in time. The reason for this is that other SBML constructs all describe how (or if) the values change in time, and it is those other constructs that should be used to describe a symbol's time-based uncertainty. For example, a **Species** whose initial value had uncertainty due to instrument precision could have an **Uncertainty** child describing this. A **Species** whose value was known to change over time due to unknown processes, but which had a known average and standard deviation could be given an **AssignmentRule** that set that **Species** amount to the known average, and the **AssignmentRule** itself could be given an **UncertStatistics** child describing the standard deviation of the variability.

3.9 The DistribBase class

The **DistribBase** class is an abstract base class which is the parent class for every class in this Distributions package. Its purpose is to replicate within the Distributions package what changed between SBML Level 3 Version 1 and SBML Level 3 Version 2: the addition of an optional `id` and `name` attribute to *SBase*. By adding these attributes here, *distrib* may be used completely exchangeably between Level 1 and Level 2 documents without any other modifications.
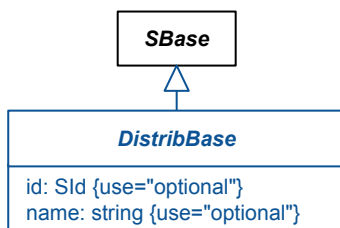
Figure 2: *The definition of the* **DistribBase** *class. The* `id` *and* `name` *attributes defined are optional, and are identical to the ones they inherit in SBML Level 3 Version 2 documents from* **SBase**.

The meaning of these attributes is identical, regardless of the level/version of the document in which they appear.

The `id` attribute is of type `SId`, and must be unique among other ids in the `SId` namespace in the parent **Model**, and has no mathematical meaning, unless stated otherwise in the definition of that object. The `name` attribute is of type `string`, and is provided to allow the user to define a human-readable label for the object. It has no uniqueness restrictions.
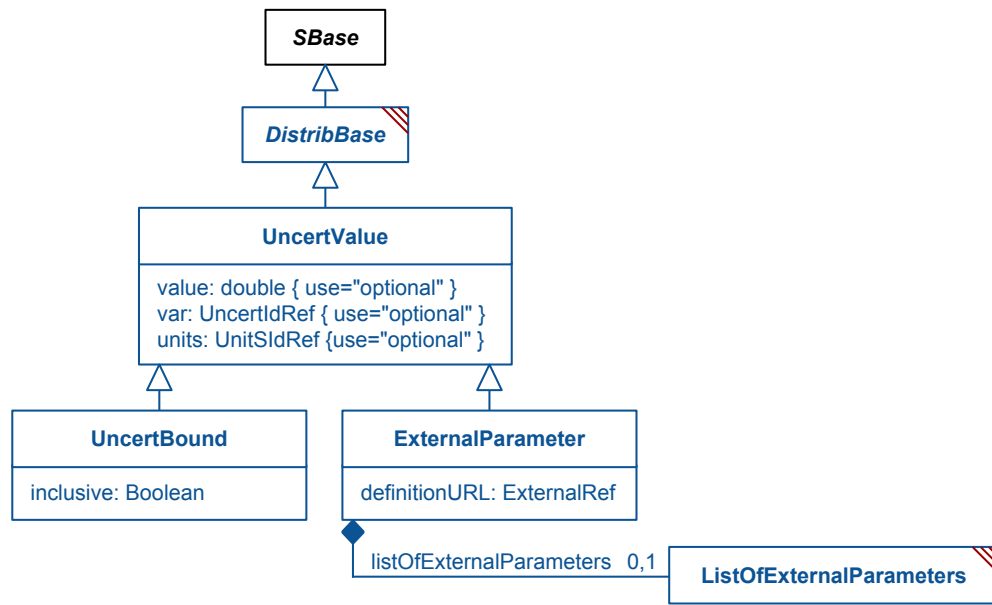
*Figure 3: The definition of the **UncertValue**, **UncertBound**, and **ExternalParameter** classes. These classes all describe a way to define or reference an element with mathematical meaning. The **UncertBound** class additionally defines whether it is considered to be inclusive or not. The meaning of an **ExternalParameter** is defined by its* `definitionURL`, *as well as its children.*

3.10 The UncertValue class

The **UncertValue** class provides two optional attributes, exactly one of which must be defined. The `value` attribute (of type `double`) is used when the **UncertValue** represents a particular number, and the `var` attribute (of type `UncertIdRef`) is used when the **UncertValue** represents a referenced element with mathematical meaning.

The optional `units` attribute may be used to indicate the units of the `val` attribute. As such, it may only be defined if the **UncertValue** has a defined `value` attribute, and not if it has a defined `var` attribute. (In the latter case, the units may be obtained from the referenced element.)

Any given **UncertValue** in a **Distribution** will have an element name specific to the parameter it represents within that **Distribution**. So, for example, a **NormalDistribution** will have one child **UncertValue** with the name "`<mean>`", and might have another **UncertValue** child with the name "`<stddev>`". All these parameters are defined as the same class for simplicity, since all of them merely need a way to reference a value.

3.10.1 Attributes inherited from SBase

An **UncertValue** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name` attributes as described in Section 3.9 on the previous page. The `id` of a **UncertValue** takes the mathematical value of its `value` attribute if that attribute is defined, and the mathematical value of the corresponding `var` if that attribute is defined. This meaning may be used in other contexts, but that meaning may not be set directly by any other SBML element of any level, version, or package. If setting the value is desired, the `var` attribute should be used, and that referenced element set as per normal SBML procedures. The meaning is provided mostly to allow access to the `val` attribute, which otherwise would be undiscoverable to any other SBML element.

3.11 The UncertBound class

The **UncertBound** class inherits from **UncertValue** and adds a single required Boolean attribute `inclusive`. This attribute indicates whether the value the bound represents is to be included in that range ("`true`") or not ("`false`"). This allows the creation of either 'open' or 'closed' boundaries of the ranges it is used to define.

3.12 The ExternalParameter class

The **ExternalParameter** class is provided to allow a modeler to encode externally-provided parameters not otherwise explicitly handled by this specification. The range of possibilities is vast, so modelers should ensure that the tool they wish to use encodes support for any **ExternalParameter** they define.

The **ExternalParameter** inherits from **UncertValue**, and adds the required attribute `definitionURL`, which is of type `ExternalRef`, and an optional child **ListOfExternalParameters**. The `definitionURL` must be a URI that defines a valid distribution-related parameter. It is strongly recommended that modelers use distribution parameters from ProbOnto (`http://probonto.org/`) and other statistical parameters from STATO (`https://www.ebi.ac.uk/ols/ontologies/stato`), as consistently referencing a single ontology will improve exchangeability.

The child **ListOfExternalParameters** is provided because some parameters may themselves need further parameterization. For example, a mixture distribution defined as an **ExternalDistribution** would contain as child **ExternalParameter** objects those other base distributions that were mixed in the overall distribution. Those base distributions would need to define their own parameterization, which could be accomplished here with child **ExternalParameter** objects. Similarly, ranges or categories might also need to be further defined with reference to child **ExternalParameter** objects that would be considered to 'belong' to the parent **ExternalParameter**.

The referenced parameter is then the parameter defined by this **ExternalParameter**, along with any further parameterization provided by its own children **ExternalParameter** elements.

Some external parameters are not single-value, but are multi-value. In these cases, you will either need several **ExternalParameter** objects, or a way to reference an SBML element extended to be defined as an array (such as the Arrays package).

3.13 The Uncertainty class

The **Uncertainty** class has two optional children: an **UncertStatistics** child and a **Distribution** child. Either or both may be used to store information about the uncertainty of an element, e.g. the distribution it was drawn from or its standard deviation. The **Uncertainty** may be annotated to provide additional information.

Note that for elements that change in value over time, the described uncertainty applies only to the element's initial state, and not to how it changes in time. For typical simulations, this means the element's initial assignment.

? Lucian: At this point, I realize that I should have nixed the separate 'UncertStatistics' and just put everything there as the child of **Uncertainty**. If nobody's using this yet, maybe I will.

The units of uncertainty values

The units of uncertainty statistics and distributions are generally either dimensionless or the same as the units of the parent, according to the formula that defines the value. A `mean` and a `standardDeviation`, for example, are always the same units as the parent, while a `coefficientOfVariation` is dimensionless.

The uncertainty of a Species

A **Species** is a unique SBML construct in that its value is either an amount or a concentration, depending on the value of its `hasOnlySubstanceUnits` attribute ("`true`" for amount, or "`false`" for concentration). The value of its uncertainty tracks with this: if the value of the parent **Species** `hasOnlySubstanceUnits` is "`true`", the uncertainty is in terms of amounts, and if "`false`", the uncertainty is in terms of concentration.

If a **Species** is being modeled in SBML in amounts, but was measured in terms of its concentration, or visa versa, an **InitialAssignment** should be created that explicitly handles this conversion and assigns the appropriate value to the **Species**, as in the example below.

```
...
    <listOfCompartments>
      <compartment id="C" spatialDimensions="3" size="2" constant="true">
```

```
          <distrib:uncertainty>                                              1
            <distrib:uncertStatistics>                                       2
              <distrib:standardDeviation distrib:value="0.15"/>              3
            </distrib:uncertStatistics>                                      4
          </distrib:uncertainty>                                             5
        </compartment>                                                       6
    </listOfCompartments>                                                    7
    <listOfSpecies>                                                          8
      <species id="S_amt" compartment="C" hasOnlySubstanceUnits="true"       9
             boundaryCondition="false" constant="false"/>                    10
    </listOfSpecies>                                                         11
    <listOfParameters>                                                       12
      <parameter id="S_conc" value="3.4" constant="true">                   13
        <distrib:uncertainty>                                               14
          <distrib:uncertStatistics>                                        15
            <distrib:standardDeviation distrib:value="0.3"/>                16
          </distrib:uncertStatistics>                                       17
        </distrib:uncertainty>                                              18
      </parameter>                                                          19
    </listOfParameters>                                                     20
    <listOfInitialAssignments>                                             21
      <initialAssignment symbol="S_amt">                                   22
        <math xmlns="http://www.w3.org/1998/Math/MathML">                  23
          <apply>                                                          24
            <times/>                                                       25
            <ci> S_conc </ci>                                              26
            <ci> C </ci>                                                   27
          </apply>                                                         28
        </math>                                                            29
      </initialAssignment>                                                 30
    </listOfInitialAssignments>                                            31
...                                                                        32
                                                                           33
```

Here, the uncertainty of the species "S_amt" is not set explicitly, and instead can be derived from the uncertainty of   34
the values in its initial assignment ("S_conc" and "C").   35

Propagation of error   *36*

It may be possible to calculate the propagation of error for a simulation of an SBML model. Be advised that this will   37
be a complicated system, and may involve calculating partial derivates of equations that are not explicitly encoded.   38
Many simulators choose instead to estimate the error through stochastic simulations. Either approach should be   39
possible with a properly encoded *distrib* model.   40

3.13.1 Attributes inherited from SBase   *41*

An **Uncertainty** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name`   42
attributes as described in Section 3.9 on page 19. The `id` of an **Uncertainty** has no mathematical meaning.   43

3.14 The UncertStatistics class   44

The **UncertStatistics** class is a collection of zero or more statistical measures related to the uncertainty of the   45
parent SBML element. It contains three types of children: **UncertValue** children, **UncertStatisticSpan** children, and   46
a **ListOfExternalParameters** child, which contains zero or more **ExternalParameter** objects. There are ten possible   47
UncertValue children, and six possible **UncertStatisticSpan** children, defined below.   48

3.14.1 Attributes inherited from SBase   *49*

An **UncertStatistics** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name`   50
attributes as described in Section 3.9 on page 19. The `id` of an **UncertStatistics** has no mathematical meaning.   51

3.15 The **UncertValue** children of **UncertStatistics**

An **UncertValue** is defined in Section 3.10 on page 20. The possible **UncertValue** types are listed below. Each is defined by its element name; the mean would be defined as `<mean>`, the standard deviation as `<standardDeviation>`, etc.

All the definitions below are from an archived copy of the definitions at `http://uncertml.org/`, with the exception of `standardError`, which was added here.

- `coefficientOfVariation`: For a random variable with mean μ and strictly positive standard deviation σ, the coefficient of variation is defined as the ratio $\frac{\sigma}{|\mu|}$. One benefit of using the coefficient of variation rather than the standard deviation is that it is unitless.

- `kurtosis`: The kurtosis of a distribution is a measure of how peaked the distribution is. The kurtosis is defined as $\frac{\mu_4}{\sigma^4}$ where μ_4 is the fourth centred moment of the distribution and σ is its standard deviation.

- `mean`: The arithmetic mean (typically just the mean) is what is commonly called the average. It is defined as $\bar{x} = \frac{1}{n} \cdot \sum_{i=1}^{n} x_i$ where x_i represents with ith observation of the quantity x in the sample set of size n. It is related to the expected value of a random variable, $\mu = E[X]$ in that the population mean, μ, which is the average of all quantities in the population and is typically not known, is replaced by its estimator, the sample mean $\bar{x}$. Note that this statistic does not deal with issues of sample size, rather the mean is taken to refer to the population mean.

- `median`: The median is described as the numeric value separating the higher half of a sample (or population) from the lower half. The median of a finite list of numbers can be found by arranging all the observations from lowest value to highest value and picking the middle one. If there is an even number of observations,
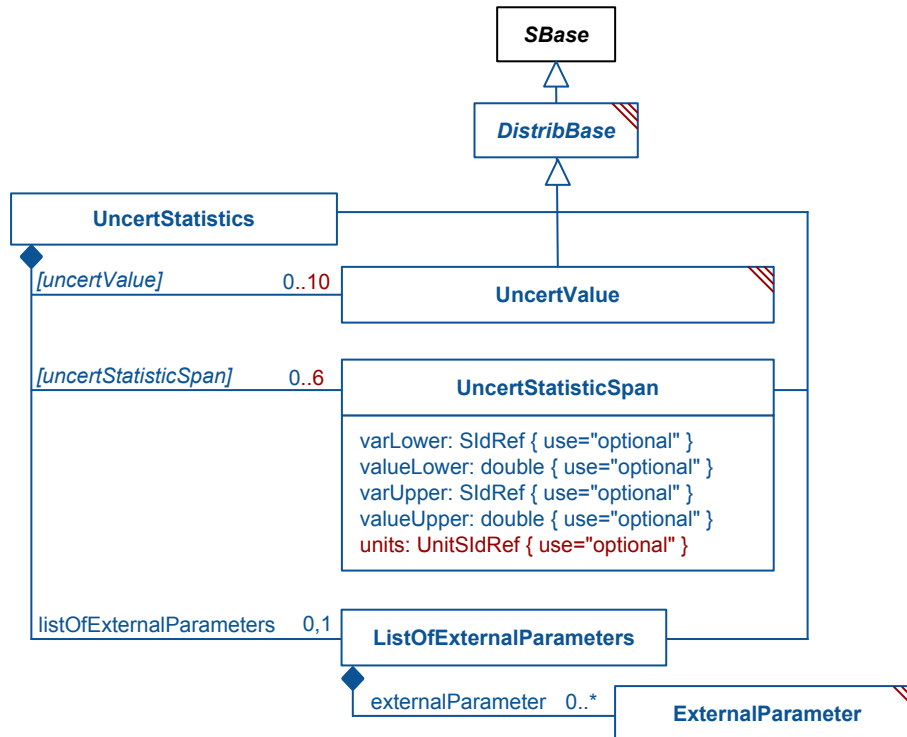
Figure 4: *The definition of the **UncertStatistics**, **UncertStatisticSpan**, and **ListOfExternalParameters** classes. (The **UncertValue** and **ExternalParameter** classes are defined elsewhere and re-used here.) The **UncertStatistics** class has a number of optional children, in three groups: those that can be classified as 'single value' statistics, those that can be classified as a 'span', and those not defined in this specification, but by an external ontology such as ProbOnto or STATO.*

then there is no single middle value, then the average of the two middle values is used. The median is also the 0.5 quantile, or 50th percentile.

- ▪ `mode`: The mode is the value that occurs the most frequently in a data set (or a probability distribution). It need not be unique (e.g. two or more quantities occur equally often) and is typically defined for continuous valued quantities by first defining the histogram, and then giving the central value of the bin containing the most counts.

- ▪ `sampleSize`: The sample size is a direct count of the number of observations made or the number of samples measured. It is used in several other statistical measurements, and can be used to convert one to another.

- ▪ `skewness`: The skewness of a random variable is a measure of how asymmetric the corresponding probability distribution is. The skewness is defined as $\frac{\mu_3}{\sigma^3}$ where μ_3 is the 3rd centred moment of the distribution and σ is its standard deviation.

- ▪ `standardDeviation`: The standard deviation of a distribution or population is the square root of its variance and is given by $\sigma = \sqrt{E[(X-\mu)^2]}$ where $\mu = E[X]$. The population standard deviation is given by $\sigma = \sqrt{\frac{1}{n}\sum_{i=1}^{n}\left(x_i - \bar{x}\right)^2}$ where $\bar{x} = \frac{1}{n}\cdot\sum_{i=1}^{n} x_i$, x_i represents with ith observation of the quantity x in the population of size n. The standard deviation is a widely used measure of the variability or dispersion since it is reported in the natural units of the quantity being considered. Note that if a finite sample of a population has been used then the sample standard deviation is the appropriate unbiased estimator to use.

- ▪ `standardError`: The standard error is the standard deviation of estimates of a population value. If that population value is a mean, this statistic is called the standard error of the mean. It is calculated as the standard deviation of a sample divided by the square root of the number of the sample size. As the sample size increases, the sample size draws closer to the population size, and the standard error approaches zero. $\sigma_{\bar{x}} = \frac{\sigma}{\sqrt{n}}$

- ▪ `variance`: The variance of a random quantity (or distribution) is the average value of the square of the deviation of that variable from its mean, given by $\sigma^2 = Var[X] = E[(X-\mu)^2]$ where $\mu = E[X]$. The complete population variance is given by $\sigma^2 = \frac{1}{n}\sum_{i=1}^{n}\left(x_i - \bar{x}\right)^2$ where $\bar{x} = \frac{1}{n}\cdot\sum_{i=1}^{n} x_i$, x_i represents with ith observation of the quantity x in the population of size n. This is the estimator of the population variance and should be replaced by the sample variance when using samples of finite size.

3.16 The UncertStatisticSpan class

The **UncertStatisticSpan** class defines a span of values that define an uncertainty statistic such as confidence interval or range. It has four optional attributes, `varLower` and `varUpper`, of type `SIdRef`, and `valueLower` and `valueUpper`, of type double. Exactly one of the attributes `varLower` and `valueLower` may be defined, and exactly one of the attributes `varUpper` and `valueUpper` may be defined. If no attributes are defined, the parameters of the span are undefined. If only one attribute is defined (one of the upper or lower attributes), that aspect of the span is defined, and the other end is undefined. The span is fully defined if two attributes (one lower and one upper) are defined.

The value of the lower attribute (whichever is defined) must be lesser or equal to the value of the upper attribute (whichever is defined), at the initial conditions of the model. The **UncertStatistics** element cannot affect the core mathematics of an SBML model, but if it is used in a mathematical context during simulation of the model, this restriction on the attribute values must be maintained, or the **UncertStatisticSpan** object as a whole will be undefined.

Like the `units` attribute on an **UncertValue**, the `units` attribute is provided if either the upper or lower end of the span is defined by value, instead of by reference (i.e. if `valueUpper` and/or `valueLower` is defined). The units on both the upper and lower ends of the span must match each other, if defined. The units for span ends defined by reference may be obtained from the referenced SBML element.

The possible **UncertStatisticSpan** types are listed below, each defining a bounded span of values instead of a single value. The definitions were again taken from an archived copy of http://uncertml.org/:

- **confidenceInterval**: For a univariate random variable x, a confidence interval is a range $[a, b]$, $a < b$, so that x lies between a and b with given probability. For example, a 95% confidence interval is a range in which x falls 95% of the time (or with probability 0.95). Confidence intervals provide intuitive summaries of the statistics of the variable x.

  If x has a continuous probability distribution P, then $[a, b]$ is a 95% confidence interval if $\int_a^b P(x) = 0.95$.

  Unless specified otherwise, the confidence interval is usually chosen so that the remaining probability is split equally, that is $P(x < a) = P(x > b)$. If x has a symmetric distribution, then the confidence intervals are usually centred around the mean. However, non-centred confidence intervals are possible and are better described by their lower and upper quantiles or levels. For example, a 50% confidence interval would usually lie between the 25% and 75% quantiles, but could in theory also lie between the 10% and 60% quantiles, although this would be rare in practice. The **confidenceInterval** allows you the flexibility to specify non-symmetric confidence intervals however in practice we would expect the main usage to be for symmetric intervals.

  The **confidenceInterval** child of a **UncertStatistics** is always the 95% confidence interval. For other confidence intervals, use an **ExternalParameter** instead.

- **credibleInterval**: In Bayesian statistics, a credible interval is similar to a confidence interval determined from the posterior distribution of a random variable x. That is, given a prior distribution $p(x)$ and some observations D, the posterior probability $p(x \mid D)$ can be computed using Bayes theorem. A 95% credible interval is then any interval $[a, b]$ so that $\int_a^b p(x \mid D) = 0.95$, that is the variable x lies in the interval $[a, b]$ with posterior probability 0.95. Note that the interpretation of a credible interval is not the same as a (frequentist) confidence interval.

  The **credibleInterval** child of a **UncertStatistics** is always the 95% credible interval. For other credibility intervals, use an **ExternalParameter** instead.

- **interquartileRange**: The interquartile range is the range between the 1st and 3rd quartiles. It contains the middle 50% of the sample realisations (or of the sample probability). It is typically used and shown in box plots.

- **range**: The range is the interval $[a, b]$ so that $a < b$ and contains all possible values of x. This is also often called the statistical range, which is the distance from the smallest value to the largest value in a sample dataset. For a sample dataset $X = (x_1, ..., x_N)$, the range is the distance from the smallest x_i to the largest. It is often used as a first estimate of the sample dispersion.

3.16.1 Attributes inherited from SBase

An **UncertStatisticSpan** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name` attributes as described in Section 3.9 on page 19. The `id` of an **UncertStatisticSpan** has no mathematical meaning.

3.17 The **ExternalParameter** children of **UncertStatistics**

Any number of **ExternalParameter** children (defined in Section 3.12 on page 21) may be included, each defined by its `definitionURL`.

As examples, the following statistics are not defined by a single value nor by a range, and would therefore be good candidates for encoding with an **ExternalParameter**. These terms were included in the now-defunct UncertML (and the definitions were again taken from an archived copy of `http://uncertml.org/`), and may also be findable in other ontologies such as STATO (which has a searchable database at `https://www.ebi.ac.uk/ols/ontologies/stato`):

- **centredMoment**: For a given positive natural number k, the k^{th} central moment of a random variable x is defined as $\mu_k = E[(x - E[x])^k]$. That is, it is the expected value of the deviation from the mean to the power k. In particular, $\mu_0 = 1$, $\mu_1 = 0$ and μ_2 is the variance of x.

- `correlation`: The correlation between two random variables x_1 and x_2 is the extent to which these variable vary together in a linear fashion. It is characterised by the coefficient $\rho_{1,2} = \frac{E[(x_1 - \mu_1)(x_2 - \mu_2)]}{\sigma_1 \sigma_2}$ where μ_1 and μ_2 are the means of x_1 and x_2 respectively, and σ_1 and σ_2 are their respective standard deviations. Note this is strictly not a description of uncertainty, but it can be useful to represent the correlation between two variables. Generally a covariance specification would be preferred since this describes the uncertainty.

- `decile`: A decile, d, is any of the nine values that divide the sorted quantities into ten equal parts, so that each part represents 1/10 of the sample, population or distribution. The first decile is equivalent to the 10th percentile.

- `moment`: For a given positive natural number k, the k^{th} moment of a random variable x is defined as $\mu_k = E[x^k]$. In particular, $\mu_0 = 1$ and μ_1 is the mean of x. The moments can be defined with respect to some point a, that is $\mu_k(a) = E[(x - a)^k]$. Moments defined about the mean are called centred moments.

- `percentile`: A percentile is the value of a quantity below which a certain percent of values fall. This can be defined for samples, populations and distributions. For finite samples there is no widely accepted method, but all methods essentially rank the quantities and then use some interpolation to compute the percentile, unless the sample size n is a multiple of 100. For probability distributions the inverse cumulative density function can be used. The most widely used method is as follows: to estimate the value, x_p, of the pth percentile of an ascending ordered dataset containing n elements with values $x_1, x_2, ..., x_n$ first compute $\rho = \frac{p}{100}(n-1) + 1$. Now ρ is split into its integer component, k, and decimal component, d, such that $\rho = k + d$. x_p is then calculated as $x_p = x_k + d(x_{k+1} - x_k)$ where $1 < \rho < n$ with special cases $x_p = x_1$ $[\rho = 1]$; x_n $[\rho = n]$.

- `probability`: Given a random variable x with probability density function $f(x)$, the probability that x lies in some part of its domain $\mathscr{X}$ is defined as $P(x \in \mathscr{X}) = \int_{x \in \mathscr{X}} f(x)$. $\mathscr{X}$ can be defined as a lower- or upper-bounded range, e.g. $P(x < 3.2)$ or as the intersection of several such ranges, e.g. $P(x \geq 1.7 \cap x < 3.2)$.

- `quantile`: Given a random variable x, the n-quantiles are the values of x which split the domain into n regions of equal probability. For instance, the k^{th} n-quantile is the value q_k for which $P(x < q_k) = \frac{k}{n}$. For some common values of n, the n-quantiles have additional names, namely quartiles for $n = 4$, deciles for $n = 10$ and percentiles for $n = 100$. More generally, a quantile can be associated to any probability p, so that q is the value of x below which a proportion p of the probability lies, i.e. $P(x < q) = p$. The plot on the right shows the 1st to 9th 10-quantiles (or deciles) for a normal distribution ($\mu = 4$, $\sigma = 1$) as orange dots. The blue curve is the cumulative density function of x. Note how the quantiles split the probability (y-axis) into 10 equal regions.

- `quartile`: The quartiles are the 4-quantiles, that is the 4 values of x below which lies a proportion 0.25, 0.50, 0.75 and 1 of the probability. One can also think of them as the 4 values of x which split the domain into 4 regions of equal probability.

3.18 The ListOfExternalParameters class

The **ListOfExternalParameters** class, like other **ListOf___** classes in SBML Level 3 Core, is a container for zero or more **ExternalParameter** objects. If empty, it simply means that no child **ExternalParameter** objects are defined for its parent, and is equivalent to not including the **ListOfExternalParameters** object at all. This situation might be useful if the list is annotated with the reason why it is empty, for example.

3.19 The Distribution class

The **Distribution** class is the abstract class from which all distributions are derived. They are organized here in much the same way they were in UncertML, by whether they are univariate or multivariate, and whether they are continuous, discrete, or categorical. In addition, the **ExternalDistribution** inherits from **Distribution**, as a 'generic' distribution definition class that allows the user to define any distribution in an external ontology such as ProbOnto.

? Lucian: When these distributions were originally defined, they were being used in extended **FunctionDefinition** elements to define actual draws from the distributions. Now that that part of the spec has been replaced by new

csymbols, these distributions are now solely being used as children of the **Uncertainty** element. If people like this, great; if we want to ditch it for the **ExternalDistribution** construct instead, that's fine. We could even expand this list as some sort of automatic conversion of ProbOnto.

In this draft of the Distributions specification, no mixed distributions and no multivariate distributions are presented, as the author has not seen any call for these distributions specifically, and believes that the generic **ExternalDistribution** distribution could cover those cases on an as-needed basis. If this turns out to not be the case, those distributions will be added to a subsequent version of this specification. The use of the Arrays package would be required for any multivariate distribution.

The full list of distributions is included in Appendix A on page 39 .

3.19.1 Attributes inherited from SBase

A **Distribution** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name` attributes as described in Section 3.9 on page 19. The `id` of a **Distribution** has no mathematical meaning.

3.20 Examples using extended *SBase*

Several examples are given to illustrate the use of the **Uncertainty** class:

3.20.1 Basic *Uncertainty* example

In this examples, a species is given an **Uncertainty** child to describe its standard deviation:

```
...
    <species id="s1" compartment="C" initialAmount="3.22" hasOnlySubstanceUnits="true"
            boundaryCondition="false" constant="false">
      <distrib:uncertainty>
        <distrib:uncertStatistics>
          <distrib:standardDeviation distrib:value="0.3"/>
        </distrib:uncertStatistics>
      </distrib:uncertainty>
    </species>
...
```

Here, the species with an initial amount of 3.22 is described as having a standard deviation of 0.3, a value that might be written as "`3.22 ± 0.3`". This is probably the simplest way to use the package to introduce facts about the uncertainty of the measurements of the values present in the model.

It is also possible to include additional information about the species, should more be known:

```
...
    <species id="s1" compartment="C" initialAmount="3.22" hasOnlySubstanceUnits="true"
            boundaryCondition="false" constant="false">
      <distrib:uncertainty>
        <distrib:uncertStatistics>
          <distrib:mean distrib:value="3.2"/>
          <distrib:standardDeviation distrib:value="0.3"/>
          <distrib:variance distrib:value="0.09"/>
        </distrib:uncertStatistics>
      </distrib:uncertainty>
    </species>
...
```

In this example, the initial amount of 3.22 is noted as having a mean of 3.2, a standard deviation of 0.3, and a variance of 0.09. Note that the standard deviation can be calculated from the variance (or visa versa), but the modeler has chosen to include both here for convenience. Note too that this use of the **Uncertainty** element does not imply that the species amount comes from a normal distribution with a mean of 3.2 and standard deviation of

0.3, but rather that the species amount comes from an unknown distribution with those qualities. If it is known that the value was drawn from a particular distribution, that distribution should be used, rather than the `Mean` and `StandardDeviation` statistical values.

Note also that 3.22 (the `initialAmount`) is different from 3.2 (the `Mean`): evidently, this model was constructed as a realization of the underlying uncertainty, instead of trying to capture the single most likely model of the underlying process.

3.20.2 Defining a Random Variable

In addition to describing the uncertainty about an experimental observation one can also use this mechanism to describe a parameter as a random variable. In the example below the parameter, Z, is defined as following a normal distribution, with a given mean and variance. No value is given for the parameter so it is then up the modeler to decide how to use this random variable. For example they may choose to simulate the model in which case they may provide values for mu_Z and var_Z and then sample a random value from the simulation. Alternatively they may choose to carry out a parameter estimation and use experimental observations to estimate mu_Z and var_Z.

```
<listOfParameters>
  <parameter id="mu_Z" value="10" constant="true"/>
  <parameter id="var_Z" value="0.1" constant="true"/>
  <parameter id="Z" constant="true">
    <distrib:uncertainty>
      <distrib:normalDistribution>
        <distrib:mean distrib:var="mu_Z"/>
        <distrib:variance distrib:var="var_Z"/>
      </distrib:normalDistribution>
    </distrib:uncertainty>
  </parameter>
</listOfParameters>
```

One could also similarly define a parameter that represented gender through two values:

```
...
    <parameter id="gender" constant="false">
      <distrib:uncertainty>
        <distrib:categoricalDistribution>
          <distrib:listOfCategories>
            <distrib:category id="male">
              <distrib:probability distrib:value="0.5"/>
              <distrib:value distrib:value="0"/>
            </distrib:category>
            <distrib:category id="female">
              <distrib:probability distrib:value="0.5"/>
              <distrib:value distrib:value="1"/>
            </distrib:category>
          </distrib:listOfCategories>
        </distrib:categoricalDistribution>
      </distrib:uncertainty>
    </parameter>
...
```

4 Interaction with other packages

4.1 Custom annotations for function definitions

Before this package was available, a collection of SBML simulator authors developed an *ad-hoc* convention for exchanging annotated **FunctionDefinition** objects that represented draws from distributions. This convention is described by Frank T. Bergmann at https://docs.google.com/file/d/0B_wMqVOQLkZ3TVZHblNNRWgzNTg/, and represents a basic starting point for any modeler interested in exchanging SBML models containing draws from distributions.

When implementing Distributions support, it would be possible to include 'backwards' support for this annotation convention by annotating any extended **FunctionDefinition** that happens to match the following distribution to also include these annotations, where appropriate.

The following table is taken from the above document by Frank Bergmann, and can be used to annotate **FunctionDefinition** elements that have been extended by Distributions to perform the same functions, providing the arguments are presented in the same order. The suggested fallback function returns the mean of the distribution.

Id	Name	URL	Fallback
uniform	Uniform distribution	http://en.wikipedia.org/wiki/Uniform_distribution_(continuous)	$lambda(a, b, \frac{a+b}{2})$
normal	Normal distribution	http://en.wikipedia.org/wiki/Normal_distribution	$lambda(m, s, m)$
exponential	Exponential distribution	http://en.wikipedia.org/wiki/Exponential_distribution	$lambda(l, \frac{1}{l})$
gamma	Gamma distribution	http://en.wikipedia.org/wiki/Gamma_distribution	$lambda(a, b, a*b)$
poisson	Poisson distribution	http://en.wikipedia.org/wiki/Poisson_distribution	$lambda(mu, mu)$
lognormal	Lognormal distribution	http://en.wikipedia.org/wiki/Log-normal_distribution	$lambda(z, s, e^{z+\frac{s^2}{2}})$
chisq	Chi-squared distribution	http://en.wikipedia.org/wiki/Chi-squared_distribution	$lambda(nu, nu)$
laplace	Laplace distribution	http://en.wikipedia.org/wiki/Laplace_distribution	$lambda(a, 0)$
cauchy	Cauchy distribution	http://en.wikipedia.org/wiki/Cauchy_distribution	$lambda(a, a)$
rayleigh	Rayleigh distribution	http://en.wikipedia.org/wiki/Rayleigh_distribution	$lambda(s, s*\sqrt{\pi/2})$
binomial	Binomial distribution	http://en.wikipedia.org/wiki/Binomial_distribution	$lambda(p, n, p*n)$
bernoulli	Bernoulli distribution	http://en.wikipedia.org/wiki/Bernoulli_distribution	$lambda(p, p)$

As an example, here is a complete (if small) model that uses the above 'custom annotation' scheme:

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version1/core"
      level="3" version="1">
  <model>
    <listOfFunctionDefinitions>
```

```
   <functionDefinition id="normal">                                                      1
     <annotation>                                                                        2
       <distribution xmlns="http://sbml.org/annotations/distribution"                    3
                definition="http://en.wikipedia.org/wiki/Normal_distribution"/>          4
     </annotation>                                                                       5
     <math xmlns="http://www.w3.org/1998/Math/MathML">                                    6
       <lambda>                                                                          7
         <bvar>                                                                          8
           <ci> mean </ci>                                                               9
         </bvar>                                                                        10
         <bvar>                                                                         11
           <ci> stdev </ci>                                                             12
         </bvar>                                                                        13
         <notanumber/>                                                                  14
       </lambda>                                                                        15
     </math>                                                                            16
   </functionDefinition>                                                                17
 </listOfFunctionDefinitions>                                                           18
 <listOfParameters>                                                                     19
   <parameter id="x" constant="true"/>                                                  20
 </listOfParameters>                                                                    21
 <listOfInitialAssignments>                                                             22
   <initialAssignment symbol="x">                                                       23
     <math xmlns="http://www.w3.org/1998/Math/MathML">                                   24
       <apply>                                                                          25
         <ci> normal </ci>                                                              26
         <cn> 3 </cn>                                                                   27
         <cn> 0.2 </cn>                                                                 28
       </apply>                                                                         29
     </math>                                                                            30
   </initialAssignment>                                                                 31
 </listOfInitialAssignments>                                                            32
 </model>                                                                               33
</sbml>                                                                                34
                                                                                       35
```

And here is the same model, using the 'csymbol' defined in *distrib*:   36

```
                                                                                       37
<?xml version="1.0" encoding="UTF-8"?>                                                  38
<sbml xmlns="http://www.sbml.org/sbml/level3/version2/core"                              39
      xmlns:distrib="http://www.sbml.org/sbml/level3/version2/distrib/version1"          40
      level="3" version="2" distrib:required="true">                                     41
  <model>                                                                               42
    <listOfParameters>                                                                 43
      <parameter id="x" constant="true"/>                                               44
    </listOfParameters>                                                                45
    <listOfInitialAssignments>                                                         46
      <initialAssignment symbol="x">                                                   47
        <math xmlns="http://www.w3.org/1998/Math/MathML">                               48
          <apply>                                                                      49
            <csymbol encoding="text" definitionURL="http://www.sbml.org/sbml/symbols/distrib/normal">  50
                    normal </csymbol>                                                   51
            <cn type="integer"> 3 </cn>                                                 52
            <cn> 0.2 </cn>                                                              53
          </apply>                                                                     54
        </math>                                                                        55
      </initialAssignment>                                                             56
    </listOfInitialAssignments>                                                        57
  </model>                                                                             58
</sbml>                                                                                59
                                                                                       60
```

4.2 The Arrays package

This package is dependent on no other package, but might rely on the Arrays package to provide vector and matrix structures if those are desired/used. Note that currently, the only way to need arrays is if an **ExternalDistribution** or **ExternalParameter** is defined that requires array input or output.

5 Use-cases and examples

The following examples are more fleshed out than the ones in the main text, and/or illustrate features of this package that were not previously illustrated.

5.1 Sampling from a distribution: PK/PD Model

This is a very straightforward use of a log normal distribution. The key point to note is that a value is sampled from the distribution and assigned to a variable when it is invoked in the initialAssignments element in this example. Later use of the variable does not result in re-sampling from the distribution. This is consistent with current SBML semantics.

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version1/core"
  xmlns:distrib="http://www.sbml.org/sbml/level3/version1/distrib/version1"
  level="3" version="1" distrib:required="true">
  <model>
    <listOfCompartments>
      <compartment id="central" size="0" constant="true"/>
      <compartment id="gut" size="0" constant="true"/>
    </listOfCompartments>
    <listOfSpecies>
      <species id="Qc" compartment="central" initialAmount="1" hasOnlySubstanceUnits="true"
        boundaryCondition="false" constant="false"/>
      <species id="Qg" compartment="gut" initialAmount="1" hasOnlySubstanceUnits="true"
        boundaryCondition="false" constant="false"/>
    </listOfSpecies>
    <listOfParameters>
      <parameter id="ka" constant="true"/>
      <parameter id="ke" constant="true"/>
      <parameter id="Cc" constant="false"/>
      <parameter id="Cc_obs" constant="false"/>
    </listOfParameters>
    <listOfInitialAssignments>
      <initialAssignment symbol="central">
        <math xmlns="http://www.w3.org/1998/Math/MathML">
          <apply>
            <csymbol encoding="text"
              definitionURL="http://www.sbml.org/sbml/symbols/distrib/lognormal"> lognormal
            </csymbol>
            <cn> 0.5 </cn>
            <cn> 0.1 </cn>
          </apply>
        </math>
      </initialAssignment>
      <initialAssignment symbol="ka">
        <math xmlns="http://www.w3.org/1998/Math/MathML">
          <apply>
            <csymbol encoding="text"
              definitionURL="http://www.sbml.org/sbml/symbols/distrib/lognormal"> lognormal
            </csymbol>
            <cn> 0.5 </cn>
            <cn> 0.1 </cn>
          </apply>
        </math>
      </initialAssignment>
      <initialAssignment symbol="ke">
        <math xmlns="http://www.w3.org/1998/Math/MathML">
          <apply>
            <csymbol encoding="text"
              definitionURL="http://www.sbml.org/sbml/symbols/distrib/lognormal"> lognormal
            </csymbol>
```

```
                    <cn> 0.5 </cn>
                    <cn> 0.1 </cn>
                  </apply>
                </math>
              </initialAssignment>
      </listOfInitialAssignments>
      <listOfRules>
        <assignmentRule variable="Cc">
          <math xmlns="http://www.w3.org/1998/Math/MathML">
            <apply>
              <divide/>
              <ci> Qc </ci>
              <ci> central </ci>
            </apply>
          </math>
        </assignmentRule>
        <assignmentRule variable="Cc_obs">
          <math xmlns="http://www.w3.org/1998/Math/MathML">
            <apply>
              <plus/>
              <ci> Cc </ci>
              <cn type="integer"> 1 </cn>
            </apply>
          </math>
        </assignmentRule>
      </listOfRules>
      <listOfReactions>
        <reaction id="absorption" reversible="false" fast="false">
          <listOfReactants>
            <speciesReference species="Qg" stoichiometry="1" constant="true"/>
          </listOfReactants>
          <listOfProducts>
            <speciesReference species="Qc" stoichiometry="1" constant="true"/>
          </listOfProducts>
          <kineticLaw>
            <math xmlns="http://www.w3.org/1998/Math/MathML">
              <apply>
                <times/>
                <ci> ka </ci>
                <ci> Qg </ci>
              </apply>
            </math>
          </kineticLaw>
        </reaction>
        <reaction id="excretion" reversible="false" fast="false">
          <listOfReactants>
            <speciesReference species="Qc" stoichiometry="1" constant="true"/>
          </listOfReactants>
          <kineticLaw>
            <math xmlns="http://www.w3.org/1998/Math/MathML">
              <apply>
                <divide/>
                <apply>
                  <times/>
                  <ci> ke </ci>
                  <ci> Qc </ci>
                </apply>
                <ci> central </ci>
              </apply>
            </math>
          </kineticLaw>
        </reaction>
      </listOfReactions>
    </model>
</sbml>
```

5.2 Multiple uses of distributions

In this example, a `normal` csymbol is used in an initial assignment, and `mean` and `standardDeviation` elements are used to denote the uncertainty in the parameter "`V`", and the uncertainty in the initial assignment to "`V`". Note that strictly speaking, one could assume that the uncertainty in the parameter itself was identical to the uncertainty in its initial assigment; both are given here by way of illustration.

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version1/core"
  xmlns:distrib="http://www.sbml.org/sbml/level3/version1/distrib/version1"
  level="3" version="1" distrib:required="true">
  <model>
    <listOfParameters>
      <parameter id="V" constant="true">
        <distrib:uncertainty>
          <distrib:uncertStatistics>
            <distrib:mean distrib:var="V_pop"/>
            <distrib:standardDeviation distrib:var="V_omega"/>
          </distrib:uncertStatistics>
        </distrib:uncertainty>
      </parameter>
      <parameter id="V_pop" value="100" constant="true"/>
      <parameter id="V_omega" value="0.25" constant="true"/>
    </listOfParameters>
    <listOfInitialAssignments>
      <initialAssignment symbol="V">
        <math xmlns="http://www.w3.org/1998/Math/MathML">
          <apply>
            <csymbol encoding="text"
              definitionURL="http://www.sbml.org/sbml/symbols/distrib/normal"> normal
            </csymbol>
            <ci> V_pop </ci>
            <ci> V_omega </ci>
          </apply>
        </math>
        <distrib:uncertainty>
          <distrib:uncertStatistics>
            <distrib:mean distrib:var="V_pop"/>
            <distrib:standardDeviation distrib:var="V_omega"/>
          </distrib:uncertStatistics>
        </distrib:uncertainty>
      </initialAssignment>
    </listOfInitialAssignments>
  </model>
</sbml>
```

5.3 Defining confidence intervals

In this example, several **Parameter** elements are given confidence intervals, and several species are given standard deviations. Each indicates the modeler's assessment of the precision of the estimated given values for those elements.

```
<?xml version="1.0" encoding="UTF-8"?>
<sbml xmlns="http://www.sbml.org/sbml/level3/version2/core"
  xmlns:distrib="http://www.sbml.org/sbml/level3/version2/distrib/version1"
  level="3" version="2" distrib:required="true">
  <model>
    <listOfCompartments>
      <compartment id="C" spatialDimensions="3" size="1" constant="true"/>
    </listOfCompartments>
    <listOfSpecies>
      <species id="S1" compartment="C" initialAmount="5.2"
```

```
          hasOnlySubstanceUnits="false" boundaryCondition="false" constant="false">   1
          <distrib:uncertainty>                                                        2
            <distrib:uncertStatistics>                                                 3
              <distrib:standardDeviation distrib:value="0.3"/>                         4
            </distrib:uncertStatistics>                                                5
          </distrib:uncertainty>                                                       6
        </species>                                                                     7
        <species id="S2" compartment="C" initialAmount="8.7"                          8
          hasOnlySubstanceUnits="false" boundaryCondition="false" constant="false">   9
          <distrib:uncertainty>                                                       10
            <distrib:uncertStatistics>                                                11
              <distrib:standardDeviation distrib:value="0.01"/>                       12
            </distrib:uncertStatistics>                                               13
          </distrib:uncertainty>                                                      14
        </species>                                                                    15
        <species id="S3" compartment="C" initialAmount="1102"                        16
          hasOnlySubstanceUnits="false" boundaryCondition="false" constant="false">  17
          <distrib:uncertainty>                                                      18
            <distrib:uncertStatistics>                                               19
              <distrib:standardDeviation distrib:value="53"/>                        20
            </distrib:uncertStatistics>                                              21
          </distrib:uncertainty>                                                     22
        </species>                                                                   23
        <species id="S4" compartment="C" initialAmount="0.026"                      24
          hasOnlySubstanceUnits="false" boundaryCondition="false" constant="false"> 25
          <distrib:uncertainty>                                                      26
            <distrib:uncertStatistics>                                               27
              <distrib:standardDeviation distrib:value="0.004"/>                     28
            </distrib:uncertStatistics>                                              29
          </distrib:uncertainty>                                                     30
        </species>                                                                   31
      </listOfSpecies>                                                               32
      <listOfParameters>                                                             33
        <parameter id="P1" value="5.13" constant="true">                            34
          <distrib:uncertainty>                                                      35
            <distrib:uncertStatistics>                                               36
              <distrib:confidenceInterval distrib:valueLower="5" distrib:valueUpper="5.32"/>  37
            </distrib:uncertStatistics>                                              38
          </distrib:uncertainty>                                                     39
        </parameter>                                                                 40
        <parameter id="P2" value="15" constant="true">                              41
          <distrib:uncertainty>                                                      42
            <distrib:uncertStatistics>                                               43
              <distrib:confidenceInterval distrib:valueLower="10.22" distrib:valueUpper="15.02"/>  44
            </distrib:uncertStatistics>                                              45
          </distrib:uncertainty>                                                     46
        </parameter>                                                                 47
        <parameter id="P3" value="0.003" constant="true">                           48
          <distrib:uncertainty>                                                      49
            <distrib:uncertStatistics>                                               50
              <distrib:confidenceInterval distrib:valueLower="-0.001" distrib:valueUpper="0.0041"/>  51
            </distrib:uncertStatistics>                                              52
          </distrib:uncertainty>                                                     53
        </parameter>                                                                 54
        <parameter id="P4" value="0.34" constant="true">                            55
          <distrib:uncertainty>                                                      56
            <distrib:uncertStatistics>                                               57
              <distrib:confidenceInterval distrib:valueLower="0.22" distrib:valueUpper="0.51"/>  58
            </distrib:uncertStatistics>                                              59
          </distrib:uncertainty>                                                     60
        </parameter>                                                                 61
        <parameter id="P5" value="92" constant="true">                              62
          <distrib:uncertainty>                                                      63
            <distrib:uncertStatistics>                                               64
              <distrib:confidenceInterval distrib:valueLower="90" distrib:valueUpper="99"/>  65
            </distrib:uncertStatistics>                                              66
```

```
          </distrib:uncertainty>                                                                              1
        </parameter>                                                                                          2
        <parameter id="P6" value="40.002" constant="true">                                                    3
          <distrib:uncertainty>                                                                               4
            <distrib:uncertStatistics>                                                                        5
              <distrib:confidenceInterval distrib:valueLower="40.0018" distrib:valueUpper="40.0021"/>         6
            </distrib:uncertStatistics>                                                                        7
          </distrib:uncertainty>                                                                               8
        </parameter>                                                                                           9
      </listOfParameters>                                                                                      10
    </model>                                                                                                   11
</sbml>                                                                                                        12
                                                                                                               13
```

6 Prototype implementations

As of this writing (February 2019), libsbml has full support for all elements defined in this version of the specification, with the exception of `standardError` and `sampleSize`. Antimony (http://antimony.sf.net/) has support for a the new csymobls for model creation only (no simulation), but does not support the `uncertainty` child of *SBase*. Libroadrunner supports a limited number of distributions from a previous version of this spec.

7 Acknowledgements

Much of the initial concrete work leading to this proposal document was carried out at the Statistical Models Workshop in Hinxton in 2011, which was organised by Nicolas le Novère. A list of participants and recordings of the discussion is available from http://sbml.org/Events/Other_Events/statistical_models_workshop_2011. Before that a lot of the ground work was carried out by Darren Wilkinson who led the discussion on *distrib* at the Seattle SBML Hackathon and before that Colin Gillespie who wrote an initial proposal back in 2005. The authors would also like to thank the participants of the *distrib* sessions during various HARMONY and COMBINE meetings for their excellent contributions in helping revising this proposal; Sarah Keating, Maciej Swat, Nicolas le Novère, and Matthias K onig for useful discussions, corrections and review comments; and Mike Hucka for LaTeX advice and the beautiful template upon which this document is based.

A Distributions

A.1 The UnivariateDistribution class

The **UnivariateDistribution** class is an abstract class that derives from the **Distribution** abstract class, and which has three derived classes itself: **ContinuousUnivariateDistribution**, **DiscreteUnivariateDistribution**, and **CategoricalUnivariateDistribution**. It is provided as a bookkeeping class to distinguish it from other types of distributions.

A.2 The MultivariateDistribution class

The **MultivariateDistribution** class is an abstract class with no derived classes in the current specification, but some could be added in the future. Most likely, it will be removed from the final version of the spec.

A.3 The ContinuousUnivariateDistribution class

The abstract **ContinuousUnivariateDistribution** class is the base class for a wide variety of distributions, all of which describe a potentially-bounded continuous range of probabilities. Many of the most commonly-used distributions such as the **NormalDistribution** and the **UniformDistribution** fall into this category.

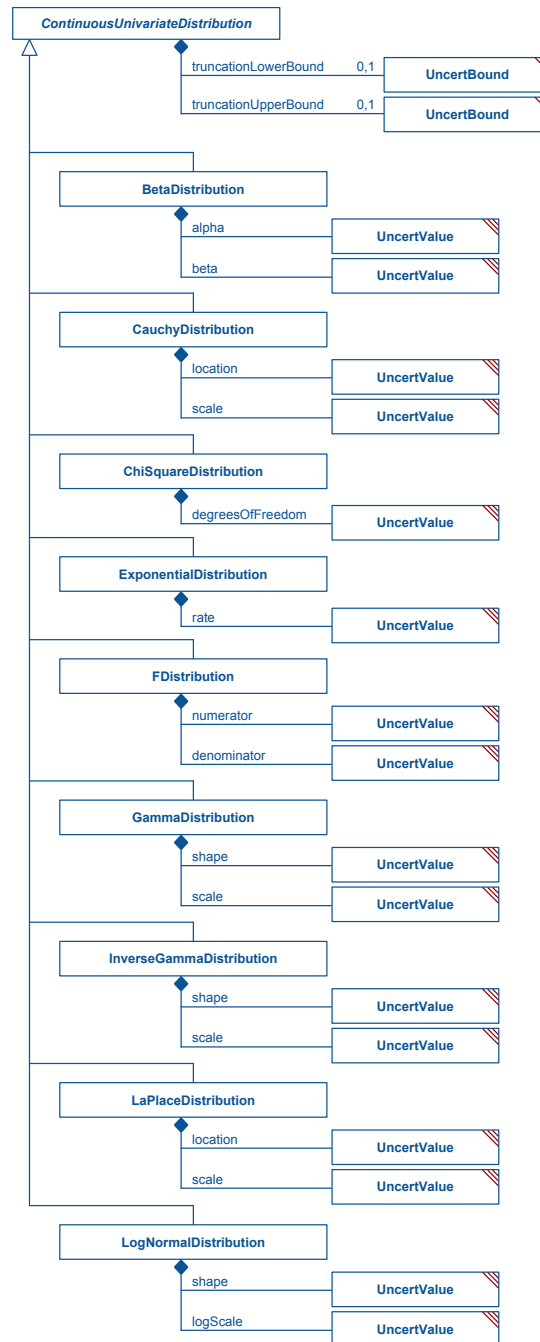All **ContinuousUnivariateDistribution** elements may have two optional children: "`lowerTruncationBound`" and "`upperTruncationBound`", both of the class **UncertBound** (defined below). Either element, if present, limit the range of possible sampled values from the distribution. The "`lowerTruncationBound`" defines the lowest value (inclusive or not, as defined by that element's `inclusive` attribute) that can be sampled, and the "`upperTruncationBound`" defines the highest. If both children are present, the "`lowerTruncationBound`" must either be lower than the "`upperTruncationBound`", or they may be equal, if both bounds are set `inclusive`="`true`". Similarly, some distributions are themselves naturally bound (some may, for example, only return values greater than zero). In those cases, the natural lower bound of the distribution must either be lower than the "`upperTruncationBound`", or be equal to it if the natural lower bound is inclusive, and if the "`upperTruncationBound`" is set `inclusive`="`true`". Similarly, the natural upper bound of the distribution must either be higher than the "`lowerTruncationBound`", or it may be equal to it if the natural upper bound is inclusive and if the "`lowerTruncationBound`" is set `inclusive`="`true`". It may be impossible to determine this from a static analysis of the model, as either or both bound's values may depend on other dynamic variables. If a simulator encounters this situation, the sampled value and the behavior of the simulator are undefined.

If bounded, the cumulative probability that would have been assigned to the region outside the bound is re-assigned proportionally to the rest of the distribution. It should be noted that while discarding any value obtained from the non-truncated version of the distribution and re-sampling is indeed one method that could be used to accomplish this, the efficiency of that algorithm decreases with the width of the allowed window, and indeed is technically zero (and would take an infinite amount of time to complete) should the bounds be equal to one another. Taking any samples obtained outside the bound window and instead returning the boundary value itself is incorrect, and will not result in a proper draw from the defined distribution.

The distributions of this type allowed in this version of the specification are defined in Figure 5 on the next page and Figure 6 on page 41. A full list of all of the distributions is provided in Section A.7 on page 44.

A.4 The DiscreteUnivariateDistribution class

The abstract **DiscreteUnivariateDistribution** class is the base class for a wide variety of distributions, all of which describe a potentially-bounded range of probabilities of discrete values. The most commonly-used distributions in this class is probably the **PoissonDistribution**. Distributions that always return integers fall in this category, which often involve events happening at particular frequencies.

All **DiscreteUnivariateDistribution** elements (like **ContinuousUnivariateDistribution** elements) may have two optional children: "`lowerTruncationBound`" and "`upperTruncationBound`", both of the class **UncertBound** (de-
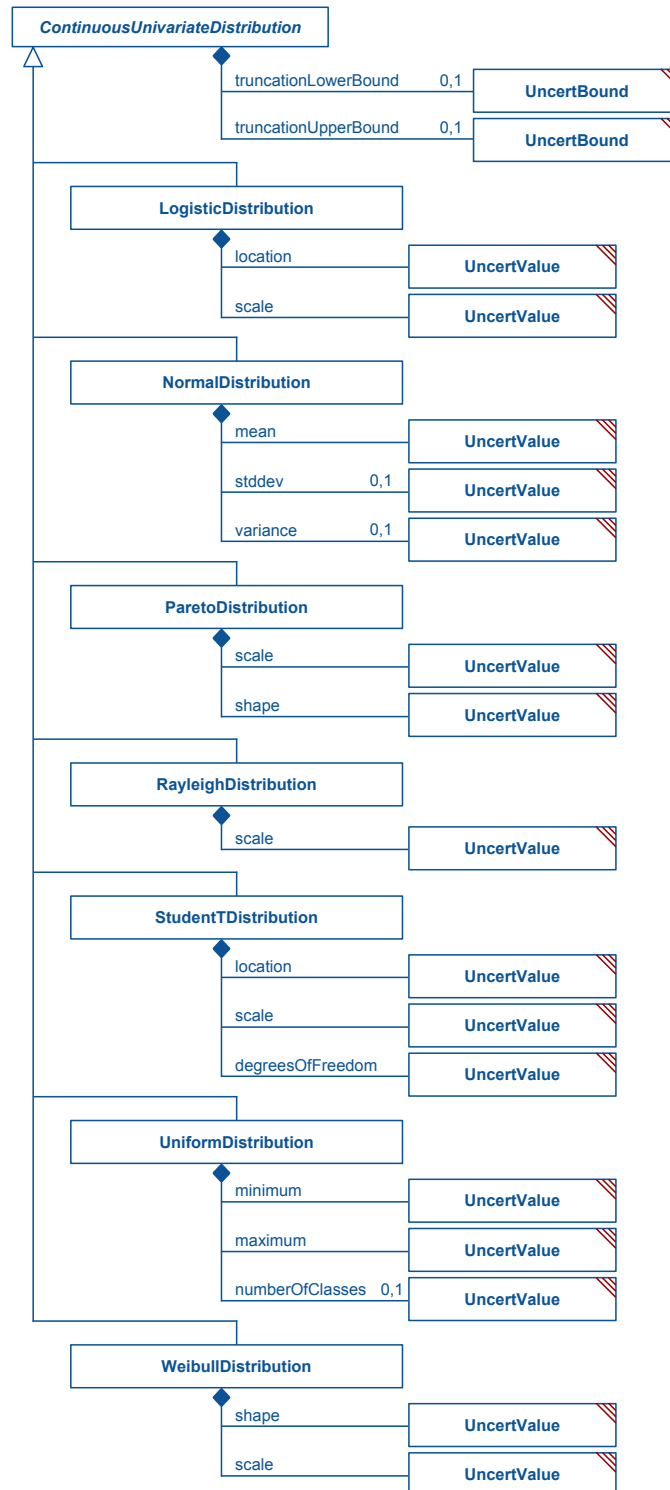
Figure 5: *The definition of the **ContinuousUnivariateDistribution** abstract class, and its **BetaDistribution**, **CauchyDistribution**, **ChiSquareDistribution**, **ExponentialDistribution**, **FDistribution**, **GammaDistribution**, **InverseGammaDistribution**, **LaPlaceDistribution**, and **LogNormalDistribution** children. All may have lower and upper **UncertBound** children, and each has one or more other parameters encoded as **UncertValue** children (both defined below).*

fined below). Either element, if present, limit the range of possible sampled values from the distribution. The "`lowerTruncationBound`" defines the value below which no sampling may take place (inclusive or not, as defined by that element's `inclusive` attribute), and the "`upperTruncationBound`" defines the value above which no sampling may take place. These bounds may fall between the possible discrete values being returned: as an example, for a distribution that returned an integer in the series [0, 1, 2, ...], if it was given a "`lowerTruncationBound`" of 1.5,

Figure 6: *The definition of the **ContinuousUnivariateDistribution** abstract class, and its **LogisticDistribution**, **NormalDistribution**, **ParetoDistribution**, **RayleighDistribution**, **StudentTDistribution**, **UniformDistribution**, and **WeibullDistribution** children. All may have lower and upper **UncertBound** children, and each has one or more other parameters encoded as **UncertValue** children (both defined below).*
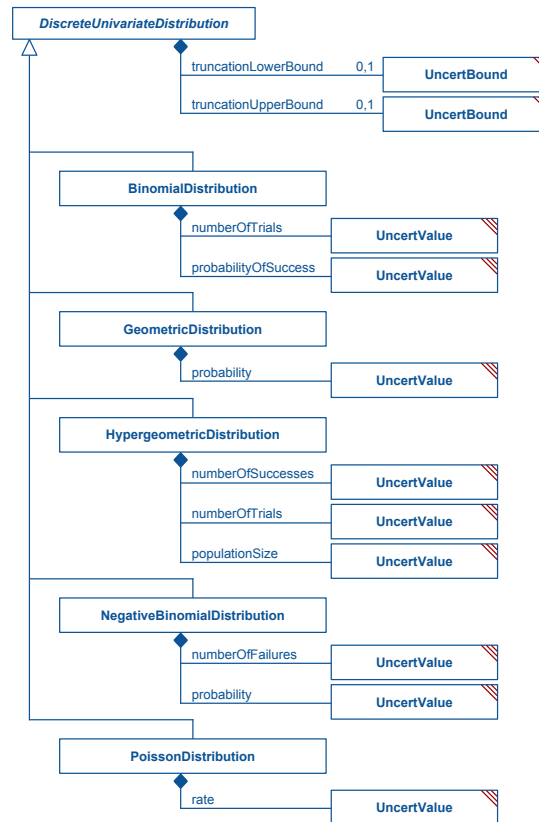
Figure 7: *The definition of the **DiscreteUnivariateDistribution** abstract class, and its **BinomialDistribution**, **GeometricDistribution**, **HypergeometricDistribution**, **NegativeBinomialDistribution**, and **PoissonDistribution** children. All may have lower and upper **UncertBound** children, and each has one or more other parameters encoded as **UncertValue** children (both defined below).*

the lowest value it could return would be 2. In this case, the value of the `inclusive` attribute on the **UncertBound** would be immaterial, as '1.5' could never be returned.

As in the case of the **ContinuousUnivariateDistribution** bounds, if both bounds are present, the "`lowerTruncationBound`" must either be lower than the "`upperTruncationBound`", or they may be equal, if both bounds are set `inclusive`="`true`". Similarly, the discrete distributions are themselves often naturally bound (some may, for example, only return values greater than zero). In those cases, the natural lower bound of the distribution must be either be lower than the "`upperTruncationBound`", or it may be equal to it if the natural lower bound is inclusive, and if the "`upperTruncationBound`" is set `inclusive`="`true`". Similarly, the natural upper bound of the distribution must either be higher than the "`lowerTruncationBound`", or it may be equal to it if the natural upper bound is inclusive and if the "`lowerTruncationBound`" is set `inclusive`="`true`". In addition, if both bounds are defined, they must define a span within which at least one possible sampled discrete value may be found. For a distribution that returns integers, for example, one may not define a lower bound of 1.5 and an upper bound of 1.8, as no integer lies within that range. It may be impossible to determine if any of these rules are violated from a static analysis of the model, as either or both bound's values may depend on other dynamic variables. If a simulator encounters this situation, the sampled value and the behavior of the simulator are undefined.

If bounded, the cumulative probability that would have been assigned to the values outside the bound is re-assigned proportionally to the rest of the distribution. It should be noted that while discarding any value obtained from the non-truncated version of the distribution and re-sampling is indeed one method that could be used to accomplish this, the efficiency of that algorithm decreases with the width of the allowed window, and indeed is technically zero (and could take an infinite amount of time to complete) should the bounds allow only a single discrete value. Taking

any samples obtained outside the bound window and instead returning the boundary value itself is incorrect, and
will not result in a proper draw from the defined distribution.

The distributions of this type allowed in this version of the specification are defined in Figure 7 on the preceding
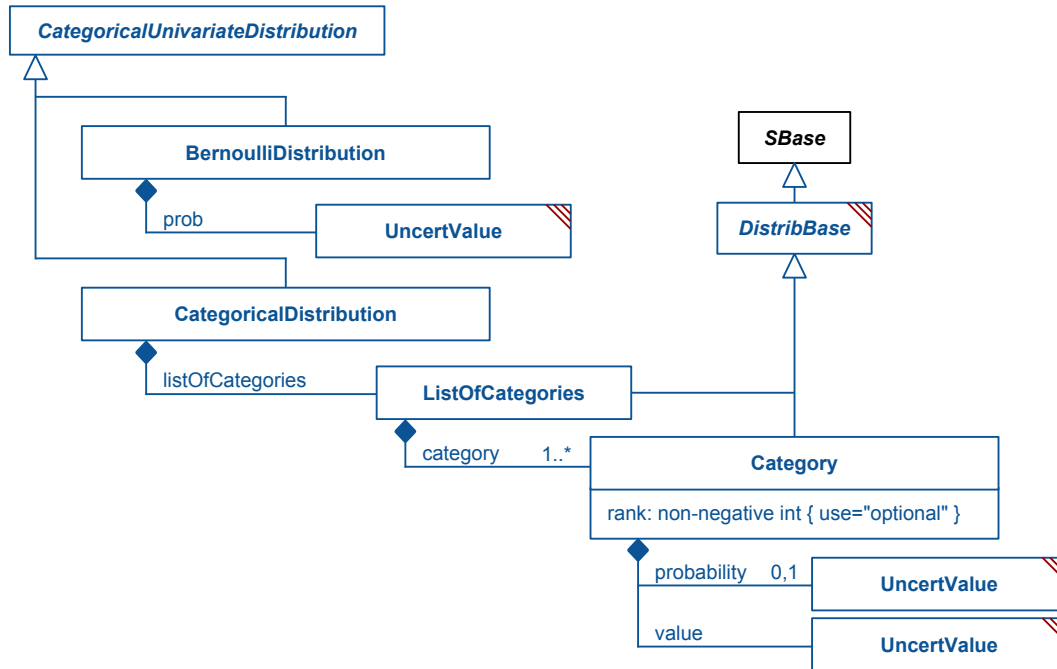page. A full list of all of the distributions is provided in Section A.7 on the next page.

Figure 8: *The definition of the CategoricalUnivariateDistribution abstract class, plus the BernoulliDistribution, CategoricalDistribution, ListOfCategories, and Category classes.*

A.5 The CategoricalUnivariateDistribution class

The **CategoricalUnivariateDistribution** abstract class includes distributions where the various possible sampled
values are each explicitly listed, along with the probability for that sampled value. The sum of these probabilities
must therefore equal 1.0, in order to be valid. This type of distribution class is used for things such as weighted die
rolls, or other situations where particular values are obtained at arbitrary probabilities.

Because each possible sampled value is explicitly listed in an **CategoricalUnivariateDistribution**, it does not have
the optional **UncertBound** values that the other univariate distributions do: if a particular value is not allowed, it is
simply dropped from the list of options, and the probabilities of the other values are scaled accordingly.

A.6 The ExternalDistribution class

The **ExternalDistribution** class is provided to allow a modeler to encode a distribution not otherwise explicitly
handled by this specification. Because the range of possibilities is so vast, the modeler should not normally expect
any given SBML simulator or other software to be able to properly manipulate this distribution, but particular
software tools may implement support for certain distributions they know their own software's users may require.

The required attribute `definitionURL`, of type `ExternalRef`, must be a URI that defines a valid distribution. It is
strongly recommended that modelers use distributions from ProbOnto (`http://probonto.org/`), as consistently
referencing a single ontology will improve exchangeability, at least slightly. The referenced distribution is then
the distribution defined by this **ExternalDistribution**, along with any parameterization provided by the children
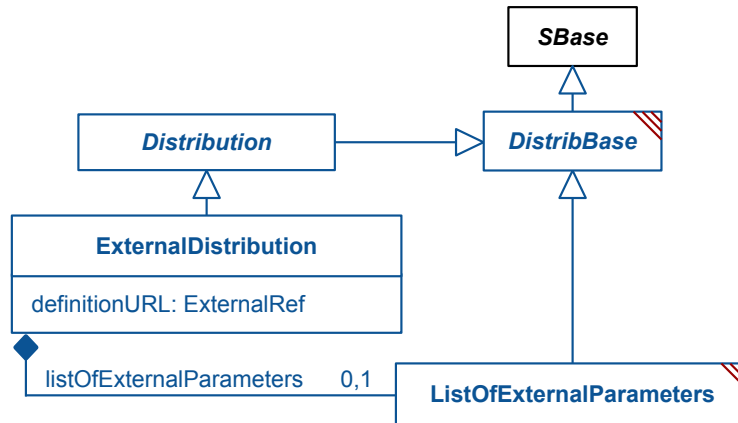
Figure 9: *The definition of the **ExternalDistribution**, **ExternalParameter**, and **ListOfExternalParameters** classes. These classes define a way to define a distribution with reference to an external database or ontology of distribution definitions.*

ExternalParameter elements.

Some referenced distributions are multivarite, meaning they define correlated distributions for two or more parameters. It is impossible with SBML Level 3 Core to define a **FunctionDefinition** that returns a vector, and similarly no `SId` in SBML Level 3 Core can be used to represent a vector. If this is desired, then, the Arrays package must be used in concert with the **ExternalDistribution** to cooperatively set up a model with a **FunctionDefinition** that can use an array as input and/or as output.

The **ExternalDistribution** defines an optional child **ListOfExternalParameters**, which can be used to parameterize the defined distribution.

A.7 Specific Distributions

In this table, all distributions are listed, along with their types (Continuous, Categorical, or Discrete), whether they're univariate or multivariate, and a brief description. The element name is the name of the distribution with spaces removed, the initial letter lower-cased, and "`Distribution`" appended, so, for example, the "`Exponential`" distribution becomes "`<exponentialDistribution>`", and the "`Student T`" distribution becomes "`<studentTDistribution>`".

All of these distributions inherit from the abstract **Distribution** class. Additionally, the appropriate distributions inherit from the **UnivariateDistribution** or **MultivariateDistribution** abstract classes, and further from the **ContinuousUnivariateDistribution**, **DiscreteUnivariateDistribution**, or **CategoricalUnivariateDistribution** classes, which are related to one another as one would expect.

All descriptions are based on the information from `http://www.uncertml.org/`, which is now defunct, but which can still be accessed at `http://web.archive.org/web/20160313012501/uncertml.org`.

Distributions are listed grouped by category (type and univarite/multivariate), and alphabetical within those categories.

A.8 The BetaDistribution class

The **BetaDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `alpha` (α) and `beta` (β). Both `alpha` and `beta` must be positive.

A random variable x is Beta distributed if the probability density function (pdf) is of the form:

$$\frac{1}{B(\alpha,\beta)} x^{\alpha-1} (1-x)^{\beta-1}, \text{ where } B(\alpha,\beta) = \frac{\Gamma(\alpha)\Gamma(\beta)}{\Gamma(\alpha+\beta)}$$

The distribution is usually denoted as $x \sim Be(\alpha, \beta)$ with parameters α and β, both positive real values. As the domain

of the random variable is defined to be $[0, 1]$ the Beta distribution is normally used to describe the distribution of a probability value.

A.9 The CauchyDistribution class

The **CauchyDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `location` (θ) and `scale` (γ). The `scale` value must be positive.

A random variable x follows a Cauchy distribution if the probability density function (pdf) is of the form:

$$\frac{1}{\pi\gamma}\left[1+\left(\frac{x-\theta}{\gamma}\right)^2\right]^{-1}$$

The Cauchy distribution is equivalent to a Student-T distribution with 1 degree of freedom. It is widely used in physics, optics and astronomy. It is also known as the Lorenz or the Breit-Wigner distribution.

A.10 The ChiSquareDistribution class

The **ChiSquareDistribution** is a **ContinuousUnivariateDistribution** defining a **UncertValue** child `degreesOfFreedom` (ν). The `degreesOfFreedom` must be a positive integer.

A random variable x is Chi-square distributed if the probability density function (pdf) is of the form:

$$\frac{1}{\Gamma(\nu/2)2^{\nu/2}}x^{\nu/2-1}exp(-x/2)$$

The distribution is usually denoted as $x \sim \chi_\nu$ where ν is known as the degrees of freedom parameter. ν has to be positive and x has to be non-negative for the density to be defined. The Chi-square distribution is a special case of the Gamma distribution where $\chi \sim \Gamma(k = \nu/2, \theta = 2)$.

A.11 The ExponentialDistribution class

The **ExponentialDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** child `rate` (λ). The `rate` value must be positive.

A random variable x follows an exponential distribution if the probability density function (pdf) is of the form:

$$\lambda e^{-\lambda x}$$

It is often represented as $x \sim \text{Exp}(\lambda)$. It is used to model the time between events for a Poisson process and is used in simulation of stochastic systems.

A.12 The FDistribution class

The **FDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `numerator` (ν_1) and `denominator` (ν_2). Both `numerator` and `denominator` must be positive integers.

A random variable x follows an F distribution if the probability density function (pdf) is of the form:

$$\frac{1}{B(\nu_1/2,\nu_2/2)}\left(\frac{\nu_1}{\nu_2}\right)^{\nu_1/2}x^{\nu_1/2-1}\left(1+\frac{\nu_1}{\nu_2}x\right)^{-\frac{\nu_1+\nu_2}{2}}$$

where $B(.)$ is the Beta function. It often arises as the ratio of two random variables that are identically Chi-Square distributed.

A.13 The GammaDistribution class

The **GammaDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `shape` (k) and `scale` (θ). Both `shape` and `scale` must be positive.

A random variable x is Gamma distributed if the probability density function (pdf) is of the form:

$$\frac{1}{\Gamma(k)\theta^k}x^{k-1}exp(-x/\theta) \text{ with } \Gamma(\cdot) \text{ the Gamma function.}$$

The distribution is usually denoted as $x \sim Gamma(k, \theta)$ where k is known as the shape parameter and θ the scale parameter. Both parameters have be positive and x has to be non-negative for the density to be defined. In practice the Gamma distribution is often use to model the distribution of non-negative quantities such as variances.

A.14 The InverseGammaDistribution class

The **InverseGammaDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `shape` (α) and `scale` (β). Both `alpha` and `beta` must be positive.

A random variable x is Inverse Gamma distributed if the probability density function (pdf) is of the form:

$$\frac{\beta^{\alpha}}{\Gamma(\alpha)} x^{-\alpha-1} exp(-\beta/x)$$

If variable x is Inverse Gamma distributed, $1/x$ is gamma distributed. The Inverse Gamma distribution function can be obtained from the Gamma distribution by a transformation of variables.

A.15 The LaPlaceDistribution class

The **LaPlaceDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `location` (μ) and `scale` (b). The `scale` value must be positive.

A random variable x is Laplace distributed if the probability density function (pdf) is of the form:

$$\frac{1}{2b} exp(-\frac{abs(x-\mu)}{b})$$

where abs denotes the absolute value. It can be thought of as a combination of two exponential distributions.

A.16 The LogNormalDistribution class

The **LogNormalDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `shape` (σ^2) and `logScale` (μ). The `shape` value must be positive.

A random variable x is Log Normal distributed if the probability density function (pdf) is of the form:

$$\frac{1}{x\sqrt{2\pi\sigma^2}} exp(-\frac{(\ln(x)-\mu)^2}{2\sigma^2})$$

If variable x is normally distributed, exp(x) is Log Normal distributed. The Log Normal distribution function can be obtained from the normal distribution by a transformation of variables. It is often used for variables that must be positive.

A.17 The LogisticDistribution class

The **LogisticDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `location` (μ) and `scale` (s). The `scale` value must be positive.

A random variable x is Logistic distributed if the probability density function (pdf) is of the form:

$$\frac{exp(-(x-\mu)/s)}{s(1+exp(-(x-\mu)/s))^2}$$

A.18 The NormalDistribution class

The **NormalDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `mean` (μ), `stddev` (σ), and `variance` (σ^2). The distribution must either define a `stddev` or a `variance`, but not both. The `variance`, if defined, must be positive.

A random variable x is normally distributed if the probability density function (pdf) is of the form:

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

The distribution is usually denoted as $x \sim \mathcal{N}(\mu, \sigma^2)$ where μ is known as the mean parameter and σ^2 the variance parameter. If the random variable x is a vector of length greater than one, the normal distribution can be generalised

to the Multivariate normal. A reason for the widespread usage of the normal distribution is the Central limit theorem which states that the distribution of the mean of a large number of independent identically distributed random variables tends to a normal distributions as the number of random variables increases.

A.19 The ParetoDistribution class

The **ParetoDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `scale` (x_m) and `shape` (α). Both `shape` and `scale` must be positive.

A random variable x follows a Pareto distribution if the probability density function is of the form:

$$\frac{\alpha x_m^\alpha}{x^{\alpha+1}}$$

The distribution allows for the specification of a minimum value below which the density is 0. It is a skewed heavy-tailed distribution.

A.20 The RayleighDistribution class

The **RayleighDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `scale`.

[From Wikipedia:] A Rayleigh distribution is often observed when the overall magnitude of a vector is related to its directional components. One example where the Rayleigh distribution naturally arises is when wind velocity is analyzed into its orthogonal 2-dimensional vector components. Assuming that each component is uncorrelated, normally distributed with equal variance, and zero mean, then the overall wind speed (vector magnitude) will be characterized by a Rayleigh distribution. A second example of the distribution arises in the case of random complex numbers whose real and imaginary components are independently and identically distributed Gaussian with equal variance and zero mean. In that case, the absolute value of the complex number is Rayleigh-distributed.

A.21 The StudentTDistribution class

The **StudentTDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `location` (μ), `scale` (σ^2) and `degreesOfFreedom` (ν).

A random variable x follows a Student-t distribution if the probability density function (pdf) is of the form:

$\frac{\Gamma(\nu/2+1/2)}{\Gamma(\nu/2)(\pi\nu\sigma^2)^{1/2}}[1 + \frac{(x-\mu)^2}{\nu\sigma^2}]^{-\nu/2-1/2}$. The distribution is usually denoted as $x \sim St(\mu, \lambda, \nu)$

This distribution corresponds to integrating out the variance of a normal distribution using a inverse Gamma prior. It can therefore be interpreted as an infinite mixture of normal distributions having the same mean but different variances. The three parameters are the mean (μ), degrees of freedom (ν) and variance (σ^2). Setting the variance to 1 and the mean to 0 we obtain the Student-t form found in standard statistics references such as Wikipedia. Setting the d.f. to 1 the Cauchy distribution is obtained. Setting the d.f. to infinity the normal distribution is obtained. The student-t distribution is commonly used in likelihood inference as the maximum likelihood parameter estimates are more robust to outlier observations compared to the normal distribution.

A.22 The UniformDistribution class

The **UniformDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `minimum` (a), `maximum` (b) and the optional `numberOfClasses`. The `minimum` value must be less than the `maximum` value. If `numberOfClasses` is defined, its value must be an integer greater than or equal to two.

A random variable x follows a uniform distribution if the probability density function (pdf) is of the form:

$$\frac{1}{b-a}$$

The distribution assigns equal probability to all events within the chosen domain between (and including) the minimum (a) and the maximum (b).

If `numberOfClasses` is included, the uniform range is divided into $numberOfClasses - 1$ sections, and each of the

borders of those sections are equally likely to be returned. If `numberOfClasses` is 2 (the minimum), the range just has $2 - 1 = 1$ section, and the borders of that section (the `minimum` and `maximum`) are the two possible return values. If `numberofClasses` is 3, the range is broken into $3 - 1 = 2$ sections, leaving the `minimum`, `maximum`, and mean as the three possible return values, etc.

A.23 The WeibullDistribution class

The **WeibullDistribution** is a **ContinuousUnivariateDistribution** that defines the **UncertValue** children `shape` (k) and `scale` (λ). Both `shape` and `scale` must be positive.

A random variable x follows an Weibull distribution if the probability density function (pdf) is of the form:

$$\frac{k}{\lambda}\left(\frac{x}{\lambda}\right)^{k-1} exp(-x/\lambda)^k$$

It includes the exponential distribution as a special case. It is often used in engineering and finance.

A.24 The BinomialDistribution class

The **BinomialDistribution** is a **DiscreteUnivariateDistribution** that defines the **UncertValue** children `numberOfTrials` (n) and `probabilityOfSuccess` (θ). The `numberOfTrials` must be a positive integer, and `probabilityOfSuccess` must be a value between zero and one, inclusive.

A random variable x follows a Binomial distribution if the probability mass function (pmf) is of the form:

$$\binom{n}{x}\theta^x(1-\theta)^{n-x}$$

where $\binom{n}{x}$ denotes n choose x. The distribution is usually denoted as $x \sim b(n,\theta)$. The distribution describes the probability of getting x successes in n trials of independent experiments that have the same probability of success.

A.25 The GeometricDistribution class

The **GeometricDistribution** is a **DiscreteUnivariateDistribution** that defines the **UncertValue** child `probability` (p). The `probability` must have a value must be between zero and one, inclusive.

A random variable x follows a geometric distribution if the probability mass function (pmf) is of the form:

$$(1-p)^{x-1}p$$

It is often represented as $x \sim \text{Geom}(p)$. It is the discrete analogue of the exponential distribution. It is used to model distribution of the number of binary (Bernoulli) trials needed to get one success, with parameter, probability p.

A.26 The HypergeometricDistribution class

The **HypergeometricDistribution** is a **DiscreteUnivariateDistribution** that defines the **UncertValue** children `numberOfSuccesses` (m). `numberOfTrials` (n), and `populationSize` (N). All three values must be positive integers, chosen such that `numberOfTrials` is less than or equal to `populationSize`.

A random variable x follows a hypergeometric distribution if the probability mass function (pmf) is of the form:

$$\frac{\binom{m}{k}\binom{N-m}{n-k}}{\binom{N}{n}}$$

probability of getting x successes. It describes the number of successes in a sequence of draws without replacement.

A.27 The NegativeBinomialDistribution class

The **NegativeBinomialDistribution** is a **DiscreteUnivariateDistribution**. It has two defined **UncertValue** children `numberOfFailures` (r) and `probability` (p). The `numberOfFailures` must be a positive integer, and `probability` must have a value between zero and one, inclusive.

A random variable x follows a Negative Binomial distribution if the probability mass function (pmf) is of the form:

$$\binom{x+r-1}{x} p^x (1-p)^r$$

The distribution describes the probability of getting x successes in trials of independent experiments that have the same probability of success, and are run until we observe r failures. Note that some systems formulate this distribution differently: observing k failures before obtaining the r^{th} success. The formulation above follows the English version of Wikipedia; the alternate formulation is used on other language Wikipedia definitions of the distribution, as well as various software packages like Matlab and R.

? Lucian: NOTE! The above formulation was used by UncertML and Wikipedia, the sort-of-default distribution definition source for the annotation scheme. However, once people actually start implementing it, they may find that their software package uses the alternative. The ProbOnto 2.5 specification (`https://sites.google.com/site/probonto/download`) goes into great detail on this issue in Appendix A.3, for anyone who wants to know more. I would be happy to change the definition to match people's software, if need be.

A.28 The PoissonDistribution class

The **PoissonDistribution** is a **DiscreteUnivariateDistribution** that defines the **UncertValue** child `rate` (λ). The `rate` value must be positive.

A random variable x follows a Poisson distribution if the probability mass function (pmf) is of the form:

$$\frac{\lambda^x}{x!} \exp(-\lambda)$$

The Poisson distribution can be used to model the number of events occurring within fixed time period of time.

A.29 The BernoulliDistribution class

The **BernoulliDistribution** is a **CategoricalUnivariateDistribution** that defines the **UncertValue** child `prob` (μ). The `prob` must have a value between zero and one, inclusive. It defines the probability that $x = 1$.

A random variable x follows a Bernoulli distribution if the probability mass function (pmf) is of the form:

$$\mu^x (1-\mu)^{1-x}$$

It describes the distribution of a single binary variable x.

A.30 The CategoricalDistribution class

The **CategoricalDistribution** is a **CategoricalUnivariateDistribution** that contains one or more **Category** elements, each of which defines **UncertValue** `value` and `probability` children associated with that category. In order to be valid, the sum of the probabilities over all categories must either equal 1.0, or there must be exactly one **Category** without a child **UncertValue** `probability`, which is then set to $1.0 - sum(other\, probabilities)$. (In this case, that sum must be between 0.0 and 1.0, inclusive.)

A Categorical distribution is a generalisation of the Bernoulli distribution to K discrete outcomes, giving the K probabilities p_i, $i = 1, .., K$ for each outcome. There is no ordering in the K outcomes.

The optional `rank` attribute, if present, is provided as a way to differentiate between an ordered vs. unordered categorical distribution. It does not affect the sampling of the distribution in any way, and is provided for reference only. The `rank` attributes, if present, must be unique among the **Category** elements of a single **CategoricalDistribution**, and must begin with "`0`". Thus, if one **Category** with a `rank` is present, the value of its `rank` must be "`0`"; if there are two, they must be "`0`" and "`1`", etc.

A.31 The ListOfCategories class

The **ListOfCategories** class, like other **ListOf**___ classes in SBML Level 3 Core, is a container for one or more **Category** objects. Unlike many of **ListOf**___ classes in SBML Level 3 Core, at least one child **Category** is required, because the behavior of the parent distribution would be undefined if it had no child **Category** objects from which to choose.

A.32 The Category class

The **Category** class has a required **UncertValue** child `value`, and an optional **UncertValue** child `probability`. In any **CategoricalDistribution**, only one child **Category** may have an undefined `probability`; the rest must be defined and their totals add up to less than one. If all **Category** children have defined `probability` children, the total of all of those probabilities must add up to exactly one.

Each **Category** defines a `value`, and that value's `probability` of being sampled from that distribution. If the `probability` is not explicitly defined, it is implicitly defined as one minus the sum of the probabilities of all the other **Category** objects in the same **CategoricalDistribution**.

A.32.1 Attributes inherited from SBase

A **Category** always inherits the optional `metaid` and `sboTerm` attributes, and inherits optional `id` and `name` attributes as described in Section 3.9 on page 19. The `id` of a **Category** has no mathematical meaning.

B Validation of SBML documents

B.1 Validation and consistency rules

This section summarizes all the conditions that must (or in some cases, at least *should*) be true of an SBML Level 3 Version 2 model that uses the Distributions package. We use the same conventions as are used in the SBML Level 3 Version 2 Core specification document. In particular, there are different degrees of rule strictness. Formally, the differences are expressed in the statement of a rule: either a rule states that a condition *must* be true, or a rule states that it *should* be true. Rules of the former kind are strict SBML validation rules—a model encoded in SBML must conform to all of them in order to be considered valid. Rules of the latter kind are consistency rules. To help highlight these differences, we use the following three symbols next to the rule numbers:

- ☑ A checked box indicates a *requirement* for SBML conformance. If a model does not follow this rule, it does not conform to the Distributions package specification. (Mnemonic intention behind the choice of symbol: "This must be checked.")

- ⚠ A triangle indicates a *recommendation* for model consistency. If a model does not follow this rule, it is not considered strictly invalid as far as the Distributions package specification is concerned; however, it indicates that the model contains a physical or conceptual inconsistency. (Mnemonic intention behind the choice of symbol: "This is a cause for warning.")

- ★ A star indicates a strong recommendation for good modeling practice. This rule is not strictly a matter of SBML encoding, but the recommendation comes from logical reasoning. As in the previous case, if a model does not follow this rule, it is not strictly considered an invalid SBML encoding. (Mnemonic intention behind the choice of symbol: "You're a star if you heed this.")

The validation rules listed in the following subsections are all stated or implied in the rest of this specification document. They are enumerated here for convenience. Unless explicitly stated, all validation rules concern objects and attributes specifically defined in the Distributions package package.

☞ For convenience and brevity, we use the shorthand "`distrib:x`" to stand for an attribute or element name `x` in the namespace for the Distributions package package, using the namespace prefix `distrib`. In reality, the prefix string may be different from the literal "`distrib`" used here (and indeed, it can be any valid XML namespace prefix that the modeler or software chooses). We use "`distrib:x`" because it is shorter than to write a full explanation everywhere we refer to an attribute or element in the Distributions package namespace.

Attributes from this package are listed in these rules as having the "`distrib:`" prefix, but as is convention for SBML packages, this prefix is optional.

General rules about this package

distrib-10101 ☑ To conform to the Distributions package specification for SBML Level 3 Version 2, an SBML document must declare "`http://www.sbml.org/sbml/level3/version2/distrib/version1`" as the XMLNamespace to use for elements of this package. (Reference: SBML Level 3 Package specification for Distributions, Version 1 Section 3.2 on page 12.)

distrib-10102 ☑ Wherever they appear in an SBML document, elements and attributes from the Distributions package must use the "`http://www.sbml.org/sbml/level3/version2/distrib/version1`" namespace, declaring so either explicitly or implicitly. (Reference: SBML Level 3 Package specification for Distributions, Version 1 Section 3.2 on page 12.)

General rules about identifiers

distrib-10301 ☑ (Extends validation rule #10301 in the SBML Level 3 Core specification. TO DO list scope of ids) (Reference: SBML Level 3 Version 1 Core, Section 3.1.7.)

distrib-10302 ☑ The value of a `distrib:id` must conform to the syntax of the **SBML** data type `SId` (Reference: SBML Level 3 Version 1 Core, Section 3.1.7.)

TODO: ANY LIST OF ELEMENTS THAT HAVE ATTRIBUTES

Rules for the extended SBML class

distrib-20101 ☑ In all SBML documents using the Distributions package, the **SBML** object must have the `distrib:required` attribute. (Reference: SBML Level 3 Version 1 Core, Section 4.1.2.)

distrib-20102 ☑ The value of attribute `distrib:required` on the **SBML** object must be of data type `boolean`. (Reference: SBML Level 3 Version 1 Core, Section 4.1.2.)

distrib-20103 ☑ The value of attribute `distrib:required` on the **SBML** object must be set to "`true`". (Reference: SBML Level 3 Package specification for Distributions, Version 1 Section 3.2 on page 12.)

Rules for extended SBase object

distrib-20301 ☑ A *SBase* object may have the optional attributes `distrib:id` and `distrib:name`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on a *SBase* object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.8 on page 18.)

distrib-20302 ☑ A *SBase* object may contain one and only one instance of the **Uncertainty** element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a *SBase* object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.8 on page 18.)

distrib-20303 ☑ The attribute `distrib:name` on a *SBase* must have a value of data type `string`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.8 on page 18.)

Rules for Distribution object

distrib-20601 ☑ A **Distribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **Distribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-20602 ☑ A **Distribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **Distribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

Rules for UnivariateDistribution object

distrib-20701 ☑ An **UnivariateDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-20702 ☑ An **UnivariateDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

Rules for MultivariateDistribution object

distrib-20801 ☑ A **MultivariateDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **MultivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-20802 ☑ A **MultivariateDistribution** object may have the optional SBML Level 3 Core subobjects for

notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **MultivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

Rules for ContinuousUnivariateDistribution object

distrib-20901 ☑ A **ContinuousUnivariateDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ContinuousUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-20902 ☑ A **ContinuousUnivariateDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **ContinuousUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-20903 ☑ A **ContinuousUnivariateDistribution** object may contain one and only one instance of each of the UncertBound and UncertBound elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **ContinuousUnivariateDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Appendix A.3 on page 39.)

Rules for DiscreteUnivariateDistribution object

distrib-21001 ☑ A **DiscreteUnivariateDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **DiscreteUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21002 ☑ A **DiscreteUnivariateDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **DiscreteUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21003 ☑ A **DiscreteUnivariateDistribution** object may contain one and only one instance of each of the UncertBound and UncertBound elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **DiscreteUnivariateDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.4 on page 39.)

Rules for CategoricalUnivariateDistribution object

distrib-21101 ☑ A **CategoricalUnivariateDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **CategoricalUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21102 ☑ A **CategoricalUnivariateDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **CategoricalUnivariateDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

Rules for UncertValue object

distrib-21201 ☑ An **UncertValue** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UncertValue**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21202 ☑ An **UncertValue** object may have the optional SBML Level 3 Core subobjects for notes and

annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UncertValue**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21203 ☑ An **UncertValue** object may have the optional attributes `distrib:value`, `distrib:var` and `distrib:units`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on an **UncertValue** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.10 on page 20.)

distrib-21204 ☑ The attribute `distrib:value` on an **UncertValue** must have a value of data type `double`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.10 on page 20.)

distrib-21205 ☑ The value of the attribute `distrib:var` of an **UncertValue** object must be the identifier of an existing *SBase* object defined in the enclosing **Model** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.10 on page 20.)

distrib-21206 ☑ The value of the attribute `distrib:units` on an **UncertValue** must have a taken from the following: the identifier of a **UnitDefinition** object in the enclosing **Model**, or one of the base units in SBML. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.10 on page 20.)

Rules for UncertBound object

distrib-21301 ☑ An **UncertBound** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UncertBound**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21302 ☑ An **UncertBound** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UncertBound**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21303 ☑ An **UncertBound** object must have the required attribute `distrib:inclusive`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on an **UncertBound** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.11 on page 20.)

distrib-21304 ☑ The attribute `distrib:inclusive` on an **UncertBound** must have a value of data type `boolean`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.11 on page 20.)

Rules for ExternalDistribution object

distrib-21401 ☑ An **ExternalDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **ExternalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21402 ☑ An **ExternalDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **ExternalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21403 ☑ An **ExternalDistribution** object must have the required attribute `distrib:definitionURL`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on an **ExternalDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.6 on page 43.)

distrib-21404 ☑ An **ExternalDistribution** object may contain one and only one instance of the **ListOfExternalParameters** element. No other elements from the SBML Level 3 Distributions namespaces are

permitted on an **ExternalDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.6 on page 43.)

distrib-21405 ☑ The attribute `distrib:definitionURL` on an **ExternalDistribution** must have a value of data type `string`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.6 on page 43.)

distrib-21406 ☑ Apart from the general notes and annotations subobjects permitted on all SBML objects, a **ListOfExternalParameters** container object may only contain **ExternalParameter** objects. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

distrib-21407 ☑ A **ListOfExternalParameters** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ListOfExternalParameters** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

Rules for ExternalParameter object

distrib-21501 ☑ An **ExternalParameter** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **ExternalParameter**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21502 ☑ An **ExternalParameter** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **ExternalParameter**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21503 ☑ An **ExternalParameter** object must have the required attribute `distrib:definitionURL`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on an **ExternalParameter** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.12 on page 21.)

distrib-21504 ☑ An **ExternalParameter** object may contain one and only one instance of the **ListOfExternalParameters** element. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **ExternalParameter** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.12 on page 21.)

distrib-21505 ☑ The attribute `distrib:definitionURL` on an **ExternalParameter** must have a value of data type `string`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.12 on page 21.)

distrib-21506 ☑ Apart from the general notes and annotations subobjects permitted on all SBML objects, a **ListOfExternalParameters** container object may only contain **ExternalParameter** objects. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

distrib-21507 ☑ A **ListOfExternalParameters** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ListOfExternalParameters** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

Rules for NormalDistribution object

distrib-21601 ☑ A **NormalDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **NormalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21602 ☑ A **NormalDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **NormalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21603 ☑ A **NormalDistribution** object must contain one and only one instance of the UncertValue element, and may contain one and only one instance of each of the UncertValue and UncertValue elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **NormalDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.18 on page 46.)

Rules for UniformDistribution object

distrib-21701 ☑ An **UniformDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UniformDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21702 ☑ An **UniformDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UniformDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21703 ☑ An **UniformDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements, and may contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **UniformDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.22 on page 47.)

Rules for CategoricalDistribution object

distrib-21801 ☑ A **CategoricalDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **CategoricalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21802 ☑ A **CategoricalDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **CategoricalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21803 ☑ A **CategoricalDistribution** object must contain one and only one instance of the **ListOfCategories** element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **CategoricalDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.30 on page 49.)

distrib-21804 ☑ The **ListOfCategories** subobject on a **CategoricalDistribution** object must not be empty. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.30 on page 49.)

distrib-21805 ☑ Apart from the general notes and annotations subobjects permitted on all SBML objects, a **ListOfCategories** container object may only contain **Category** objects. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.31 on page 49.)

distrib-21806 ☑ A **ListOfCategories** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ListOfCategories** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.31 on page 49.)

Rules for Category object

distrib-21901 ☑ A **Category** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **Category**.

(Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21902 ☑ A **Category** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **Category**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-21903 ☑ A **Category** object may have the optional attribute `distrib:rank`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on a **Category** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.32 on page 50.)

distrib-21904 ☑ A **Category** object must contain one and only one instance of the UncertValue element, and may contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **Category** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.32 on page 50.)

distrib-21905 ☑ The attribute `distrib:rank` on a **Category** must have a value of data type `integer`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.32 on page 50.)

Rules for BernoulliDistribution object

distrib-22001 ☑ A **BernoulliDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **BernoulliDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22002 ☑ A **BernoulliDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **BernoulliDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22003 ☑ A **BernoulliDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **BernoulliDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.29 on page 49.)

Rules for BetaDistribution object

distrib-22101 ☑ A **BetaDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **BetaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22102 ☑ A **BetaDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **BetaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22103 ☑ A **BetaDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **BetaDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.8 on page 44.)

Rules for BinomialDistribution object

distrib-22201 ☑ A **BinomialDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **BinomialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22202 ☑ A **BinomialDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **BinomialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22203 ☑ A **BinomialDistribution** object must contain one and only one instance of each of the Uncert-Value and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **BinomialDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.24 on page 48.)

Rules for CauchyDistribution object

distrib-22301 ☑ A **CauchyDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **CauchyDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22302 ☑ A **CauchyDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **CauchyDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22303 ☑ A **CauchyDistribution** object must contain one and only one instance of each of the Uncert-Value and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **CauchyDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.9 on page 45.)

Rules for ChiSquareDistribution object

distrib-22401 ☑ A **ChiSquareDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ChiSquareDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22402 ☑ A **ChiSquareDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **ChiSquareDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22403 ☑ A **ChiSquareDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **ChiSquareDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.10 on page 45.)

Rules for ExponentialDistribution object

distrib-22501 ☑ An **ExponentialDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **ExponentialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22502 ☑ An **ExponentialDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **ExponentialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22503 ☑ An **ExponentialDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **ExponentialDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.11 on page 45.)

Rules for FDistribution object

distrib-22601 ☑ A **FDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **FDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22602 ☑ A **FDistribution** object may have the optional SBML Level 3 Core subobjects for notes and

annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **FDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22603 ☑ A **FDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **FDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.12 on page 45.)

Rules for GammaDistribution object

distrib-22701 ☑ A **GammaDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **GammaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22702 ☑ A **GammaDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **GammaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22703 ☑ A **GammaDistribution** object must contain one and only one instance of each of the Uncert-Value and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **GammaDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.13 on page 45.)

Rules for GeometricDistribution object

distrib-22801 ☑ A **GeometricDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **GeometricDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22802 ☑ A **GeometricDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **GeometricDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22803 ☑ A **GeometricDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **GeometricDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.25 on page 48.)

Rules for HypergeometricDistribution object

distrib-22901 ☑ A **HypergeometricDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **HypergeometricDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22902 ☑ A **HypergeometricDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **HypergeometricDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-22903 ☑ A **HypergeometricDistribution** object must contain one and only one instance of each of the UncertValue, UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **HypergeometricDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.26 on page 48.)

Rules for InverseGammaDistribution object

distrib-23001 ☑ An **InverseGammaDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permit-

ted on an **InverseGammaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23002 ☑ An **InverseGammaDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **InverseGammaDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23003 ☑ An **InverseGammaDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **InverseGammaDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.14 on page 46.)

Rules for LaPlaceDistribution object

distrib-23101 ☑ A **LaPlaceDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **LaPlaceDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23102 ☑ A **LaPlaceDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **LaPlaceDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23103 ☑ A **LaPlaceDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **LaPlaceDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.15 on page 46.)

Rules for LogNormalDistribution object

distrib-23201 ☑ A **LogNormalDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **LogNormalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23202 ☑ A **LogNormalDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **LogNormalDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23203 ☑ A **LogNormalDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **LogNormalDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.16 on page 46.)

Rules for LogisticDistribution object

distrib-23301 ☑ A **LogisticDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **LogisticDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23302 ☑ A **LogisticDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **LogisticDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23303 ☑ A **LogisticDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **LogisticDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.17 on page 46.)

Rules for NegativeBinomialDistribution object

distrib-23401 ☑ A **NegativeBinomialDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **NegativeBinomialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23402 ☑ A **NegativeBinomialDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **NegativeBinomialDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23403 ☑ A **NegativeBinomialDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **NegativeBinomialDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.27 on page 48.)

Rules for ParetoDistribution object

distrib-23501 ☑ A **ParetoDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ParetoDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23502 ☑ A **ParetoDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **ParetoDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23503 ☑ A **ParetoDistribution** object must contain one and only one instance of each of the UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **ParetoDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.19 on page 47.)

Rules for PoissonDistribution object

distrib-23601 ☑ A **PoissonDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **PoissonDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23602 ☑ A **PoissonDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **PoissonDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23603 ☑ A **PoissonDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **PoissonDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.28 on page 49.)

Rules for RayleighDistribution object

distrib-23701 ☑ A **RayleighDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **RayleighDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23702 ☑ A **RayleighDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **RayleighDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23703 ☑ A **RayleighDistribution** object must contain one and only one instance of the UncertValue element. No other elements from the SBML Level 3 Distributions namespaces are permitted on a

RayleighDistribution object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.20 on page 47.)

Rules for StudentTDistribution object

distrib-23801 ☑ A **StudentTDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **StudentTDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23802 ☑ A **StudentTDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **StudentTDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23803 ☑ A **StudentTDistribution** object must contain one and only one instance of each of the Uncert-Value, UncertValue and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **StudentTDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.21 on page 47.)

Rules for WeibullDistribution object

distrib-23901 ☑ A **WeibullDistribution** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **WeibullDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23902 ☑ A **WeibullDistribution** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on a **WeibullDistribution**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-23903 ☑ A **WeibullDistribution** object must contain one and only one instance of each of the Uncert-Value and **UncertValue** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on a **WeibullDistribution** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section A.23 on page 48.)

Rules for Uncertainty object

distrib-24001 ☑ An **Uncertainty** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **Uncertainty**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24002 ☑ An **Uncertainty** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **Uncertainty**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24003 ☑ An **Uncertainty** object may contain one and only one instance of each of the UncertStatistics and Distribution elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **Uncertainty** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.13 on page 21.)

Rules for UncertStatistics object

distrib-24101 ☑ An **UncertStatistics** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UncertStatistics**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24102 ☑ An **UncertStatistics** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UncertStatistics**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24103 ☑ An **UncertStatistics** object may contain one and only one instance of each of the UncertValue, UncertValue, UncertValue, UncertValue, UncertValue, UncertValue, UncertValue, Uncert-Value, UncertStatisticSpan, UncertStatisticSpan, UncertStatisticSpan, UncertStatisticSpan and **ListOfExternalParameters** elements. No other elements from the SBML Level 3 Distributions namespaces are permitted on an **UncertStatistics** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.14 on page 22.)

distrib-24104 ☑ Apart from the general notes and annotations subobjects permitted on all SBML objects, a **ListOfExternalParameters** container object may only contain **ExternalParameter** objects. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

distrib-24105 ☑ A **ListOfExternalParameters** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on a **ListOfExternalParameters** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.18 on page 26.)

Rules for UncertStatisticSpan object

distrib-24201 ☑ An **UncertStatisticSpan** object may have the optional SBML Level 3 Core attributes `metaid` and `sboTerm`. No other attributes from the SBML Level 3 Core namespaces are permitted on an **UncertStatisticSpan**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24202 ☑ An **UncertStatisticSpan** object may have the optional SBML Level 3 Core subobjects for notes and annotations. No other elements from the SBML Level 3 Core namespaces are permitted on an **UncertStatisticSpan**. (Reference: SBML Level 3 Version 1 Core, Section 3.2.)

distrib-24203 ☑ An **UncertStatisticSpan** object may have the optional attributes `distrib:varLower`, `distrib:valueLower`, `distrib:varUpper` and `distrib:valueUpper`. No other attributes from the SBML Level 3 Distributions namespaces are permitted on an **UncertStatisticSpan** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.16 on page 24.)

distrib-24204 ☑ The value of the attribute `distrib:varLower` of an **UncertStatisticSpan** object must be the identifier of an existing **SBase** object defined in the enclosing **Model** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.16 on page 24.)

distrib-24205 ☑ The attribute `distrib:valueLower` on an **UncertStatisticSpan** must have a value of data type `double`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.16 on page 24.)

distrib-24206 ☑ The value of the attribute `distrib:varUpper` of an **UncertStatisticSpan** object must be the identifier of an existing **SBase** object defined in the enclosing **Model** object. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.16 on page 24.)

distrib-24207 ☑ The attribute `distrib:valueUpper` on an **UncertStatisticSpan** must have a value of data type `double`. (Reference: SBML Level 3 Package specification for Distributions, Version 1, Section 3.16 on page 24.)

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