« Shall we

## build ontologies regarding neuroscience facts or are human sentences sufficient ? »



### From knowledge to data : towards a formal representation of computational neuroscience known facts.

Models studied in computational neuroscience are fed up with facts obtained from the large literature reporting knowledge obtained from biological experimentation. Such knowledge is put in words, leading to phenomenological descriptions of the underlying behaviors. Then, a part of this information is put in equations and in distributed algorithms to check some quantitative or qualitative aspects of the presumed global understanding. Surprisingly enough, this work is mainly realized like by craftsmen, and more or less by intuition.

Other domains of knowledge have taken into account recent and powerful developments regarding semantic representations and introduce an intermediate framework between knowledge-in-words and knowledge-inequations: knowledge-in-facts. From very large and heterogeneous corpus like Wikipedia turned in DBpedia semantic data base, to very specialized fields of knowledge formalized as ontologies, it is clear that powerful methodological tools are now available to manage the data at a higher scale, beyond what "one" brain can handle manually, allowing to take larger corpus into account, but also to apply more rigorous methods on such semantic data.



https://team.inria.fr/mnemosyne

### Mnemosyme is modeling ``biological intelligence´´



### Using a systemic approach.



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### Modeling comp. Neuro. with ontologies ?

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Str	iatu	m								

From Wikipedia, the free encyclopedia

"Neostriatum" redirects here. For the avian brain region formerly called the neostriatum, see nidopallium.

The **striatum**, also known as the **neostriatum** or **striate nucleus**, is a subcortical (i.e., inside, rather than on the outside) part of the forebrain. It is the major input station of the basal ganglia system. The striatum, in turn, gets input from the cerebral cortex. In primates (including humans), the striatum is divided by a white matter tract called the internal capsule into two sectors called the caudate nucleus and putamen. The term **corpus striatum** occasionally refers to the striatum combined with the globus pallidus, a structure closely related to the putamen, and the lenticular nucleus refers to the putamen together with the globus pallidus.<sup>[1]</sup>

Functionally, the striatum helps coordinate motivation with body movement. It facilitates and balances motivation with both higher-level and lower-level functions, such as inhibiting one's behavior in a complex social interaction and fine-motor functions of inhibiting small voluntary movement.<sup>[2]</sup>



At the bottom J

V·T·E	Human brain, cerebrum, Interior of the cerebral hemispheres—Rostral Basal ganglia and associated structures (TA A14.1.09.321-552, GA 9.832-837)	[show]
V · T · E	Brain and spinal cord: neural tracts and fasciculi	[show]
V T E	Anxiety disorder: Obsessive-compulsive disorder (F42, 300.3)	[show]
Categories	: Cerebrum   Basal ganglia	

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V·T·EHuman brain, cerebrum, Interior of the cerebral hemispheres—Rostral Basal ganglia and associated structures (TA A14.1.09.321-552, GA 9.832-837)[hide]						[hide]	
	Grey matter		striatum: Putamen • Cauda	ate nucleus			
		Corpus striatum	<i>lentiform nucleus:</i> Putamen				
			Globus pallidus (GPe • GPi)				
		Ventral striatum	Nucleus accumbens • Olfact	ory tubercle • Islands of Calleja			
		Other	Amygdala • Claustrum				
Pacal ganglia	White matter	Semioval center					
basar yangna		Internal capsule (Ant	terior limb				
		Genu · Posterior limb · Optic radiation)					
		Corona radiata					
		External capsule • Extreme capsule					
		Pallidothalamic tracts: Thalamic fasciculus (Ansa lenticularis					
		Lenticular fasciculus	) • Subthalamic fasciculus				
Phinonconhalon	Grey matter	Anterior olfactory nu	cleus • Anterior perforated sul	ostance • Olfactory bulb			
Khinencephalon	White matter	Olfactory tract (Media	al olfactory stria • Lateral olfact	ory stria) • Olfactory trigone			
Other basel ferebrain	Grey matter	Substantia innominata (Basal optic nucleus of Meynert) • Nucleus of diagonal band					
Other basal forebrain	White matter	Diagonal band of Broca • Stria terminalis					
	Grey matter	Hippocampus proper: CA1 • CA2 • CA3 • CA4					
Archicortex:		Dentate gyrus: Fascia dentata					
Hippocampus anatomy		Subiculum					
	White matter	Alveus • Fimbria • Pe	rforant path • Schaffer collater	al			
M: CNS		anat (n/s/m/p/4/	/e/b/d/c/a/f/l/g)/phys/devp	noco (m/d/e/h/v/s)/cong/tumr, sysi/epon, injr	proc, drug (N1A/2AB/C/3/4/7A/B/	C/D)	

### At the bottom

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 Human brain, cerebrum, Interior of the cerebral hemispheres—Rostral Basal ganglia and associated structures (TA A14.1.09.321-552, GA 9.832-837)
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 [show]

 Categories: Cerebrum | Basal ganglia
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		1°: Pacinian cor	puscle/Meissner's corpuscle → Gracile fasciculus/Cuneate fasciculus → Gracile nucleus/Cuneate nucleus				
	PCML	$2^{\circ}: \rightarrow$ sensory decussation/arcuate fibers (Posterior external arcuate fibers, Internal arcuate fibers) $\rightarrow$ Medial lemniscus/Trigeminal lemniscus $\rightarrow$ Thalamus (VPL, VPM)					
Sensory/		3°: → Posterior limb of internal capsule → Postcentral gyrus					
ascending	Anterolateral/ pain	Fast/lateral	1° (Free nerve ending $\rightarrow$ A delta fiber) $\rightarrow$ 2° (Anterior white commissure $\rightarrow$ Lateral and Anterior Spinothalamic tract $\rightarrow$ Spinal lemniscus $\rightarrow$ VPL of Thalamus) $\rightarrow$ 3° (Postcentral gyrus) $\rightarrow$ 4° (Posterior parietal cortex)				
			2° (Spinotectal tract → Superior colliculus of Midbrain tectum)				
		Slow/medial	1° (Group C nerve fiber $\rightarrow$ Spinoreticular tract $\rightarrow$ Reticular formation) $\rightarrow$ 2° (MD of Thalamus) $\rightarrow$ 3° (Cingulate cortex)				
	Pyramidal	Ial flexion: Primary motor cortex → Posterior limb of internal capsule → Decussation of pyramids → Corticospinal tract (Lateral, Anterior) → Neuromuscular junction					
		flexion: Primary motor cortex → Genu of internal capsule → Corticobulbar tract → Facial motor nucleus → Facial muscles					
		flexion: Red nucleus → Rubrospinal tract					
	Extrapyramidal	<i>extension:</i> Vestibulocerebellum → Vestibular nuclei → <b>Vestibulospinal tract</b>					
Motor/		<i>extension:</i> Vestibulocerebellum → Reticular formation → <b>Reticulospinal tract</b>					
descending		Midbrain tectum → Tectospinal tract → muscles of neck					
		<i>direct</i> : 1° (Motor cortex → <b>Striatum</b> ) → 2° (GPi) → 3° (Lenticular fasciculus/Ansa lenticularis → Thalamic fasciculus → VL of Thalamus) → 4° (Thalamocortical radiations → Supplementary motor area) → 5° (Motor cortex)					
	Basal ganglia	<i>indirect:</i> 1° (Motor cortex $\rightarrow$ <b>Striatum</b> ) $\rightarrow$ 2° (GPe) $\rightarrow$ 3° (Subthalamic fasciculus $\rightarrow$ Subthalamic nucleus) $\rightarrow$ 4° (Subthalamic fasciculus $\rightarrow$ GPi) $\rightarrow$ 5° (Lenticular fasciculus/Ansa lenticularis $\rightarrow$ Thalamic fasciculus $\rightarrow$ VL of Thalamus) $\rightarrow$ 6° (Thalamocortical radiations $\rightarrow$ Supplementary motor area) $\rightarrow$ 7° (Motor cortex)					
		nigrostriatal pathway: Pars compacta → Striatum					
	Afferen	Vestibular nu	cleus → <b>Vestibulocerebellar tract</b> → ICP → Cerebellum → Granule cell				
		Pontine nuclei → <b>Pontocerebellar fibers</b> → MCP → Deep cerebellar nuclei → Granule cell					
		Inferior olivary nucleus → Olivocerebellar tract → ICP → Hemisphere → Purkinje cell → Deep cerebellar nuclei					
		Dentate nucleus in Lateral hemisphere/pontocerebellum $\rightarrow$ SCP $\rightarrow$ Dentatothalamic tract $\rightarrow$ Thalamus (VL) $\rightarrow$ Motor cortex					
Corobollar	Efferen	Interposed nucleus in Intermediate hemisphere/spinocerebellum $\rightarrow$ SCP $\rightarrow$ Reticular formation, or $\rightarrow$ Cerebellothalamic tract $\rightarrow$ Red nucleus $\rightarrow$ Thalamus (VL) $\rightarrow$ Motor cortex					
Cerebellar		Fastigial nucleus in Flocculonodular lobe/vestibulocerebellum → Vestibulocerebellar tract → Vestibular nucleus					
		lower limb → 1° (muscle spindles → DRG) → 2° (Posterior thoracic nucleus → Dorsal/posterior spinocerebellar tract → ICP Unc. prop.					
	Bidirectional	r	upper limb $\rightarrow$ 1° (muscle spindles $\rightarrow$ DRG) $\rightarrow$ 2° (Accessory cuneate nucleus $\rightarrow$ Cuneocerebellar tract $\rightarrow$ ICP $\rightarrow$ Anterior lobe of cerebellum)				
	эрпосегерена	Reflex arc	lower limb $\rightarrow$ 1° (Golgi tendon organ) $\rightarrow$ 2° (Ventral/anterior spinocerebellar tract $\rightarrow$ SCP $\rightarrow$ Cerebellar vermis)				

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 [show]

- This is no more knowledge but
- This is no more for

knowledge but data. human but algorithms.

- Words are defined :
  - By an identifier
  - As instance of classes (e.g. a brain-structure)
  - In relations to other words (e.g. connected to)
  - Though properties with values (e.g. a numeric value)
- Not all knowledge is reduced to data
  - But almost all

well-defined

anatomical facts : yes !

"Ganglion cells are output neurons of the retina, integrating visual information through a network of specific cells (amacrine, bipolar, horizontal). Their axons, corresponding to the optic nerve, carry the information outside the retina, without reciprocal connection (no feedback).

In the human eye, there are about 125 millions of photoreceptors (including more than 100 millions rods) and 1.5 millions of ganglion cells (GC), but the connectivity is uneven: some GC receive from less than 10 photoreceptors, others from thousands. More precisely, there are four kinds of ganglion cells: midget GC (80% of the GC) towards the parvocells in the thalamus; parasol GC (10% of the GC) towards the magnocells in the tha-

#### @RDF{

retinal-cell has-sub-class photo-receptor , horizontal-cell , bipolar-cell , amacrine-cell , ganglion-cell. photo-receptor has-sub-class rod , cone. visual-field human:horizontal-value 160deg. visual-field human:vertical-value 135deg. fovea-field human:value 3deg. fovea human:same-as macula. rod human:count 100e6. rod light-sensitivity low. rod color-sensitivity none. rod foveal-density negligible. cones human:count 25e6. cones light-sensitivity high. cones color-sensitivity high. cones foveal-density high. ganglion-cell has-sub-class midget-ganglion-cell parasol-ganglion-cell bistratified-ganglion-cell photosensitive-ganglion-cell. midget-ganglion-cell human:count 1.2e6. parasol-ganglion-cell human:count 1e5. bistratified-ganglion-cell human:count 1e5. photosensitive-ganglion-cell human:count low. midget-ganglion-cell is-a parvocellular-cell. midget-ganglion-cell projects-onto thalamus-LGN. parasol-ganglion-cell is-a magnocellular-cell. parasol-ganglion-cell projects-onto thalamus-LGN. bistratified-ganglion-cell projects-onto thalamus-LGN. bistratified-ganglion-cell projects-onto superior-colliculus. photosensitive-ganglion-cell projects-onto suprachiasmatic-nucleus.



isMemberOf

• This is no more sentences but facts.  $\frac{Syntax}{2} \rightarrow a.k.o.$  Semantic

- Clearly this must be a human task, not an meaningless algorithm.
  - Which information is pertinent ?
  - How should the data be structured ?
  - What is the implicit information here ?
    - e.g. which contextual information to add ?
  - What about knowledge put in graphics ?

### Using an ontology editor to manage/design the collected semantic data

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brainy-bot		
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© Retina-konio	eRDF { retinal-cell has-sub-class photo-receptor , horizontal-cell , bipolar-cell	
© Retina-magno	, amacrine-cell , ganglion-cell.	
© dSColliculus	visual-field human:horizontal-value 160deg.	
© FO-TRN	visual-field human:vertical-value 135deg.	
_C v1	fovea-field human:value 3deg. fovea human:same-as macula.	thalamus diencephalic i brain grey matter
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	cones color-sensitivity high.	
	cones foveal-density high.	regional part of
	bistratified-ganglion-cell photosensitive-ganglion-cell.	diencephalon
	midget-ganglion-cell human:count 1.2e6.	
	bistratified-ganglion-cell human:count 1e5.	
Lookup All Ontologies?	photosensitive-ganglion-cell human:count low.	
	midget-ganglion-cell is-a parvocellular-cell. midget-ganglion-cell projects-onto thalamus-LGN.	lateral nuclear group
	parasol-ganglion-cell is-a magnocellular-cell.	
	parasol-ganglion-cell projects-onto thalamus-LGN. bistratified-ganglion-cell projects-onto thalamus-LGN	a la
	bistratified-ganglion-cell projects-onto superior-colliculus.	
	photosensitive-ganglion-cell projects-onto suprachiasmatic-nucleus	pulvinar nucleus

What for ?

 Specify vocabulary and known facts : (qualitative data + quantitative values).

- Fix facts : not only "talk about" them, including
   Contradictory (either-or assumption)
   Fuzzy (fuzzy value, irreductible verbal info)
  - Unknown (wildcard value or fact)
- Data/Facts inference/representation
- Code parameterization

Graphical representation.

Automatic code input generation.

What for ?

Explicit the link between facts and model.

### ✓ Data inference/representation

- ✓ Offers new views on the collected knowledge,
- ✓ Allows modeling at the "conceptual" level,
- ✓ Checks biological statement coherence/consequence,
- ✓ Generate well-defined falsifiable assumptions.

### Code parameterization:

- ✓ guaranty the reproducibility of numerical experiments,
- ✓ avoid hidden implementation ad-hoc tricks,
- ✓ change the order of magnitude of the numerical exploration.

Modeling now reproducible !

### Modeling comp. Neuro. with ontologies ?

Let us start
 Looking at WikiPedia
 Annotating a PDF paper

Let us see
 What's already there
 What's missing here

So what & what for ?

On-to-lo-g... what ?
 Data as a graph
 Facts as triplets
 Logic as objects

A "turtoise" langage

What? The "fact" layer

A semantic description of some knowledge ? It is a

labeled directed multi-graph,

*i.e., a fully distributed set of information atoms.* 

✓ Each atom of information is a triplet



✓ the subject is the resource to describe,
✓ the predicate is the property type of the description,
✓ the object is the value of the property.

Example : The «sky has the color blue» writes : sky color blue.

What? The "fact" layer

subject verb object

**R.D.F.** Resource Description Format

Such a piece of information
 Defines meta-data over a document.
 Corresponds to text annotation.

✓ Each "token" is a

Uniform International Resource Identifier

✓ while object can also be a value or "literal"

(i.e., string, numerical value (number, date, ...))

Lesson 1.1: tokens.

✓ Each token is a  $<U_IRI>$ 

- ✓ It must have a universal identifier (no ambiguity)
- ✓ It may have a local denomination (usability)
- ✓ It includes international char (e.g.: español, français, …)

@let evs <http://ncicb.nci.nih.gov/xml/owl/EVS/Thesaurus.owl#>

@let prefix: <http://host/domain/>

yields `prefix:name` ↔ `<http://host/domain/name>`

#### Which striatum ?





(EVS) Entreprise Vocabulary System

Lesson 1.2: litterals.

### Strings are enclosed between double quotes "...",

- $\checkmark$  while '\" denotes a double-quote within a string,
- $\checkmark$  and ""..."" encloses multi-line strings.
- Have langage tag (e.g., @cl, @fr)

Numerical values have meta-data with

- ✓ an explicit physical-unit,
- bounds : minimal-value, maximal-value,
- ✓ a defaut-value,
- ✓ A precison epsilon-value (e.g. type)

Only ! Because:

Fuzzy value: (high|low|negligible|none)

**Contextual value** 

(e.g., in cat or human

- Compounded information is defined by a syntax
  - ✓ e.g.., a regular-expression,
  - but decomposes into atomic triplets

concave response

 $(x; k) \rightarrow \log(1 + kx)/k$ 

Lesson 1.3: phrases.

The construct subject a object with the verb "a" defines a subject type ✓ Token may have several types

The construct subject verb\_1 object\_1 ; verb\_2 object\_2 .
(i.e. ``subject verb\_1 object\_1 . subject verb\_2 object\_2 .``)
defines a property set, i.e. t-uple of information .

The construct

subject verb object\_1, object\_2. (i.e. ``subject verb\_1 object\_11 . subject verb\_1 object\_12 .``) subject verb (object\_1, object\_2). defines a set or collection (i.e. sequence) of values.

Lesson 1.4: reusing data.

The construct:

@use <URI>

allows to "import" data:

it states that the given turtoise document has to be merged to this one before considering these declarations.

This *document* <URI> is defined via the construct:

@let <> <URI>

And we are done !

What? The "fact" layer

✓ A well-founded atomic definition of a re-usable data token.

- Obvious to read/write from any software.
   Obvious to "tag" when reading a document.
   Recognized as a basic standard.
- Decidable algorithms of knowledge managements
   derive from "graph methods".
- A menagerie of tools to manipulate the data
   editing, data-basing functions, querying, re-formatting
- ✓ The "Turtle" syntax (here its "Turtoise" dialect) allows you, human, to efficiently read/write that.

What? The "logic" layer

✓ A class lattice :

✓ A class defines a concept, i.e. a category of individuals.

✓ Classes combine with operators.

✓ A property lattice :

✓ States relationships between individuals of a given class.

Relates an instance of a class to another instance.

Can have class or littreral as object.

✓ Individuals.

Ontologies are set of class + property
 They organise into "stores"

Is data coherent ?

Is assumption falsifiable ? Queries allow to

Retrieve initial data
 Derived new information

Lesson 2.1: Defining classes.

human a class. human sc mammalian.

✓ states that human is a class and a sub-class of mammalian.

- ✓ "thing"/"nothing" is the superclass/subclass of all classes
- classes form a lattice (nocycle, transitivity, …)
- class names are distinct from others namespaces
- ✓ litteral values are individuals of the `string`, `number` classes

# Lesson 2.2: Class operators.

Light logic inference

boy and male, kid. teen or boy, girl. bad not gentle.

Decidable logic inference

✓ Defines new classes by

intersection (conjunction), union (disjunction), complement.

boy ne girl.

✓ States that the two classes have no intersection.

my\_brothers are Bod, Carl, David.
 ✓ Defines a class by enumeration of individual.

# Lesson 2.3: Properties.

- A `property` states relationships between individuals of a given class.
- $\checkmark$  It relates an instance of a class to another instance of a class.
- ✓ It is declared using the `a property` verb/object.
- ✓ Properties form a lattice, using the `sp` (`sub-property of`) verb.
  - ✓ `domain` verb: class to which the property can be applied.
  - ✓ `range` verb: class that the property may have as value.
  - ✓ `min` verb: The minimal cardinality (i.e. at least N distinct values).
  - ✓ `max` verb: The maximal cardinality.
  - ✓ `some` verb: At least one value must belong to the given class.
  - ✓ `a symmetric` verb-object: The property is a symmetric.
  - ✓ `a transitive` verb-object: The property is a transitive.
  - ✓ `a functional` verb-object: The property has only one value.
  - ✓ `a unambiguous` verb-object: It defines a unique identifier.
  - ✓ `inverse` verb: The property is the inverse of another.
  - ✓ `has` verb: The property has the given value.

# Lesson 2.4: Property limits.

- ✓ Property names must be disjoint to other namespaces.
- ✓ Property range can have litteral values.
- Domain and range can be conjunction/disjunction of classes
- ✓ The `some` verb means
  - ✓ `some-values-from` if the the object is a class and
  - ✓ `has-value` if the object is an individual .

But:

 $\checkmark$  The `min` and `max` cardinality constraints are global.

For "light logic inference", no feature:
 `has`, `max` for transitive properties, `has-value`
 `min` and `max` cardinality beyond `0` or `1`

## Lesson 2.5: Individuals.

- Invididuals
  - $\checkmark$  Belongs to a class
  - Have property taking values
  - Can be set as `eq` or `ne` (equal or different)
- ✓ What's suprising here ?
  - Typing (i.e. defining what can be "done" with the individual)
     The type is a *property* and may have several value,
    - ✓ The notion of *class* captures the idea of category.

### ✓ Property ≠ category and

- ✓ It is not "attached" to a class but lives in a lattice,
- Has several values and . . domain (unless functional),

What? The "logic" layer

✓ Summarizing :

- An "integrated view" of concepts and properties,
   as modular as object-oriented description
   with a minimal formalism:
- A simple tool but with universal use.
   A stratified power of expression up to the 1st order logic,
   With inference and validation algorithms tuned to the used semantic,
  - ✓ With a direct "readable" mapping as a semantic graph.

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- Tagging and Folksonomy of publications
  - e.g. http://www.citeulike.org
- XML parameterization initiatives
  - http://brainml.org
  - http://www.neuroml.org
    - Used in VirtualRetina or neuralensemble.org
- + standardization of the terminology
- + clarification of specification (model choice, parameters)
- + interoperability between software

But not (yet) widely used.

- Wikipedia hierarchical anatomical data
  - Regions, Connections, Anatomical data.
- The Brain Architecture Management System
  - Brain Parts (gray matter regions, major fiber tracts, and ventricles),
  - Cell Types, Molecules,
  - Connections (between regions and cell types),
  - **Relations** (between parts identified different neuroanatomical atlases).
- The bio-ontology portail
  - Thesaurus (vocabulary)
  - Linked (clinical, physiologicla, ...) data
  - Query and inter-ontologies connections

Anatomy and clinical data

Vocabulary.

- Brain info :
  - http://braininfo.rprc.washington.edu
  - BrainInfo#NeuroNames
- Neuro lex :
  - http://neurolex.org/wiki/Main\_Page



- Using Semantic [. for .] an Ontology of Brain-Cortex Anatomy
  - Olivier Dameron et al Stanford, Inria Rennes, 200x
- Foundations for an ontology of brain areas, circuits and functions
  - Bénédicte M Batrancourt, CRIM Paris ; CNS'2013
- Cognitive Ontologies Mapping structure and function of the brain from a systemic view
  - Jaime Gomez, U. Polytechnica Madrid, 2008
- And also
  - Ontology of the ventricular system in relation with schizophrenia
    - Santos, Sallet, Cohn et al
  - A developmental ontology for the mammalian brain,
    - Puelles, Watson et al
  - Ontology for neuropsychology and cognitive psychology
    - Bond



# So what ?

✓ The finality:

✓ build systemic bio-plausible models
 ✓ for cognitive functions.

✓The goal:

✓ collect biological facts in a formal way

✓ so that :

Knowledge is well-defined,

✓ Algorithms can process them.

✓The tool:

 Ontology in order to describe knowledge to be put in models.

## So what ?

### what is the targeted knowledge ?

### Architecture:

✓ multi-scale brain structures and connections.

### Terminology:

✓ brain anatomical and functional element names.

### ✓ Data:

✓ quantitative and qualitative facts on each element.

✓ In a nutshell :

information put in a model and its simulation code.

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### But : Does the brain use and/or compute ontologies ?

Good news : this is not the question.

