



Open Science enabled by Text Mining

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www.nactem.ac.uk

School of Computer Science

The University of Manchester





Science Ouverte et Fouille de Textes

"Les mots nous cachent davantage les choses invisibles qu'ils ne nous révèlent les visibles" (Camus)

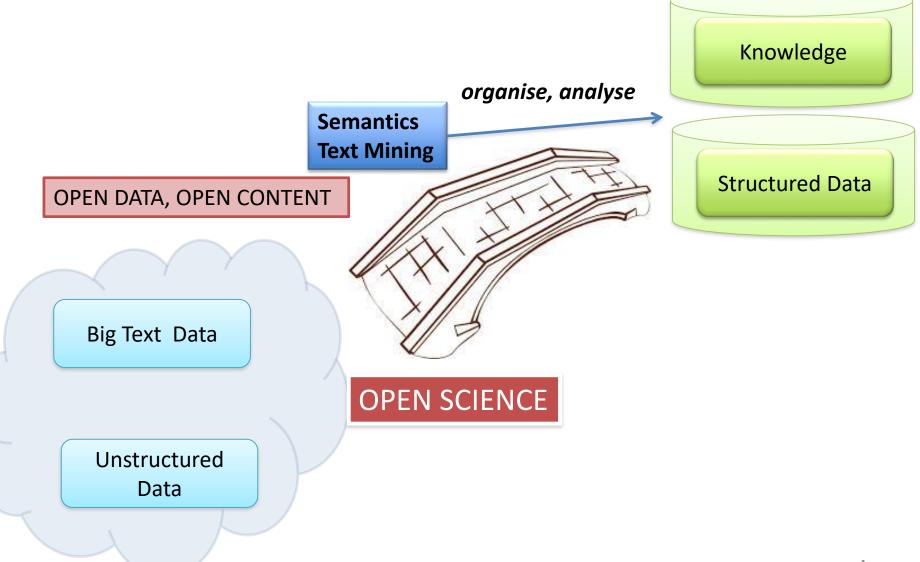
" [...] on a des instruments à portée de main qui, si correctement développés, donneront accès aux et la maîtrise des connaissances héritées des âges." (Préface à l'article de Vannevar Bush "As we may think", *The Atlantic*, juillet **1945**)

What hinders Open Science?

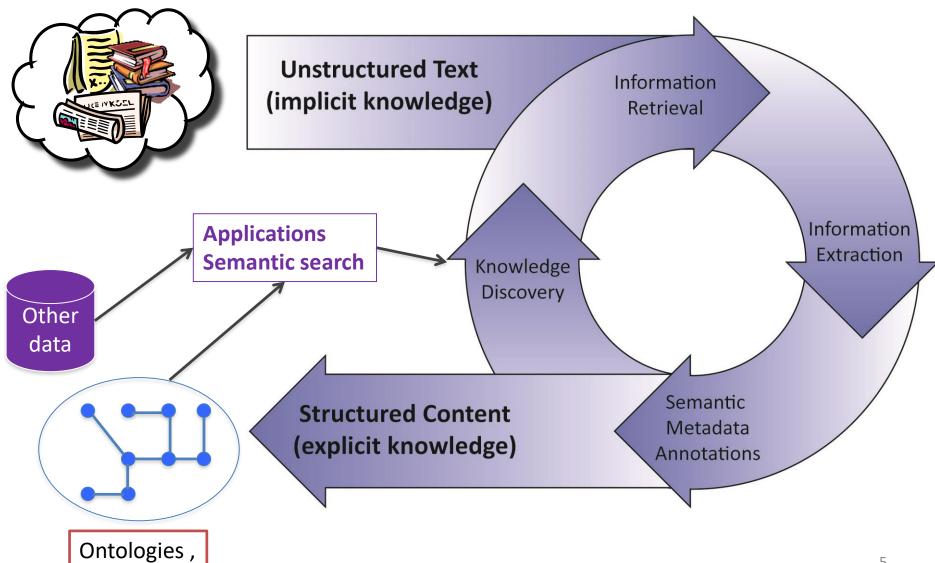
- Silos, fragmentations, literature grows daily
- We cannot read faster, we read more narrowly..
- Reductionist approaches, incomplete and uncertain information, complex problem space
- We cannot make **links** between disciplines

Scientific discovery by serendipity?

Big Textual Data

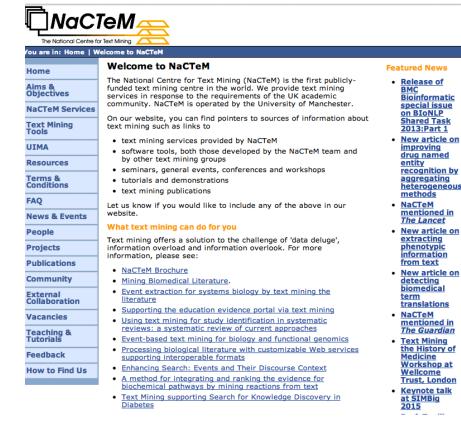


Text Mining



UK National Centre for Text Mining

- 1st publicly funded national text mining centre
- Location: Manchester Institute of Biotechnology
- Phase I Biology (2005-2008)
- Phase II Biology, Medicine, Social Sciences (2008-2011)
- Phase III Biology, Medicine, Health, Biodiversity, Humanities, Social Sciences; Fully sustainable centre (2011-)



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Text Mining for Open Science

Tools, Resources, Infrastructure, Services

- Tools for extracting concepts, relations, events, epistemic knowledge
- **Resources**: lexical repositories, annotated data stores
- Supporting semantic search, developing models, experiments, knowledge discovery, hypothesis generation
- Interoperable infrastructure

Communities

- Biology
- Medicine
- Digital Humanities
- Computational Social Science
- Biodiversity
- Chemistry
- Public Health
- Pharmacovigilance

Semantic search from PubMed

Thalia: Text Mining for Highlighting, Aggregating, and Linking Information in Articles

- Eight different types: chemicals, diseases, drugs, genes, metabolites, proteins, species and anatomical entities
- Covers entire PubMed
 - Over 26M abstracts
 - Automatic daily updates

Nactem-copious.man.ac.uk

Soto, Przybyla, Ananiadou 2018 Bioinformatics

Beyond keyword search

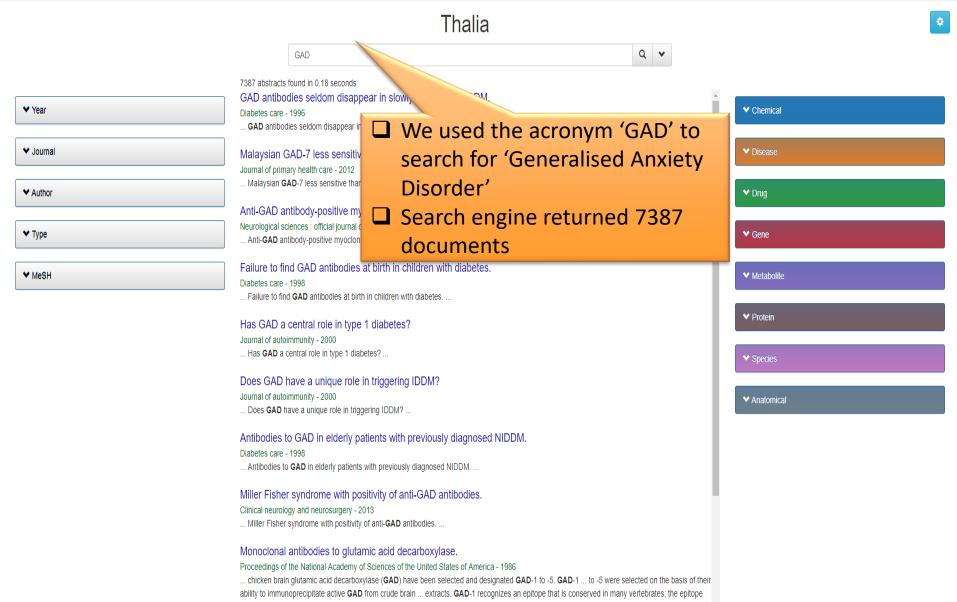
	Thalia	۵
	cat Q 🗸	
✓ Year	125679 abstracts found in 0.22 seconds Ovarian tissue culture in the presence of the stimulates follicle growth and steroidoge The Journal of endocrinology - 2017 + VEGFA165 groups, we observed a high level of total antibout the stimulates follicle growth and CAT	▲ Chemical
✓ Journal	Oxidative stress of imidaclothiz This search strategy returns 125,67	79 abstracts
✓ Author	Oxidative stress of imidaclothiz on eart Unnecessary workload	
♥ Туре	iTRAQ-based proteomic analysi Journal of proteomics - 2017 of action against Bacillus cereus.H, reactive oxygen species (ROS), the activities of catalase (CAT , resoluing non-downregulation of antioxidant proteins, especially CAT and SOD. BIOLOGICAL SIGNIFICANCE	
♥ MeSH	Mechanisms underlying toxicity and stimulatory role of single-walled carbon nanotubes in Hyoscyamus Journal of hazardous materials - 2017 of changes in the expression of various antioxidant enzymes including SOD, POD, CAT, and APX, and also	 ✓ Metabolite ✓ Protein
	Physiological and biochemical effects of nickel on rainbow trout (Oncorhynchus mykiss) tissues: Ass Chemosphere - 2017 of superoxide dismutase (SOD), catalase (CAT) enzyme activity and of lipid peroxidation (LPO), and glutathione in LPO indicating peroxidative damage and antioxidant enzymes SOD and CAT activity in tissues (p < 0.05), but 2 mg/Ni concentration caused a significant decrease in CAT activity in kidney tissues (p	✓ Species
	Chronic effects of triclocarban in the amphipod Gammarus locusta: Behavioural and biochemical impai Ecotoxicology and environmental safety - 2017 markers of oxidative stress (catalase (CAT), glutathione-s-transferase (GST) and lipid peroxidation	✓ Anatomical
	Molecular detection of severe fever with thrombocytopenia syndrome virus (SFTSV) in feral cats from Ticks and tick-borne diseases - 2017 Molecular detection of severe fever with thrombocytopenia syndrome virus (SFTSV) in feral cats from Seoul, Korea. This study tested serum samples of feral cats from a highly urbanized habitat, Seoul 126 samples tested, SFTSV was detected by RT-PCR in 22 (17.5%) cats from various sites of Seoul	

Search based on semantic types

	Thalia	\$
	Search Q 🔨	
Article meta-data	Named entities	
Year	Chemicals	
Journal	Diseases	
Author	Drugs	
Туре	Genes	
MeSH	Metabolites	
	Proteins CAT (UniProt:Q6IB77) x	
	Species	
	Anatomic	
Q Search		
	18457 abstracts found in 0.34 seconds Ovarian tissue culture in the presence	
✓ Year	The Journal of endocrinology - 2017 VEGFA165 groups, we observed a high level o	ns
✓ Journal	and lipid peroxidation levels and increased number Search results narrowed down to 18,457	
	Oxidative stress of imidaclothiz on eart Using faceted search , we reduced number of	
★ Author	Comparative biochemistry and physiology. Toxicol	
	Oxidative stress of imidaclothiz on earthworm E papers that we need to read by 85%	
♥ Туре	iTRAQ-based proteomic analysis of LI-F type peptides produced by Paenibacillus polymyxa JSa-9 mode	
	Journal of proteomics - 2017 de of action against Bacillus cereus.H, reactive oxygen species (ROS), the activities of catalase (CAT) and superoxide dismutase (SOD), and the	
✓ MeSH	relative expression of target genes by quantitative real-t impairment of cellular functions, resulting from downregulation of antioxidant proteins,	
	especially CAT and SOD. BIOLOGICAL SIGNIFICANCE: The mode of action of LI-F type antimicrobial peptides (AMP-jsa9)	
	Mechanisms underlying toxicity and stimulatory role of single-walled carbon nanotubes in Hyoscyamus	
	Journal of hazardous materials - 2017 s was a consequence of changes in the expression of various antioxidant enzymes including SOD, POD, CAT, and APX, and also biosynthesis of	18457 UniProt:Q8/877

proteins, phenolics, and specific metabolites such as proline. R ...

Disambiguation



recognized

Dealing with Acronyms



Dealing with Acronyms

	Thalia		
	GAD Q 🔺		
Article meta-da	ta Named entities		
Year	Chemicals		
Journal	Diseases generalized anxiety disorder (UMLS:C0175704)	X	
Author	Drugs		
Type MeSH	By disambiguating the acronym in our search strategy we reduced the number of articles from 7.287 to 1.674 (increased)		
Q Search	of articles from 7,387 to 1,674 (increased specificity)		
✔ Year	1674 abstracts found in 3.49 seconds Which medications should be prescribed to treat generalized anxiety disorder (GAD)?. Journal of psychiatry & neuroscience : JPN - 1999 Which medications should be prescribed to treat generalized anxiety disorder (GAD)? Which medications should be prescribed to treat generalized anxiety disorder (GAD)?	✓ Chemical	
Journal		▲ Disease	
	[Generalized anxiety disorders (GAD)a neglected illness? Background und aims of the GAD-P study]. Fortschritte der Medizin. Originalien - 2001	generalized anxiety disorder	1674 UMLS:C0175704
 Author 	[Generalized anxiety disorders (GAD)a neglected illness? Background und aims of the GAD-P study].In the past Generalized anxiety disorder (GAD)previously classified as anxiety neurosiswas regarded as not being a separate diagnostic entity. On the basis of new explicit criteria for GAD	GAD	1446 UMLS:C0270549
 Туре 	Delimiting the boundaries of generalized anxiety disorder: differentiating high worriers with and w	anxiety disorders	589 UMLS:C0003489
MeSH	Journal of anxiety disorders - 2002 with and without GAD.Investigations of the boundary between generalized anxiety disorder (GAD) and normal worry have relied primarily on comparisons of GAD-diagnosed individuals with non-anxious controls. One with GAD are unique to the disorder or are typical of severe worry more	panic disorder	293 UMLS:C0008111
	generally. The present	major depressive disorder	209 UMLS:C0339017
	Assessing generalized anxiety disorder in elderly people using the GAD-7 and GAD-2 scales: results	MDD	202

The American journal of geriatric psychiatry : official journal of the American Association for Geriatric Psychiatry - 2014

... Assessing generalized anxiety disorder in elderly people using the GAD-7 and GAD-2 scales: results ... -item Generalized Anxiety Disorder scale

UMLS:C1269683

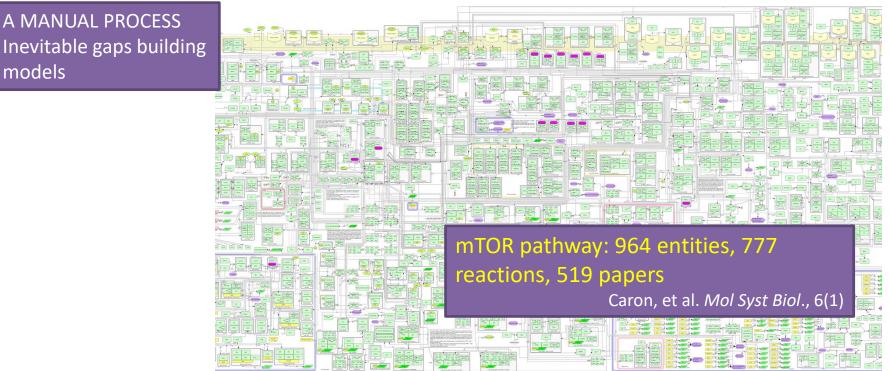
Thalia: document view

- Each named entity is color-coded
- Multiple-typed entities are visualised with a white rectangle
- Entities are linked to the corresponding URI in the ontology

Full text view ×
Low-Dose Farnesyltransferase Inhibitor Suppresses HIF-1α and Snail Expression in Triple-Negative Breast Cancer MDA-MB-231 Cells In Vitro. Journal of cellular physiology - 2017 Go to PubMed
The aggressiveness of triple-negative breast cancer (TNBC), which lacks estrogen receptor, progesterone receptor and epidermal growth factor receptor 2 and epidermal growth factor factor fills and recurrence of TNBC. Elevated hypoxia-induces aggressive phenotype are not completely understood in TNBC. The cytotoxic effects of farnesyltransferase (Files) inhibitors (FIIs) have been studied in cancer and leukemia cells in contrast, the effect of FIIs on HIF-1 α formation in human filt he low-dose FIIs did n and the prope
On Chemicals On Diseases On Drugs On Genes
On Metabolites On Proteins On Species On Anatomic

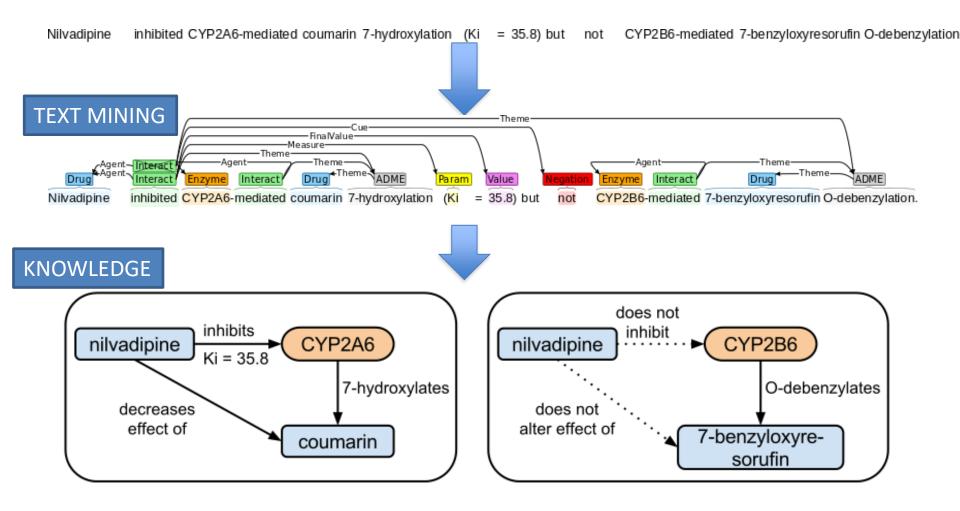
Understanding Mechanisms for Pathway Construction and Modelling

- Is it enough to mine just correlations from big data?
- Understanding not only of the components but the system



Ananiadou

Evidence-based science: linking text with knowledge

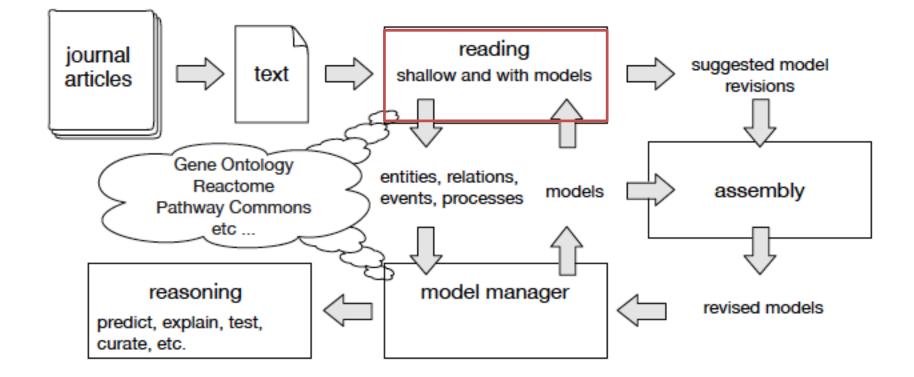


Linking Knowledge with Text *Curation*

Pathways

- Key to understanding biological systems
- Models need verification and maintenance (i.e., annotation/curation)
- Scale and speed of literature challenging
- Annotation/curation remains largely a manual task of incorporating knowledge from scientific publications

The Big Mechanism: reading, assembly, experiments



http://nactem.ac.uk/big_mechanism/

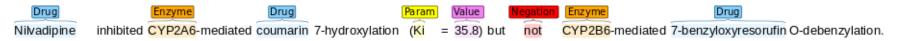




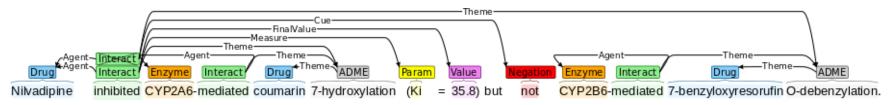
From concepts to events

Nilvadipine inhibited CYP2A6-mediated coumarin 7-hydroxylation (Ki = 35.8) but not CYP2B6-mediated 7-benzyloxyresorufin O-debenzylation.

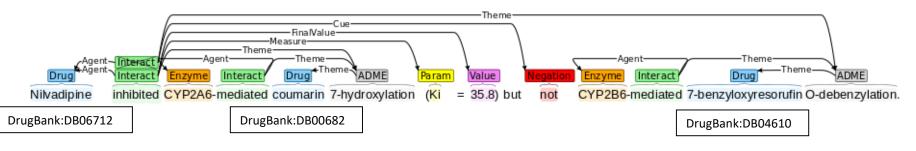
1 Concept recognition



2 Interaction recognition



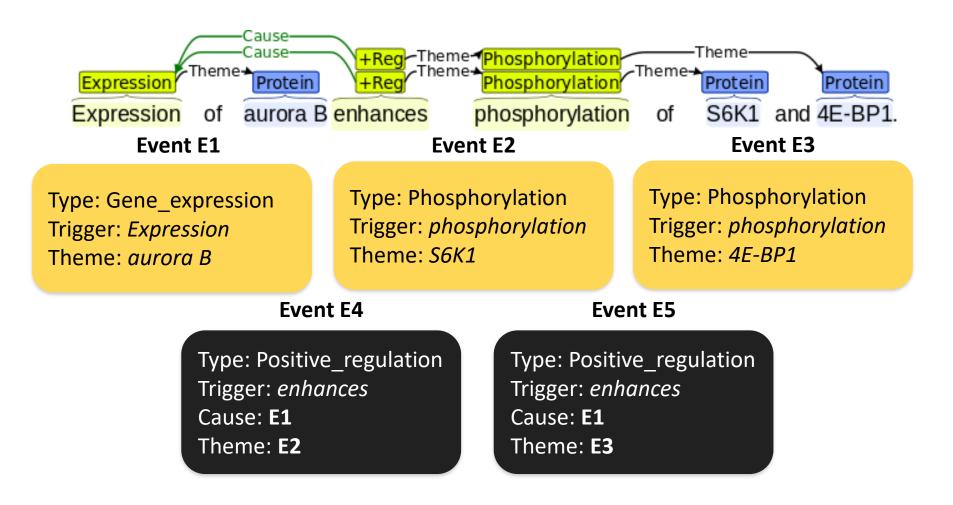
3 Concept and interaction identification







Event Extraction



Uncertainty Examples from Big Mechanism data

Therefore, AFP may interact with STAT3 in the signal pathway for chemotherapeutic efficiency of agents on AFPGC.

Therefore, LiCl **might** inhibit GSK3β in different ways

These data suggest that PI3K and β ARK1 form a macromolecular complex within the cell.

We then examined whether netrin-2 enhances the interaction between Cdo and Stim1.

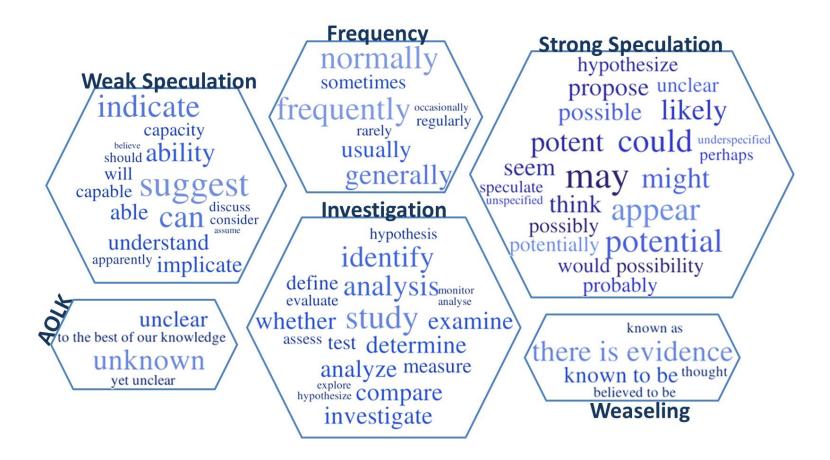
These results **indicate** that FLCN can interact directly with RagA via its GTPase domain.

Altogether, these results show that cobalt could affect both p53 and HIPK2 activity.

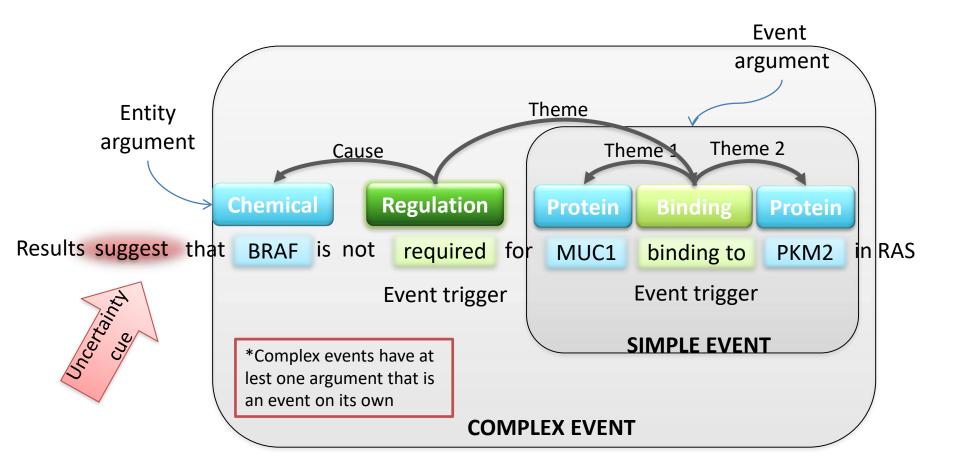
To **test if** endogenous hPGAM5 interacts with hPINK1, we first generated an anti-hPGAM5 antibody

We **hypothesize** that unphosphorylated cdr2 interacts with c-myc to prevent c-myc degradation

Uncertainty cues



Events and their interpretation



Uncertainty for model curation and experiments

- Uncertainty scoring as an expressive confidence measure
- Value for each event mentioned in a sentence
 - Consolidated uncertainty values from different papers
- Effort to decrease manual effort and select more certain events

Zerva, C., Batista-Navarro, R., Day, P. and S. Ananiadou (2017) Using uncertainty to link and rank evidence from biomedical literature for model reconstruction, *Bioinformatics*

LitPathExplorer: a confidence-based tool for exploring pathway models

- 1. Enabling flexible search and exploration of biomolecular pathway networks
 - different views of the data
 - various interactive functionalities
- 2. Provide a means for making existing evidence in the scientific literature available to support corroboration
- 3. Facilitate the discovery of new interactions that are not yet part of a given model
- 4. Allow the user to become an active participant of the analytical process

quantify confidence in the events

Video: http://nactem.ac.uk/LitPathExplorer_BI/LitPathExplorer.mp4

Search

LitPathExplor	er	×,	
Event			
MEK		Role	+
Complex (Ras	s/R	af/ MEK/GTP)	
KSR/14-3-3 ((din	ner)/MEK1-2-ad	tive
MEKK1/4			
MEK2			
Complex (Ras	s/G	TP/ Raf/MEK)	
Activated RAF	F1 (complex:MEK	
MEK1-2-activ	'e		
MEKK1			
MEKK1-5			
p-S218,222-	MEI	K1	
MEK1-2			
MEK			
MEK1			
MEK inhibitor	-		
Ras/Raf/MEK	/ER	ĸ	
MEKS271			
MEKi			
MEK-1			
DNEE-MEK1			

AND v		
Event		
Entity	Role	+-

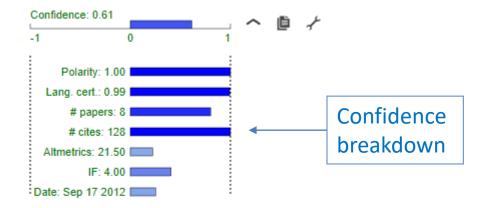
- A pathway model can be searched by providing:
 - event types,
 - entities,
 - and/or roles for each entity in the reaction
- Multiple queries can be combined in a Boolean search

Inspector, quantifying the confidence

Inspector

Node clicked: Event 🆓

Protein_catabolism (E5163)

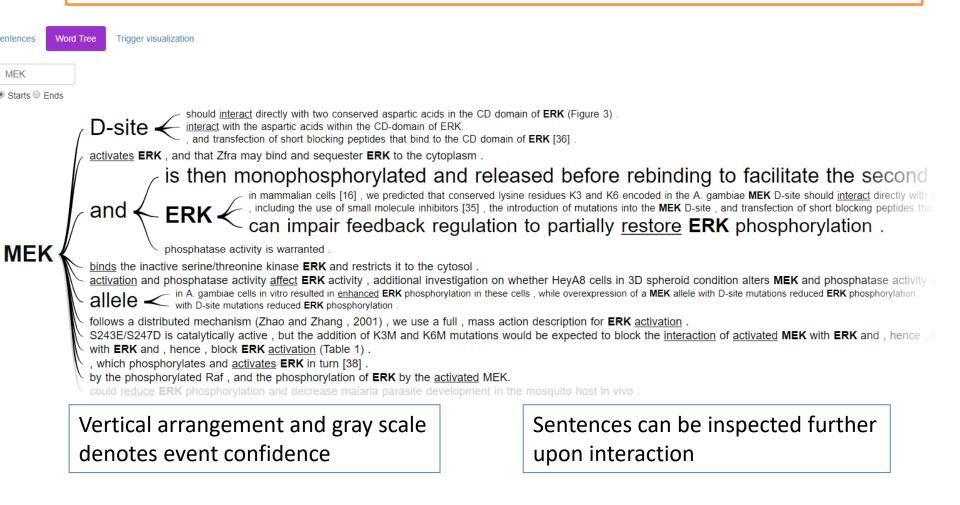


Entities related:

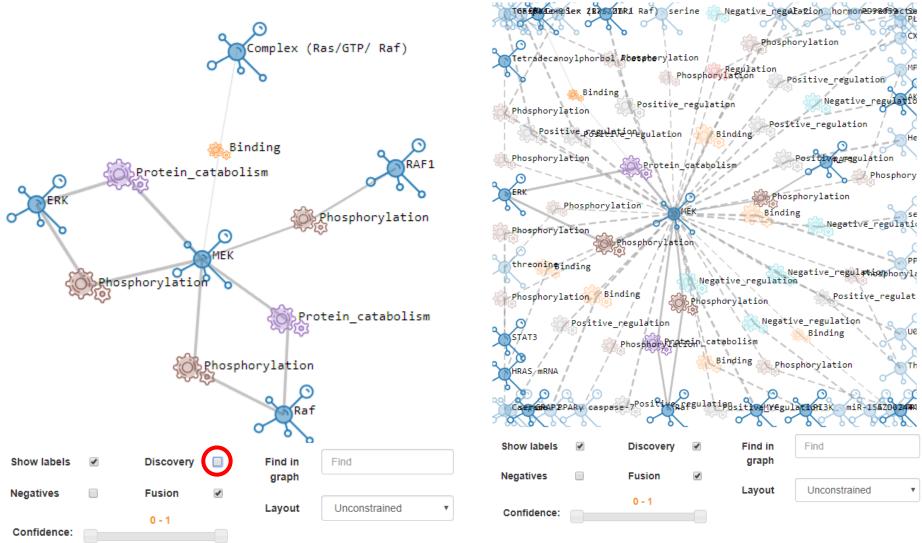
GeneOrProtein: ERK (PANTHER PATHWAY COMPONENT:P00371) Role: theme

GeneOrProtein: MEK (PANTHER PATHWAY COMPONENT:P00225) Role: agent

Word tree visualisation: Contrast event mentions across the corpus

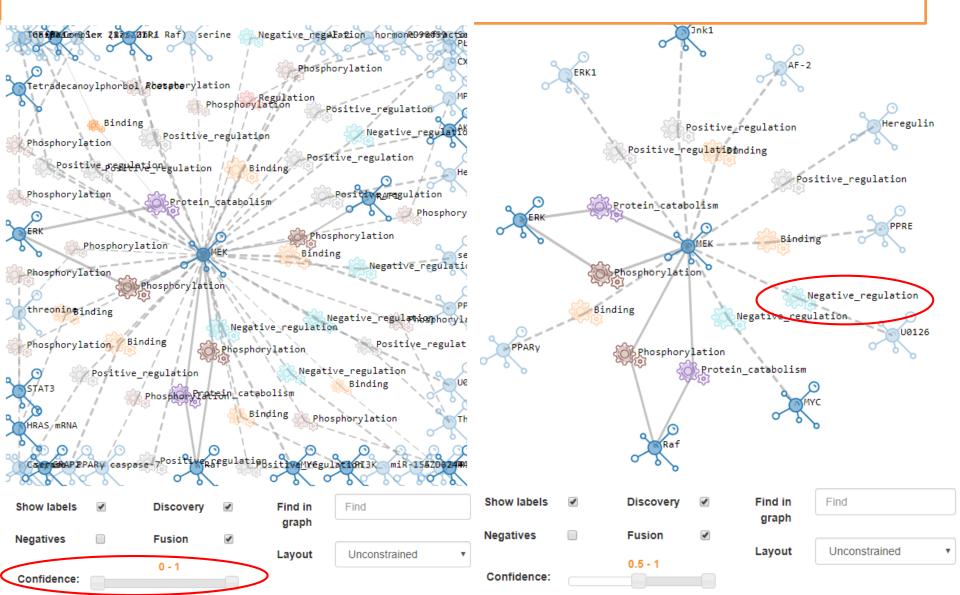


Network Viewer: Discovery mode Extending the model with events found in the literature



Discovery mode

Difficult to explore when too many candidate events are found

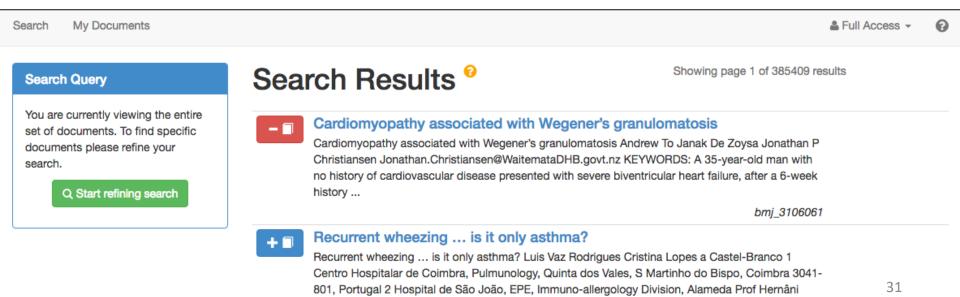


History of Medicine

- <u>http://nactem.ac.uk/hom</u>
- Archives
 - British Medical Journal articles (380,000)

Monteiro, Porto...

- London Medical Office of Health reports (5,000)



Search Query

Term pulmonary consumption New Search Refine Search

Related Terms

pulmonary tubercular disease

pulmonary cedema

tubercular consumption consumption

chronic pulmonary tuberculosis pulmonary heart disease

tuberculous consumption

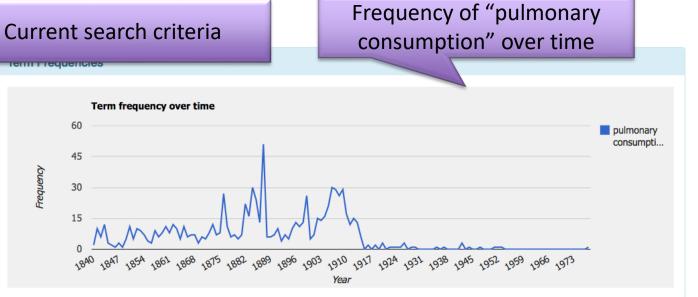
tuberculous pulmonary disease

pulmonary pulmonary syphilis

infectious pulmonary tuberculosis

pulmonary tuberculosis

pulmonary phthisis



Rest in Pulmonary Consumption

REST IN PULMONARY CONSUMPTION. SIR,-In a paragraph which appears in the BRITISH1 MEDICAL JOURNAL of Oct. 4th, it is mentioned that Dr. Berkart has instituted a novel plan of treating pulmonary consumption by the application of mechanical compression made in such a manner by strapping and...

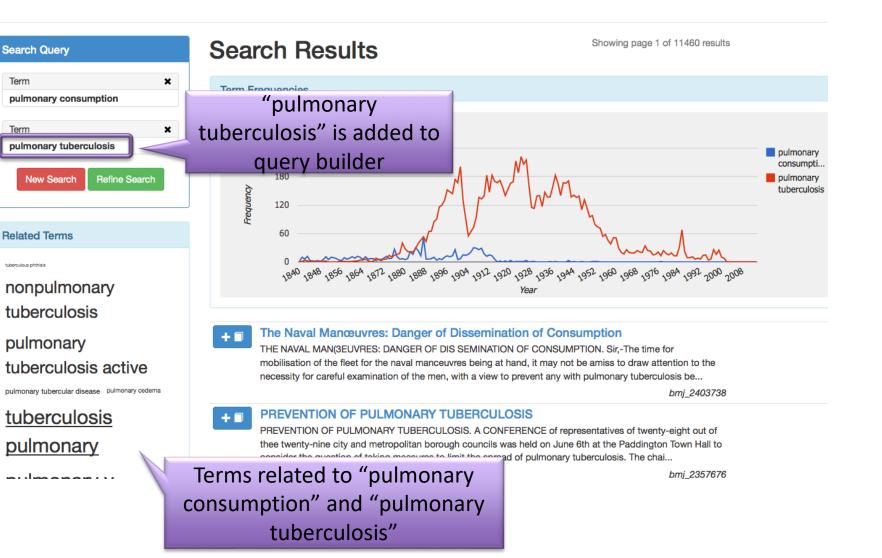
bmj_2294575

GRADUATED LABOUR IN PULMONARY CONSUMPTION

GRADUATED LABOUR IN PULMONARY CONSUMPTION. Sir,-Dr. M. S. Paterson's paper on graduated labour In pulmonary consumption will be read with great interest by every one engaged In treating consumption. For the last two years, ance a short visit to Sir A. E.,Wright's laboratory, endeavour ha...

bmj_2435881

related terms



Content

THE attention.

370 Medical JOURNAL I

INTERNATIONAL MEDICAL CONGRESS:

.*~~~~.

[AUG. 16, I913.'

THE NECESSITY FOR A MORE THOROUGH

CONTROL OF THE MILK SUPPLY IN COMBATING SURGICAL TUBERCULOSIS



Number of "Environmental" entities in the document

CHILDREN.

and others, Haye done valuable work in developing the methods by means of wlliclh sis is of Halliman or bovine origin.

Professor Delepine has for several years carried out ii-antiport investigations in Manchester in Regar d to the tuberculous milk question, and lie has shown that mucll still remains to be done to diminish0 the amount of tuberculous milk whilch is supplied to our large cities.

The control of tlle Edinburgh milk supply is very inadequate, and it is a remarkable fact that the. Public Health Department has no power to enforce the laughter of tile cows with tule from time to time in the city byres; the medical officer of health has merely power to boundaries.

Fortunately comparatively large quantities of the bacilli are necessary to produce tu even -more prevalent amongst children tilan it is.

Post-or0tem records in this country show tilat in about 25 per cent.

of the children possessing tuberculous lesions the infection has taken place through the ammentary trac

As the result of my clinical experience in the Royal Edinburgh Hospital for Sick Children, I have for many years been convinced that Koch and his disciples committee a serious error in practically disregarding the milk as a source of tuberculous infection in children. I could adduce many instances from my own personal observation where the etiological relations between the disease and the "milk history " of the

"Environmental" entity instances in the document

Entities

Anatomical (37)

Biological (32) animal (1) bacilli (4) bacillus (7) bovine bacillus (2) cow (8) guinea-pig (2) human bacillus (4) organism (1) tubercle bacilli (3) Condition (71) Environmental (22) cow (1) dust (1) gold daledalin (1) infective material (1) milk (13) milk product (1) raw milk (1) rural district (2) tuberculous milk (1) Subject (53) case (29) child (21) consumptive patient (1) given case (1) young child (1) Therapeutic or Investigational (19)

I ne earliest observation on the contagious nature of tuberculosis occurs in the Orainances of Iwan in Ingla, thirteen centuries B.C.

In these, pulmonary consumption and swelling of the glands of the neck are declared to be unclean, incurable diseases.

Those who wanted to marry are cautioned not to select a spouse from a consumptive fancily even though the family had a good lineage and possessed great wealth.

These laws show a knowledge of tuberculosis, and as the disease was regarded as an impediment to marriage it seems likely that it was regarded as contagious.

In Europe the earliest reference to the contagious nature of tuberculosis was made about 390 B.C.

In this he states: "The majority of those who had nursed this disease [phthisis] had did not generally subscribe to this view, preferring the idea of the tuberculous diath About one hundred years later Aristotle realized that consumption could be spread Galen and Avicenna held similar views.

Phrases identifying

Causality events are

highlighted

Cases where specific causes of tuberculosis are mentioned in the document

It is more than a tragedy that these early Greek observations were disregarded and forgotten in later centuries, for it was only towards the end of the seventeenth century that in Latin countries such as Spain and Italy the contagious nature of the disease was again being recognized.

In Britain, Edward Mainwaring was possibly the first to assert that tuberculosis was spread by contact, as in 1667 he wrote: "You must not frequently converse with a phthisical person whose unwholesome breath may infect the sound by drawing in put-id vapours which the other breathes forth, but above all a phthisical bedfellow is most dangerous to affect a sound person and chiefly to be avoided."

Later, in 1694, Richard Morton in his book Phthisis logia, which is an outstanding presentation of the symp symptomatology of tuberculosis, wrote: " A contagious principle also propagates the disease, for, as I have found by experience, an affected person may poison a bedfellow by a kind of

ineteenth century, however, leaders of the profession like Bright and Addison (1839),

sness of tuberculosis, and attached great importance to diathesis. dominated men's minds that little or no progress had been made for centuries

rsity of London on March 6, 1947.

urred the disease was tuberculosis, was of the opinion that at least in France

tuberculosis was not usually con contagious.

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also

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That tuberculosis could be produced in animals by inoculation with tuberculous material was shown in 1843 by Klenke, and in 1868 Villein finally proved experimentally that tuberculosis was an infectious disease.

It was Villein who first showed that tuberculous material from bovine sources was more virulent for rabbits than' that from human. In 1867 William Budd, of North Tawton, as a result of a study of the geographical distribution tion of consumption in past and present times, concluded: "The tuberculous matter itself is (or includes) the specific matter of the disease and constitutes the material by which phthisis is propagated from one person to another and disseminated throughout society." Even after the dis cover of the tubercle bacillus by Koch- in 1882 we find from the Collective Investigation Record (British Medical Association, 1883) that many British physicians reported that they had never observed a case of probable trans mission of the disease' from a case of phthisis.

Later, Theobald Smith (1896, 1898) reported the existence of " a distinctively human or sputum, and a bovine variety of tubercle bacillus."

Though Koch proved that the tubercle bacillus was the cause pecans and stressed the importance of inhalation of infected sputum in the spread of the disease, it is strange how in 1901, while accepting that- human and bovine types were different, he yet concluded that the latter was rarely pathogenic for man.

Koch (1882) had undoubtedly found the bovine bacillus in human disease, for he had obtained from a case of caseous pneumonia (No.

27) a culture of tubercle bacilli which after inoculation into the anterior chamber of the eye proved virulent for rabbits.

In 1911 the British Royal Commission proved conclusively that the bovine bacillus was a definite cause of tuberculosis in man.

It is not surprising that Koch went astray when we consider the relative rarity of primary abdominal tuberculosis in Germany as compared with this

Causality (11)

inoculation causes tuberculosis

- bovine bacillus causes tuberculosis
- inhalation causes respiratory tuberculosis

Cause of losion
 Cause of meningitis

- Caused by infection
- Caused by treatment
- Caused by infection
- Cause of respiratory tuberculosis
- Cause of infection
- Cause of tuberculosis

35

Supporting hypothesis generation

Data driven methods complementing human hypothesis generation

Rapid mining of candidate hypotheses from text, validated against experimental data

- Migraine and magnesium deficiency
- Indomethacin and Alzheimer's disease
- Using thalidomide for treating a series of diseases such as acute pancreatitis and chronic hepatitis C

FACTA+

🕲 caffeine - FACTA Search - Mozilla Firefox								\mathbf{X}		
Eile Edit View History Bookmarks Iools Help										
💽 🕞 C 🗶 🏡 http://text0.mib.man.ac.uk/software/facta/a.cgi?query=caffeine/111101/20/0&cat=human&cat= 🏫 🔹 💽 Google 🖉								\mathbf{P}		
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17,094 document(s) mi	ш 17,702	,200 MEDLINE article	s (0.04 se	conds).		k to show).				
Concepts found in th	e docun	nents ranked by [Free	quency <u>P</u>	ointwise Mutu	al Informa	tion Freq. * PMI	[].			
Human Gene/Pro	tein	Disease		Sympto	om	Drug		Compo		
ATP	586 📃	malignant	323 🗐	<u>headache</u>	277 📃	caffeine	17010 📃	<u>caffeine</u>	17077 📃	
ryanodine receptor	523 📃	hyperthermia		fatigue	233 📃	Theophylline	149 🗐	calcium	4337 🗐	
AMP	406 📃	depression	312 📃	pain	230 📃	CTX	62 📃	DNA	978 📃	
adenosine receptor	365 📃	cancer	262 📃	seizures	173 📃	CaCl2	57 📃	sodium	977 📃	
CYP1A2	358 📃	hypertension	215 📃	sleepiness	119 📃	Norepinephrine	57 🔲	water	776 🔲	
ATPase	347 📃	tumor	197 📃	anesthesia	112 📃	CaCl2	56 📃	ATP	589 🔲	
phosphodiesterase	315 🗐	migraine	151 📃	exhaustion	79 🔲	CaCl2	56 🗐	potassium	490 🗐	
endoplasmic reticulum	250 🗐	AT1	128 📃	insomnia	66 📃	Ethanol	52 📃	glucose	449 国	
insulin	235 📃	<u>diabetes</u>	121 📃	nausea	64 📃	Freeze	46 📃	AMP	423 🗐	
cytochrome P450	211 📃	obesity	114 📃	analgesia	59 🔲	vitamin C	37 📃	cAMP	404 🗐	
cardiac muscle	195 📃	<u>cold</u>	107 📃	vomiting	50 🗐	Forskolin	37 🗐	ethanol	332 🗐	
RyR1	188 📃	stroke	92 📃	tremor	38 🗐	lactose	34 🗐	Ryanodine	311 🔲	
<u>spermatozoa</u>	178 📃	<u>ischemia</u>	87 🗐	overweight	35 🗐	citric acid	33 📃	noradrenaline	284 🗐	
microsomal	163 📃	Parkinson's disease	85 🗐	twitching	34 📃	Histamine	32 📃	Adenosine	266 📃	
<u>p53</u>	144 🗐	osteoporosis	85 📃	hypothermia	34 🗐	PCP	28 📃	Mg2+	264 📃	
voltage-gated	136 📃	carcinogenesis	84 🗐	<u>bitter taste</u>	26 📃	<u>vitamin E</u>	25 📃	oxygen	258 🗐	
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Discovering and visualizing indirect associations between biomedical concepts

Bioinformatics

Yoshimasa Tsuruoka^{1,*}, Makoto Miwa^{2,3,4}, Kaisei Hamamoto³, Jun'ichi Tsujii^{3,4,5} and Sophia Ananiadou^{3,4}

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Effects of oral administration of caffeine and D-ribose on mental fatigue.

OBJECTIVE: We examined the effects of administering two different candidate antifatigue substances, caffeine and D-ribose, on mental fatigue. METHODS: In a double-blinded, placebo-controlled, three-way crossover design, 17 healthy volunteers were randomized to oral caffeine (200 mg/d), D-ribose (2000 mg/d), or placebo for 8 d. As fatigue-inducing mental tasks, subjects performed a 30-min Uchida-Kraepelin psychodiagnostic test and a 30-min advanced trail-making test on four occasions. RESULTS: During the tasks, the task performance of the caffeine group was better than that of the placebo group. However, after the fatigue-inducing tasks, although subjective perception of fatigue, motivation, or sleepiness was not significantly different, plasma branched-chain amino acid levels in the caffeine group were lower than those of the placebo group. ... CONCLUSION: Because plasma branched-chain amino acid levels are decreased by mental fatigue, these results suggest that administration of caffeine improved task performance through the enhancement of central nervous system activity without increasing the sensation of fatigue. However, further decreases in branched-chain amino acid levels indicate that **caffeine** might promote deeper **fatigue** than placebo.... 8 Mar

PMID:18178380 Nutrition (Burbank, Los Angeles Country

Efficacy of caff

... In this study, t marmoset monk Three times dur assessed with a was not able to comparable effe CONCLUSIO psychomotor pe PMID:1805791

Enhanced m Caffeine produ restriction or **ca** subjective meas prolonged wakeft.

... However, further decreases in branchedchain amino acid levels indicate that caffeine might promote deeper **fatigue** than placebo

more tiredness and mood disturbance at 3 a.m., and exhibited longer reaction times and more attentional lapses. Heavier catterine consume: exhibited the greatest decreases in Profile of Mood States (POMS) Vigor. ... These effects did not vary with level of habitual caffeine consumption. These findings indicate that consumption of a caffeine-containing food supplement improves subjective state and cognitive performance in fatigued individuals that is likely a result of its **caffeine** content.

PMID:18266548 Experimental and clinical psychopharmacology 2008 Feb

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E-cadherin		Find Ass	ociated C	oncepts View Documents			
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8.1015	13.95	HJMD	0.750	P-cadherin CDHP Polymerase	<u>0.681</u>	E cadherin	
7.9947	8.66	acute	0.667	theadherin (heart)	<u>0.667</u>	inconío	discossovia
7.9349	9.49			is associated with SNAIL3, transcript			
7.5968	8.19	Testis	0.222		0.887 0.556 0.333	E cadherin	
7.5968	8.19	Testis	0.222	ZBTB7C cadherin-9 TRIM86	0.667 0.556 0.333	E cadherin	

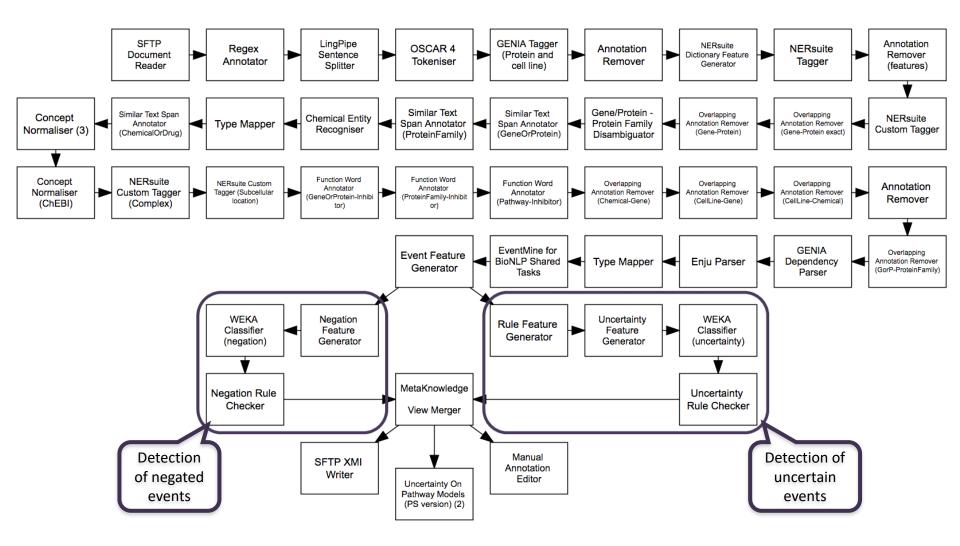
Text mining workflows

- A pipeline that executes particular tools and resources in order; text Mining is not monolithic
- Semantic search



- Different workflows can be created, compared and evaluated by the ability to seamlessly "mix and match" various versions of components
- I know I can build the workflow from the components but which one is the preferred for my task?

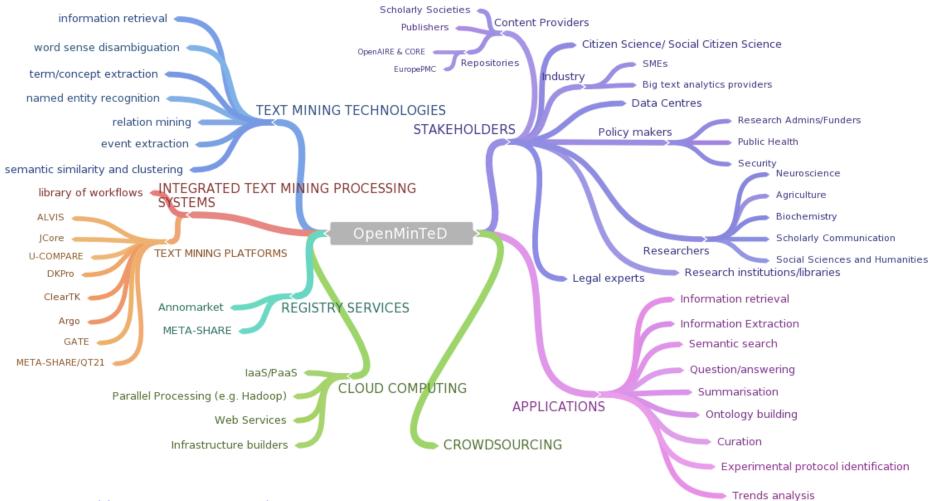
Text mining workflow for uncertainty



Interoperability of tools, platforms and resources

openMIN7ED

Open Mining Infrastructure for Text & Data



http://openminted.eu/

National Plan for Open Science

- What is the critical point in the plan?
 - Open data, open content
 - Linking data with content
 - Reproducibility
 - Research objects
 - Code, experiments, annotations, databases
 - Common Infrastructure for sharing and customisation
 - Standards FAIR

What action to prioritise?

- Access to data and privacy protection
 - difficulty gaining access to EHRs
 - cross-linked with other clinical, public health, biology data and the literature
 - better ways to anonymise data
 - TDM would contribute massively to personalised medicine
 - Security
- Training. Lack of trained personnel in TDM
- Lack of knowledge in existing tools and resources

Thank you. Merci.

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National Centre for Text Mining

www.nactem.ac.uk