

A database to understand tissue growth processes contributing to body or muscle composition

<u>Jérémy Tournayre</u>, Isabelle Cassar-Malek, Matthieu Reichstadt, Brigitte Picard, Nicolas Kaspric and Muriel Bonnet



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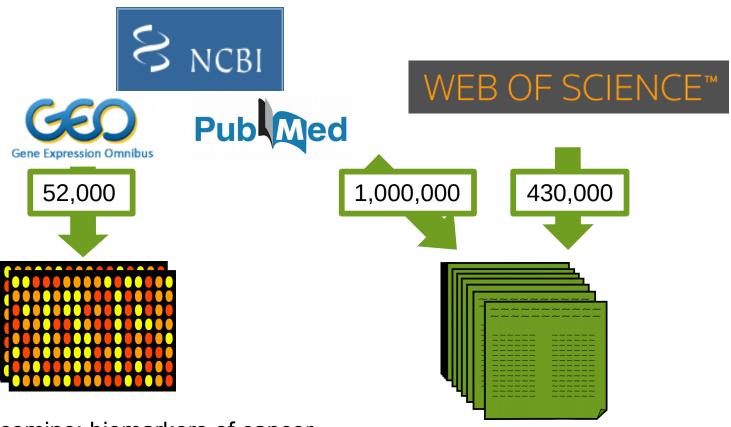




Context



Genomics generates a huge amount of data



Oncomine: biomarkers of cancer

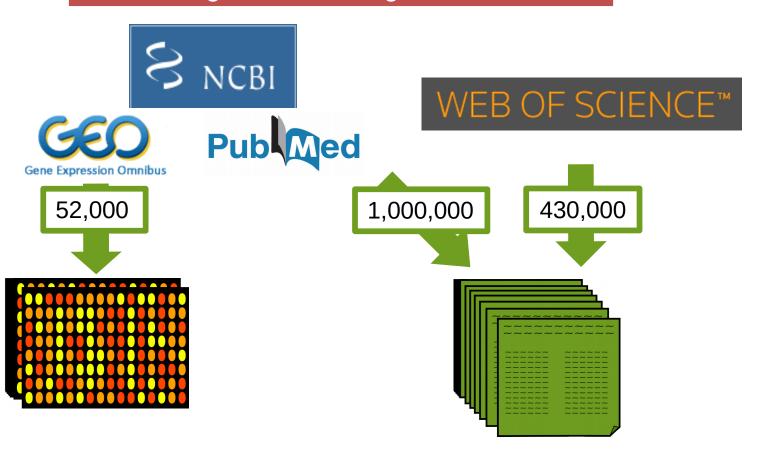
ONCOMINE: A Cancer Microarray Database and Integrated Data-Mining Platform (*Daniel R Rhodes*, 2004)

EURRECA (EURopean micronutrients RECommendations Aligned): Assessing potential biomarkers of micronutrient status by using a systematic review methodology: methods (*Lee Hooper et al.*, 2009)

Context



Genomics generates a huge amount of data



Objective: In the context of carcass and meat qualities we hypothesised that aggregating data could help to understand mechanisms underlying fat and muscle hypertrophy and hyperplasia

Fat&MuscleDB

Workflow of the development of Fat&MuscleDB







Select Extract Classify Visualise Aggregate

Keywords

Cellular and tissular traits linked to muscle and adipose tissue growth (86)

e.g. Adipose tissue, muscle, marbling, double-muscled, carcasses, meat qualities...

Species and cell lines (26)

e.g. Bovine, 3T3-L1, C2C12...

Methods (12)

e.g. Transcriptome, proteome...

Keywords to exclude

e.g. Diseases, carcinoma...



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~28,000 combinations

Keywords to exclude

e.g. Diseases, carcinoma...

Number of references:

~18,000 publications



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~2,500 transcriptomic data







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Cone Expression Omnibus

~410 Publications

Curators

~170 Transcriptomics data

~2,500 transcriptomic data

Data extraction from publications





Select Extract Classify Visualise Aggregate

Target genes of myostatin loss-of-function in muscles of late bovine fetuses. Isabelle Cassar-Malek et al., 2007

Examples of up-regulated genes in DM semitendinosus

Gene symbol		Gene name	ī	old chang	e Homology
26 10 10 3N 1 4Rik		Slc I 6a I 0 Solute carrier family 16		2.76	53%
2410044K02Rik		Thoc3 THO complex 3		2.64	96%
SURFI		Surfeit I		2.54	88%
LOC58504 H	lypot	hetical protein from clones 23549 and 237	762	2.52	86%
Slc26a4		Solute carrier family 26, member 4		2.48	ND
FLJI 3855		Hypothetical protein FLJ 13855		2.36	94%

Semi-automatic extraction with these tools: Tabula, Pdf2text

Data extraction from GEO





Select

Extract

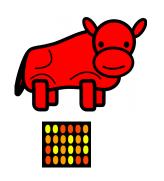
Classify

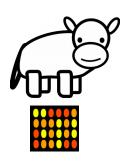
Visualise

Aggregate

What are the transcripts differentially expressed between two conditions?

Double-muscled Charolais (260 days post coitum)





Control Charolais (260 days post coitum)

(source on GEO: GSE5456)

Muscular hypertrophy from genetic origin

Variance mixture from Anapuce (R library) (J. Aubert) (http://cran.r-project.org/web/packages/anapuce/index.html)

up-regulated genes
Lists of: down-regulated genes
stable genes

Classification





Criteria

Muscle or adipose tissue growth from both in vivo and in vitro experiments

e.g. Muscular hypertrophy from genetic origin, 3T3-L1 differentiating into adipocytes...

Physiological traits

e.g. Double-muscled, high marbled, obese...

Cell lines

e.g. C2C12, 3T3-L1, C3H/10T1/2...

Tissues

e.g. Semitendinosus muscle, white adipose tissue...

Breeds

e.g. Charolais, holstein, piedmontese...

Data source

Publication name, chip(s), author(s)

Classification



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First condition	Second condition	State	Species	Breed	Tissue	Cell	Reference	Authors
Muscular hypertrophy from genetic origin	Reference	Double-muscled VS Control	Bovine	Charolais (260 Days)	Semitendinosus muscle	NA	Target genes of myostatin loss-of-function in muscles of late bovine fetuses. Link DOI Link Pubmed	Cassar-Malek I ; Passelaigue F ; Bernard C ; Léger J ; Hocquette JF

Data visualisation



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Transcripts - Increased abundance

Transcripts - Decreased abundance

Analysed by Fat&MuscleDB	Uniprot conversion	Gene Symbol	Gene name	Fold
No	Q3U9N9	2610103N14Rik	Slc16a10 Solute carrier family 16	2.76
No	Q29RH4	2410044K02Rik	Thoc3 THO complex 3	2.64
No	F1N6P2	SURF1	surfeit 1	2.54
No	NA	LOC58504	Hypothetical protein from clones 23549 and 23762	2.52
No	E1BMB2	SLC26A4	solute carrier family 26, member 4	2.48
No	17G9K3	FLJ13855	hypothetical protein FLJ13855	2.36
No	Q9Y236	C8orf1	Chromosome 8 open reading frame 1	2.27
No	G3X6W9	MYBPH	myosin binding protein H	2.27
No	Q13495	CXorf6	chromosome X open reading frame 6	2.19
No	A7MB74	ACCN2	amiloride-sensitive cation channel 2, neuronal	2.17

Data visualisation



Select Extract Classify Visualise Aggregate

Search accessions



Presence

First condition	Second condition	Molecule	State	Species	Breed (age)	Biological sample	Reference	Authors
Proteins secreted by adipose tissues	Proteins secreted by adipose tissues	Proteins	[NA]	Rat	Sprague-Dawley Male (5 Weeks)	Gonadal adipose tissue	Secretome analysis of rat adipose tissues shows location-specific roles	Roca-Rivada, Arturo ; Alonso, Jana ; Al-Massadi, Omar ; Castelao, Cecilia ; Ramon Peinado, Juar Maria Seoane. Luisa
Proteins secreted by adipose tissues	Proteins secreted by adipose tissues	Proteins	Undergoing laparotomy to remove an intramural myoma	Human	(39 Years)	Omental	Comparison of isotope-labeled amino acid incorporation rates	Roelofsen H; Dijkstra M; Weening D; de Vries MP; Hoek A; Vonk RJ

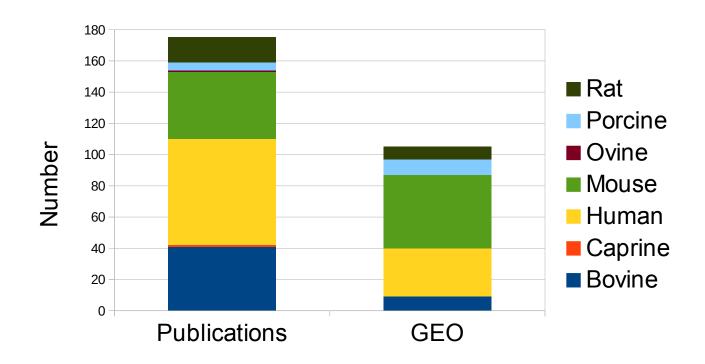
Current contents Fat&MuscleDB



~170 Publications

To be implemented

~100 Gene Expression Omnibus series



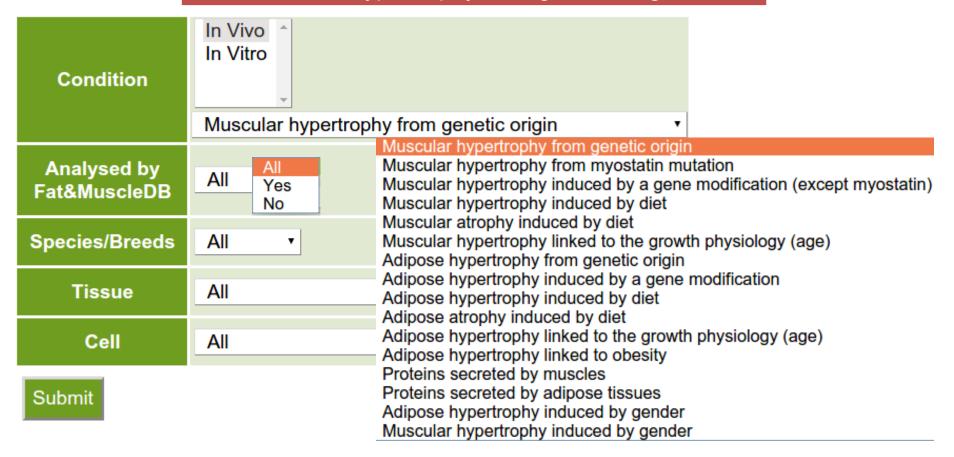
The number of publications and GEO relative to species

Aggregation



Select Extract Classify Visualise Aggregate

What are the transcripts and proteins involved in muscular hypertrophy from genetic origin?



Aggregation



Select Extract Classify Visualise Aggregate

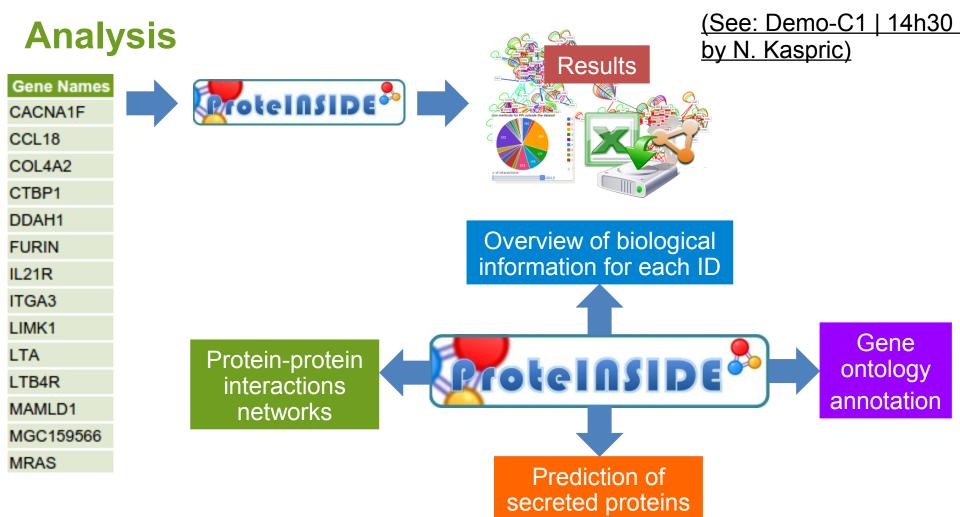
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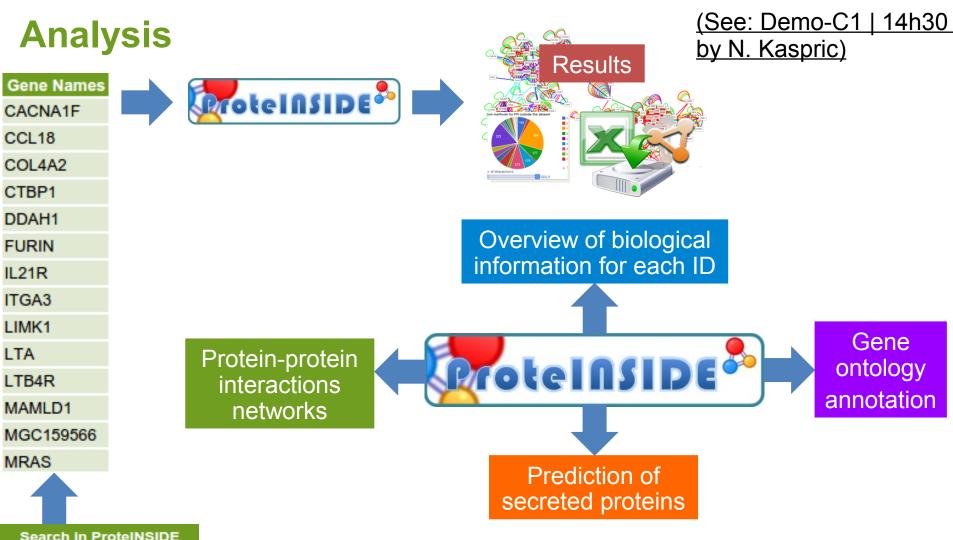
30 pairwise comparisons

Selected	Muscular hypertrophy from genetic origin	Adipose hypertrophy from genetic origin	Proteins	Higher intramuscul fat developmei	Bovine	Korean (27 Months)	Longissimus dorsi	NA	Differentially expressed proteins during fat accumulation in bovine	Zhang Q; A Lee HG; Han JA; Kim EB; Kang SK; Yin J:
Selected	Muscular hypertrophy from genetic origin	Adipose hypertrophy from genetic origin	Transcripts	Control	Bovine	Piedmontese (3 Months) VS Wagyu (3 Months)	Longissimus dorsi	NA	GSE25554	Reverter A ; Hudson N

Decreased abundance	Increased abu	ındance	Stable abunda	ince
Analysed by Fat&MuscleD	B Uniprot	Entry N	ame Review	red Count ▼
No	Q0P571	MLRS_B	OVIN Yes	2
No	Q3SWW8	TSP4_B0	OVIN Yes	2
No	A7MBI7	COMT_B	OVIN Yes	2
No	Q5KR47	TPM3_B	OVIN Yes	2



ProteINSIDE to Easily Investigate Proteomics Data from Ruminants: Application to Mine Proteome of Adipose and Muscle Tissues in Bovine Foetuses (*Kaspric N et al.*, 2015)



Search in ProteINSIDE

Decreased

Increased

Submit in Basic Analysis

Submit in Custom Analysis

ProteINSIDE to Easily Investigate Proteomics Data from Ruminants: Application to Mine Proteome of Adipose and Muscle Tissues in Bovine Foetuses (Kaspric N et al., 2015)

(Realised by A. Perot)

Conclusions





Perspectives

- Continue to feed Fat&MuscleDB
- Address research questions: e.g. what are the proteins involved in the cross-talk between adipose tissues and muscles (secretome)

Minimise unnecessary redundancy in research efforts by a better use of available data

Muriel Bonnet



Isabelle Cassar-Malek



Matthieu Reichstadt



Brigitte Picard



Nicolas Kaspric



Antoine Perot



Anne de la Foye



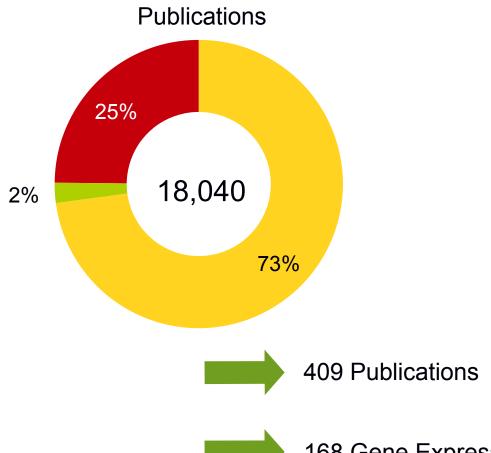
Thank you for your attention

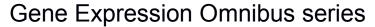


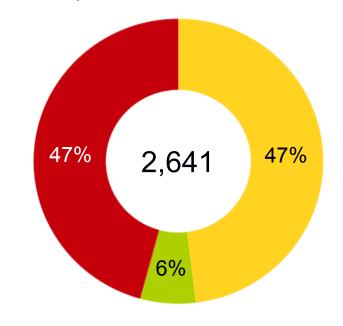
Rejected

Not seen

Selected







168 Gene Expression Omnibus series