

FRAUNHOFER-INSTITUTE FOR ALGORITHMS AND SCIENTIFIC COMPUTING (SCAI)

Meta-Data Curation Guidelines of Publicly Available Gene-expression Data



Short description

This document describes a guideline for annotating Microarray metadata for datasets coming from public repositories like Gene Expression omnibus (GEO) and ArrayExpress(AE).

The relevant fields were described with the help of disease experts. This document describes the guidelines of how to annotate each field and where to find the relevant information. Some possible examples harbouring each field is also provided.

Currently, only Alzheimer's Disease experiments are considered. Each experiment is curated by two annotators to obtain good inter-annotator consistency.



What each term means?

Term	Meaning
Field	Field is the column name in <i>NeuroTransDB</i> . Each field represents different kind of data obtained
	through curation
Description	Describes what data has to be filled in each of the fields concerned
Example	Examples of data filled in each field
GUIDELINE	Describes the importance of the field
SOURCE	Where to obtain the information

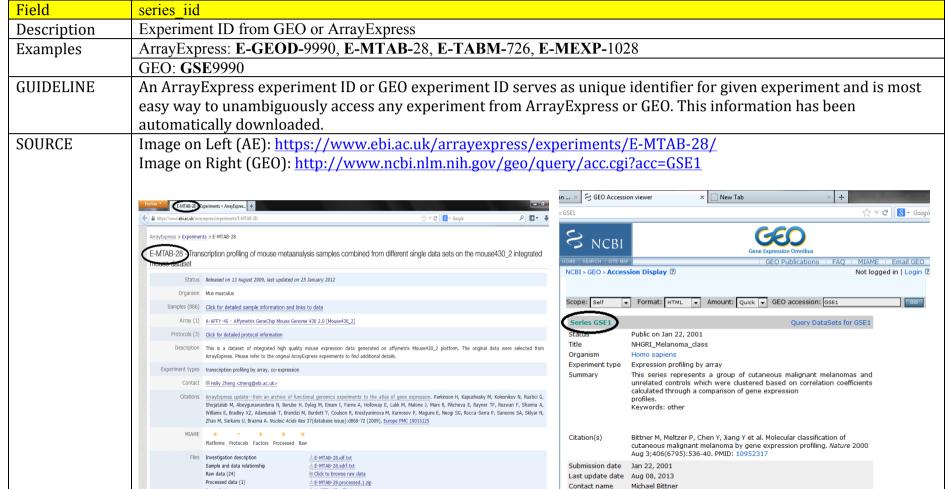
Databases concerned

Database Name	URL	Maintained by
ArrayExpress (AE)	https://www.ebi.ac.uk/arrayexpress/	Europe
GEO (Gene Expression Omnibus)	http://www.ncbi.nlm.nih.gov/geo/	United States of America

^{*}Characters given in **BOLD** are important



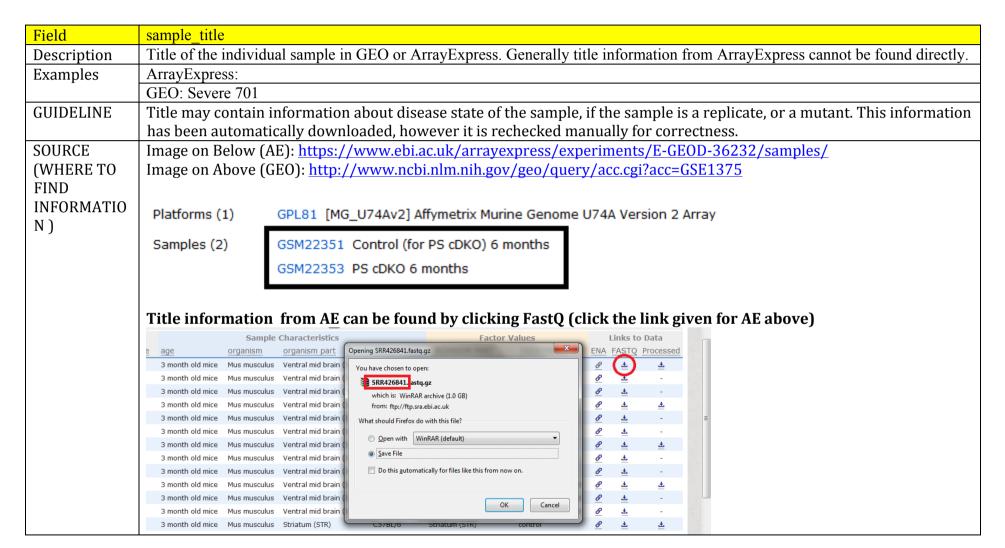
S	C	Α	ı





Field	sample_id			
Description	Sample ID of the samples used in Experiment/Series in A	rrayExpress/0	GEO respectively	
Examples	ArrayExpress: GSM 32049			
	GEO: GSM 32054			
GUIDELINE	ArrayExpress sample ID's or GEO sample ID's serve as	unique iden	tifier for the given samples from an Experiment and	
GOIDEEINE	these ID's are the most easy way to unambiguously acc	•		
	information has been automatically downloaded.	less ally sain	pie ironi Array Express or deo databases. Tins	
COLIDCE	, and the second	/	/F CEOD 1020 / l /	
SOURCE	Image on Left (AE): https://www.ebi.ac.uk/arrayexpre			
	Image on Right (GEO): http://www.ncbi.nlm.nih.gov/g	<u>geo/query/a</u>	cc.cgi?acc=GSE1829	
		Contact name	Winnie Liang	
	ArrayExpress	Street address	ne Translational Genomics 445 N. Fifth Street	
	MilayLApiess	City	Phoenix	
	Home Experiments Arrays Submit Help About ArrayExpress	State/province	AZ	
		ZIP/Postal code	85012	
	ArrayExpress > Experiments > E-GEOD-1829 > Samples and Data*	Country	USA	
	E-GEOD-1829 - Transcription profiling of mouse hippocampus t	Platforms (1)	GPL341 [RAE230A] Affymetrix Rat Expression 230A Array	
		Samples (11)	GSM32049 hippocampus, left: Ketogenic 1-2_e1_le1	
	Samp	∃ Less	GSM32050 hippocampus, left: Ketogenic 3-4_e1_le1	
	_Source Name ^ Organism		GSM32051 hippocampus, left: Ketogenic 5-6_e1_le1	
	GSE1829GSM32049 Mus musculus		GSM32052 hippocampus, left: Ketogenic 7-8_e1_le1	
	GSE1829GSM32050 Mus musculus		GSM32053 hippocampus, left: Ketogenic 9-10_e1_le1	
	GSE1829GSM32051 Mus musculus		GSM32054 hippocampus, left: Control 1-2_e1_le1	
	GSE1829GSM32052 Mus musculus		GSM32055 hippocampus, left: Control 3-4_e1_le1	
	GSE1829GSM32053 Mus musculus		GSM32056 hippocampus, left: Control 5-6_e1_le1	
	GSE1829GSM32054 Mus musculus		GSM32057 hippocampus, left: Control 7-8_e1_le1	
	GSE1829GSM32055 Mus musculus		GSM32058 hippocampus, left: Control 9-10_e1_le1	
	GSE1829GSM32056 Mus musculus		GSM32059 hippocampus, left: Control 11-12_e1_le1	
	GSE1829GSM32057 Mus musculus	Relations	•	
	GSE1829GSM32058 Mus musculus	BioProject	PRJNA90399	
	GSE1829GSM32059 Mus musculus			







Field	sample_type					
Description	Biomolecules extracted from samples for analysis in Experiments					
Examples	ArrayExpress: low-molecuar-weight RNA derived from total RNA					
	GEO: RNA, total RNA, SRA, genomic DNA, pooled RNA from nine control mice/rat, Single RNA					
GUIDELINE	The entity extracted from organism/cells are proteins or nucleic acids or in special case SRA					
SOURCE	Image on Left (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM884353					
	Image on Right (AE): http://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-48028/samples/					
	Mostly you can find information about the "type" from GEO sample files but for exact information one must read					
	paper and/or examine ArrayExpress sample data file sheet. There are experiments where in one single sample,					
	RNA's are pooled from many pateints/organisms, in that cases "type" should be "pool from 'n-number'					
	mice/rat/patients" or just pool. In case of SRA (Sequence Read Archive) one can enter "SRA:total RNA" in "type" field					
	(as shown in figure). Consider experiment E-GEOD-48028 where sample "type" was found to be "RNA" in GEO, but					
	AE had more definite and precise information about the "type" i.e. "low-molecuar-weight RNA derived from total					
	RNA" was found. If this information has been automatically downloaded, then it is rechecked manually for					
	correctness.					
	Sample GSM884353 Query DataSets for GSM884353 Status Public on Sep 04, 2012					
	Title Control VMB RNA-Seq					
	Sample type SRA					
	Source name Brain Organism Mus musculus Source Name ^ Sample description Sample source					
	Characteristics strain: C578I/6					
	age: 3 month old mice GSM1165972 1 low-molecuar-weight RNA derived from total RNA wild type mice tissue: Ventral mid brain (VMB)					
	treatment: Control GSM1165973 1 low-molecuar-weight RNA derived from total RNA Tg2576 Treatment protocol 3 month old C578l6 mice were administered pesticides using subcutaneous					
	alzet pumps for one week. After one week the mice were harvested and ventral mid brain and striatum collected for RNA isolation. RNA was purified GSM1165974 1 low-molecuar-weight RNA derived from total RNA wild type mice					
	Followed by qualitative and quantitative analysis. Extracted molecule total RNA GSM1165975 1 low-molecular-weight RNA derived from total RNA APP/PS1M146V/					
	Extracted indicate total riva					
ĺ						



Field	disease_keyword
Description	Keyword used during data retrieval
Examples	ArrayExpress: Alzheimer
	GEO: Alzheimer
GUIDELINE	The keyword used for retrieval
SOURCE	This information is mostly automatically mined out

Field	domain_specificity
Description	If the experiment or rather sample is relevant to the disease keyword used for retrieval
Examples	This field takes binomial values. "Yes" if experiment is related to disease domain else "No"
GUIDELINE	Gives information about the relevance of sample to the disease being looked up. Relevance means that the experiment is substantially related to Alzheimer's diseases. It's a Boolean value either YES or NO. This has helped us improve our data retrieval method by improving our keyword (synonym) list
SOURCE	Paper, GEO or ArrayExpress description page

Field	experiment_title
Description	Title of the experiment, which is downloaded automatically
Examples	For experiment GSE12685, the title provided by two databases are:
	ArrayExpress: Transcription profiling of human Alzheimers disease patients to identify genes regulating synaptic function and
	neuroplasticity in incipient AD
	GEO: Expression of mRNAs Regulating Synaptic Function and Neuroplasticity in Incipient AD
GUIDELINE	The title is automatically retrieved from the databases
SOURCE	This information is mostly automatically mined out



Field	age						
Description	_		ne sample is retrieved. ation about age at death			also included if cells an also be included	re used in an
Examples	Age organism: 6 months, 6 mon, Post Natal Day 15, P19, 24 - 28 weeks, 6-8 wks, 3d, age 540 days, Gestation day 15-16, 10 day-old, 12 months, age at onset: 25yr/age at death: 74yrs						
	Age cell:	15 days, age of culture	e day 5				
GUIDELINE	Microarra	ay analysis reveals ag	ge-related differences	in gene express	ion and hence	inclusion of age data	is vital
	Image on	Left (GEO): http://w	ww.ncbi.nlm.nih.gov	/geo/query/acc	.cgi?acc=GSM	<u> 26703</u>	
	Image on	Right (AE): https://	www.ebi.ac.uk/array	express/experin	nents/E-GEOD	0-36232/samples/	
	Age infor	mation can either be	explicitly mentioned	in AE/GEO or ha	ave to be sear	ched from research pa	aper linked to
	experiment. Age information is usually found in Materials and Methods section of paper, figure or supplement provided in research paper. One can find cell age information from image on Left. Sometimes age information				•		
	the form of range. If age information was automatically retrieved we check again if it is correct.						
	the form	oi range. Ii age iniori	nation was automatic	cally retrieved w	e che <u>ck again</u>	if it is correct.	
	Sample GSM26	703	nation was automatic Query DataSets for GSM26703	cally retrieved w		if it is correct. Characteristics	
	Sample GSM26 Status Title	703 Public on Jul 30, 2004 sAPPalpha #1		ally retrieved w			strain or line
	Sample GSM26 Status Title Sample type	703 Public on Jul 30, 2004 sAPPalpha #1 RNA			Sample	Characteristics	strain or line
	Sample GSM26 Status Title Sample type Source name Organism	703 Public on Jul 30, 2004 sAPPalpha #1 RNA hippocampal slice cultures Mus musculus		age	Sample organism	Characteristics organism part	
	Sample GSM26 Status Title Sample type Source name	703 Public on Jul 30, 2004 sAPPalpha #1 RNA hippocampal slice cultures Mus musculus		age 3 month old mice	Sample organism Mus musculus	Organism part Ventral mid brain (VMB)	C57BL/6
	Sample GSM26 Status Title Sample type Source name Organism	Public on Jul 30, 2004 sAPPalpha # 1 RNA hippocampal slice cultures Mus musculus ule total RNA Organotypic hippocampal slice cultures (C57B6/SJL) were maintained in cult	Query DataSets for GSM26703 ures of male, postnatal day 15 mice ure for 15 days and treated with 1 nM	age 3 month old mice 3 month old mice	Sample organism Mus musculus Mus musculus	Characteristics organism part Ventral mid brain (VMB) Ventral mid brain (VMB)	C57BL/6 C57BL/6
	Sample GSM26 Status Title Sample type Source name Organism Extracted molecu	Public on Jul 30, 2004 sAPPalpha # 1 RNA hippocampal slice cultures Mus musculus sle total RNA Organotypic hippocampal slice cult (C57B6/SJL) were maintained in cult sAPPalpha (sAPPalpha) for an additic filter inserts in Neurobasal media wi mM glutamine. The sample consists	Query DataSets for GSM26703 ures of male, postnatal day 15 mice ure for 15 days and treated with 1 nM nal 24 h Cultures were maintained no. the B-27 supplement (Invitrogen) and 1 of 12 slices from 4 mice. Data were	age 3 month old mice 3 month old mice 3 month old mice	Sample organism Mus musculus Mus musculus Mus musculus	Characteristics organism part Ventral mid brain (VMB) Ventral mid brain (VMB) Ventral mid brain (VMB)	C57BL/6 C57BL/6 C57BL/6
	Sample GSM26 Status Title Sample type Source name Organism Extracted molecu	Public on Jul 30, 2004 sAPPalpha #1 RNA hippocampal slice cultures Mus musculus ule total RNA Organotypic hippocampal slice cultu (C5786/SJL) were maintained in cult sAPPalpha (sAPPalpha) for an addition filter inserts in Neurobasal media wil	Query DataSets for GSM26703 ures of male, postnatal day 15 mice ure for 15 days and treated with 1 nM nal 24 h Cultures were maintained no. the B-27 supplement (Invitrogen) and 1 of 12 slices from 4 mice. Data were	age 3 month old mice 3 month old mice 3 month old mice 3 month old mice	Sample organism Mus musculus Mus musculus Mus musculus Mus musculus	Characteristics organism part Ventral mid brain (VMB) Ventral mid brain (VMB) Ventral mid brain (VMB) Ventral mid brain (VMB)	C57BL/6 C57BL/6 C57BL/6 C57BL/6



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Field	gender
Description	Gender of the sample organism
Examples	Male, Female, f, m, Homo, Mixed, Other, (pooled age information is given as $x\%$ male and $y\%$ female (x = numerical value, y = numeric value and $x+y$ = 100)
GUIDELINE	Gender data is used to analyse gender biased or gender specific gene expression analysis and hence gender information is vital
SOURCE	Image reference: http://www.ncbi.nlm.nih.gov/pmc/articles/PMC3341364/ Gender information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. Gender information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper. If gender information was automatically generated we check if it is correct. Consider the ID E-GEOD-36232 where gender information was not available in GEO and AE but was found from paper.
	Il experiments were conducted within National Institutes of ealth guidelines for animal research and were approved by ne Weill Cornell Medical College Animal Care and Use ommittee under animal protocol number 2010-0043. The nice were kept on a 12-hr light/dark cycle, with food and ater available ad libitum. All surgeries were performed nder isoflurane anesthesia and all efforts were made to inimize animal suffering The month old C57Bl6 male mice (Jackson Labs, Bar arbor, Maine, USA) were administered PQ, PY and MNPQ The self-the month old C57Bl6 male mice (Jackson Labs, Bar arbor, Maine, USA) were administered PQ, PY and MNPQ The weill Cornell Medical College Animal Care and Use of pestanal grade (Sigma Inc, MO, USA) using subcutaneous alzet pumps (Durect Corporation, Cupertino, CA, USA) for 7 days. Concentrations were PQ (10 mg/kg/day). PY (3.5 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day)+MN (30 mg/kg/day) and a combination of PQ (10 mg/kg/day) and a combina



Field	age_info_section
Description	Page or section where the information about the age was found
Examples	http:// or sdrf or ftp://
GUIDELINE	Serves as a reference link to literature information from Paper or AE/GEO
SOURCE	Address bar of the web page where you found age information or in case of downloaded supplementary files one can right
	click the hyperlink of "supplementary file" -> Copy link location and paste in the "age_info_section" field. If age information
	was found from AE or GEO web pages, insert the specific URL from the AE/GEO sections

Field	gender_info_section
Description	Page or section from where gender information about the sample was found
Examples	http:// or sdrf or ftp://
GUIDELINE	Serves as a reference link to literature information from Paper or AE/GEO
SOURCE	Address bar of the web page where we found gender information or in case of downloaded supplementary files one can right click the hyperlink of "supplementary file" -> Copy link location and paste in the "gender_info_section" field. If Gender information was found from AE or GEO web pages, insert the specific URL from the AE/GEO sections.

Field	series_id
Description	The experiment IDs as downloaded from the databases
Examples	GSE12685 or E-GEOD-12685
GUIDELINE	Identifier of the experiment
SOURCE	Series webpage of GEO



Field	source_database
Description	Name of the database from which the experiment's information was downloaded
Examples	GEO or ArrayExpress
GUIDELINE	The data origin information

Field	geo_series_id
Description	The experiment IDs from GEO
Examples	GSE12685
GUIDELINE	Since most of the experiments are firstly downloaded from ArrayExpress, the GEO experiment IDs are represented as E-GEOD-12865. However, we automatically revert the series ID here and save the original ID from GEO
SOURCE	Series webpage of GEO

Field	original_sample_id
Description	The sample Ids downloaded as is from the databases
Examples	For experiment E-GEOD-12685, an example sample id from ArrayExpress is GSE12685GSM318211
GUIDELINE	ArrayExpress modifies the sample IDs from GEO. For example, in GSE12865 the sample ID is GSM318211. However, ArrayExpress refers it as GSE12685GSM318211.
SOURCE	Sample webpage of GEO



Field	phenotype							
Description	Phenotype of the sample							
Examples	Diseased_AD, Diseased_PD, Diseased_EP, normal, mutant, control, normal_elderly_control, young_control, AD_control, KO mice/rat, KI mice/rat, Treated vs non treated mice/rat, 3xTg, 5x backcrossed mice/rat, pool young, pool adult, Sham							
	mice/rat							
GUIDELINE	This field describes the observable characteristics or traits that can distinguish between normal and diseased							
	patient/cell							
SOURCE	not find such specifically in Image on Left	information diagrams, (GEO): http://diagrams.	s either available in title of in ArrayExpress or GEO for graphs, tables and supplents: p://www.ncbi.nlm.nih.gov.ps://www.ebi.ac.uk/arrayo	iles a final check nentary material /geo/query/acc	should be don s given in rese .cgi?acc=GSE6	ie in resea arch papo 771	arch paper and er.	uo
				Sample_source_name	e Organism age	bio-source nar	me cell tyne	
					Homo sapiens 63 years	EC control 1	layer III neurons	
				brain, Entorhinal Cortex	Homo sapiens 85 years	EC control 2	layer III neurons	
				brain, Entorhinal Cortex	Homo sapiens 80 years	EC control 3	layer III neurons	
				brain, Entorhinal Cortex	Homo sapiens 80 years	EC control 4	layer III neurons	
	Platforms (1)	GPL4757 Ior	n Channel Splice Array	brain, Entorhinal Cortex	Homo sapiens 102 years	EC control 5	layer III neurons	
			· · · · · · · · · · · · · · · · · · ·	brain, Entorhinal Cortex	Homo sapiens 79 years	EC control 6	layer III neurons	
	Samples (10)	GSM156001	Temporal Cortex Control (mTLE) 1		Homo sapiens 76 years	EC control 7	layer III neurons	
	∃ Less	GSM156003	Temporal Cortex Control (mTLE) 3		Homo sapiens 83 years	EC control 8	layer III neurons layer III neurons	
		GSM156051	Temporal Cortex Control (mTLE) 2		Homo sapiens 79 years Homo sapiens 88 years	EC control 9 EC control 10	layer III neurons	
			Temporal Cortex Control (mTLE) 4		Homo sapiens 82 years	EC control 11	layer III neurons	
				brain, Entorhinal Cortex	Homo sapiens 69 years	EC control 12	layer III neurons	
		GSM156159	Temporal Cortex Control (mTLE) 5	brain, Entorhinal Cortex	Homo sapiens 78 years	EC control 13	layer III neurons	
		GSM156160	Temporal Cortex Control (mTLE) 6	brain, hippocampus	Homo sapiens 85 days	HIP control 1	layer III neurons	
		GSM156161	Temporal Cortex Control (mTLE) 7	brain, hippocampus	Homo sapiens 80 years	HIP control 2	layer III neurons	
			Temporal Cortex Control (mTLE) 8	brain, hippocampus	Homo sapiens 80 years	HIP control 3	layer III neurons	
				brain, hippocampus	Homo sapiens 102 years		layer III neurons	
		GSM156164	Temporal Cortex Control (mTLE) 9	brain, hippocampus	Homo sapiens 63 years	HIP control 5	layer III neurons	
		GSM156169	Temporal Cortex Control (mTLE) 10	brain, hippocampus	Homo sapiens 79 years	LIS COURSI 6	layer III neurons	



Field	phenotype_info_section
Description	Page or section from where phenotype information about the sample was found
Examples	http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE32534
GUIDELINE	Serves as a reference link to literature information from Paper or AE/GEO
SOURCE	Address bar of the web page where you found phenotype information or in case of downloaded supplementary files one
	can right click the hyperlink of "supplementary file" -> Copy link location and paste in the "phenotype_info_section" field. If
	phenotype information was found from AE or GEO web pages, insert the specific URL from the AE/GEO sections.

Field	stage
Description	This field describes disease stage of AD
Examples	onset(SNCA inclusion bodies), Braak1 stage according to Braak and Braak/mmse score-15
GUIDELINE	Provides information on the disease progression. It can be name of the stage (mild/severe) or values of different scoring system (mini-mental state examination score/Hoehn and Yahr's Staging score). If both are available, we include them as is. It is crucial to mention the criterion behind the staging.

Field	stage_info_section
Description	Page or section from where phenotype information about the sample was found
Examples	http://
GUIDELINE	Serves as a reference link to literature information from Paper or AE/GEO
SOURCE	Address bar of the web page where you found stage information or in case of downloaded supplementary files one can
	right click the hyperlink of "supplementary file" -> Copy link location and paste in the "stage_info_section" field. If
	phenotype information was found from AE or GEO web pages, insert the specific URL from the AE/GEO sections.



Field	replicate_type
Description	If the sample is a replicate, then what type it is: biological or technical
Examples	Technical
GUIDELINE	This supports the meta-analysis
SOURCE	Replicate information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.

Field	replicate_of
Description	If the sample is a replicate, then provide the id of its replicate
Examples	GSM969416
GUIDELINE	This supports the meta-analysis
SOURCE	Replicate information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.

Field	sample_id
Description	The sample Ids resolved to the correct names in GEO
Examples	For experiment E-GEOD-12685, an example sample id from ArrayExpress is GSE12685GSM318211. This is resolved to GSM318211
GUIDELINE	ArrayExpress modifies the sample IDs from GEO. For example, in GSE12865 the sample ID is GSM318211. However, ArrayExpress refers it as GSE12685GSM318211.
SOURCE	Sample webpage of GEO



Field	pmd				
Description	PMD= Postmortem duration. Duration between the death of the patient (if dead) and extraction of the sample.				
Examples	2 min or 14 hrs				
GUIDELINE	Postmortem duration/ Postmortem interval	determine the RNA integrity of the	extracted sample.		
SOURCE	Image reference left (Research Paper): h Image on right (AE): https://www.ebi.ac		•		
	PMD/ PMI information can either be exp to experiment. This information is usuall files provided in research paper. Consider but was later found after reading correspondent.	ly found in Materials and Methods er ID E-GEOD-47038 for which pmo	section of paper, figure or sup	plementary	
	PMC CT I I	↑ ♥ ♥ 🔞 ▼ 6009le rain identifies target genes in Alzheimer's disease 136(10): 3018–3027. Characterist	ien.		
	vithin the first 2 min post-mortem and kept at -80° C. Brain areas were dissected from cryostat brain sections (-20° C) with the help of a $\times 5$ magnifying lens, following their	13 7371 3.3	MD 28 PMD 30 PMD 18 PMD 2 5 PMD	20 PMD 17 PMD 7	
	stereotaxic coordinates (Paxinos and Franklin, 2001), as	and three 3TG control mice were used	00		
	previously described (Roux et al., 2003; Panayotis et al.,	validations and RNA assays. Four APF	9		
	2011a, b). For validations of epigenetically deregulated genes	wild-type animals were deeply anaestlank	03		
	in Alzheimer's disease, frontal cortex samples from two transgenic models were used, namely APP/PSEN1 (Borchelt	intraperitoneal injection (0.2 ml/10 g ank mixture of ketamine (100 mg/kg) and ank	53		
	et al., 1997) and 3xTg-AD (Oddo et al., 2003) mice. Five	before intra-cardiac perfusion with 49		41	
	12-month-old AβPP/PS1 and five wild-type littermates, and	PBS. Brains were removed, post-fixed	00	41	
	three 18-month-old 3TG and three 3TG control mice, were assayed for DNA methylation using the mouse brain	4 h and processed for immunohistoch of four APP/PS1 and four wild-type m	30		



Field	ph_value
Description	pH value of the sample at the time of analysis.
Examples	4
GUIDELINE	It determines the sample source quality
SOURCE	pH information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.

Field	cause_of_death
Description	Describes the reason for death of the source from which the sample was extracted
Examples	Heart attack
GUIDELINE	This helps us to determine if AD is the cause of death or other/comorbid diseases
SOURCE	This information is mostly mentioned in the associated research papers linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.

Field	comorbidity
Description	Describes the co-existence of other diseased
Examples	Diabetes Mellitus
GUIDELINE	This helps us to determine if AD is a side effect or vice versa of other co-existing diseases
SOURCE	This information is mostly mentioned in the associated research papers linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.



Field	priority
Description	Describes the priority of the experiment after the classification step
Examples	AD1
GUIDELINE	This is the Priority classification information
SOURCE	Manually carried out by expert curators. Please refer to the paper for guidelines

Field	sample_priority
Description	Describes the priority of the sample identified during meta-curation
Examples	AD1
GUIDELINE	Some experiments contain samples that are derived from humans/animals and/or cell lines. In other cases the samples are from different diseases such as Breast Cancer. To improve the retrieval method we have additionally assigned priority to individual samples
SOURCE	Manually carried out by expert curators. Please refer to the paper for guidelines

Field	raw_file_location
Description	URL to the raw file provided by databases
Examples	http://www.ncbi.nlm.nih.gov/geo/download/?acc=GSE12685&format=file
GUIDELINE	This provides us information as to where to download raw files
SOURCE	GEO or ArrayExpress series webpage



Field	cell_types
Description	Types of cells used as samples
Examples	primary hippocampal neurons, primary mouse embryonic fibroblasts (MEFs), DA neurons, non-DA neurons and glia cells
GUIDELINE	Important for gene expression analysis and difference in gene expression in different cell types
SOURCE	Image on Left (GEO http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM512701 Image on Right (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-36232/samples/
	Cell type information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.
	HtrA2 KO cells show increased sensitivity to mitochondrial stress characterized by enhanced CHOP expression To test the contribution of HtrA2 to the transcriptional activation of a ctroce response, wild-type (WT) and HtrA2 KO primary mouse embryonic fibroblasts (MEFs) were challenged with an inducer or introcriorana stress and treat transcriptional response was monitored using microarray technology (experimental outline, Supplementary Figure S2a). WT and HtrA2 KO MEFs were chosen, as these cells have been shown earlier to display differential sensitivity to drug-induced stresses triggered by specific respiratory inhibitors such as rotenone, which specifically blocks complex I. This analysis revealed that in HtrA2 KO MEFs, the number of rotenone-induced transcriptional changes was significantly enhanced, suggesting that loss of HtrA2 results in an enhanced transcriptional response (Figure 1a). Among the genes most highly induced by rotenone treatment in HtrA2 KO cells was the transcription factor CHOP (Supplementary Table S1), which is known to be induced in response to various stresses such as activation of the UPR ^{ER} , UPR ^{mt} and ISR.



Field	raw_filename
Description	Raw file name for each sample, extracted from the raw file location
Examples	GSM318213.CEL.gz
GUIDELINE	This provides easy mapping of raw file names to the respective sample phenotype for gene expression analysis
SOURCE	GEO or ArrayExpress sample webpage

Field	cell_lines
Description	If the cell lines are used as samples or derived from the tissue extracetd
Examples	Embryonic mesenchymal cell line
GUIDELINE	Important for gene expression analysis and difference in gene expression in different cell lines
SOURCE	Image on Left (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM26703
	Image on Right (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-36232/samples/
	Cell line information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. Cell line information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.

Field	additional_info
Description	Any interesting Data found by curator
Examples	
GUIDELINE	Some interesting outcome of experiment, if samples passed or failed QC, ethnicity, any outliers in data would not go unnoticed if
	they are noted in add_information column.
SOURCE	GEO/ AE files
	Research paper



Field	id
Description	Internal identifier automatically generated by SQL database during upload
Examples	235
GUIDELINE	Important if we want to build relations between several tables

Field	annotation_remarks
Description	Sometimes used as the basis as to why the annotator annotated certain field
Examples	
GUIDELINE	This information is necessary if we want to trace back some ambiguous annotation as to why a certain field was
	annotated
SOURCE	GEO/ AE files
	Research paper

Field	body_fluid			
Description	sample molecules are extracted from fluids of body			
Examples	Whole Blood, Venous blood, peripheral blood			
GUIDELINE	Separate analysis of gene expression changes, to differentiate it from the tissue expression			
SOURCE	Body fluid information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to			
	experiment. Gender information is usually found in Materials and Methods section of paper, figure or supplementary files			
	provided in research paper.			



Field	brain_regions_or_tissue					
Description	What region of brain or which tissue is used as sample					
Examples	Brain Region: Caudate-Putamen, Prefrontal cortex, frontral cortex, neocortex, hippocampus, Whole Brain, ventral mid brain, striatum, cerebral cortex, left hippocampus, bilateral hippocampus, entorhinal cortex, embryonic brain, post natal brain Tissue: main olfactory epithelium, Granular Cell Layer					
GUIDELINE	Vital for gene expression analysis based on brain region/ tissue bias					
SOURCE	Image on Left (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-31458/samples/ Image on Right (GEO): http://www.ncbi.nlm.nih.gov/pmc/articles/PMC2481328/ Brain and tissue type information can either be explicitly mentioned in AE/GEO or have to be searched from research					
	paper linked to experiment. This information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper. In one of the experiment exact information about brain region was found in figure caption (Image on right) Sample_source_name Pre-Frontal contex of naïve AChE-5 mice Pre-Frontal contex of naïve AChE-5 mice Pre-Frontal contex of naïve FVB/N mice					
	Pre-Frontal contex of naïve PVB/N mice Pre-Frontal contex of naïve AChE-R mice Pre-Frontal contex of MPTP-exposed FVB/N mice ce Pre-Frontal contex of MPTP-exposed AChE-R mice nice Pre-Frontal contex of MPTP-exposed AChE-R mice pre-Frontal contex of MPTP-exposed AChE-R mice nice Pre-Frontal contex of MPTP-exposed AChE-R mice nice Pre-Frontal contex of MPTP-exposed AChE-S mice nice Pre-Frontal contex of MPTP-exposed AChE-S mice nice Pre-Frontal contex of MPTP-exposed AChE-S mice Caudate-Putamen of naïve FVB/N mice ce Caudate-Putamen of MPTP-exposed FVB/N mice ce Caudate-Putamen of MPTP-exposed FVB/N mice ce Caudate-Putamen of MPTP-exposed FVB/N mice ce Caudate-Putamen of naïve AChE-S mice Caudate-Putamen of naïve AChE-S mice nice Pre-Frontal contex of MPTP-exposed FVB/N mice ce Caudate-Putamen of MPTP-exposed FVB/N mice ce Caudate-Putamen of naïve AChE-S mice nice Pre-Frontal contex of MPTP-exposed FVB/N mice nice Pre-Frontal contex of MPTP-exposed FVB/N mice nice Pre-Frontal contex of MPTP-exposed FVB/N mice nice Pre-Frontal contex of MPTP-exposed AChE-S mice nice Pre-Frontal contex of MPTP-exp					



Field	superseries				
Description	If the current experiment is a superseries or not				
Examples	Yes/ No (binomial value)				
GUIDELINE	Gives information about superseries, this helps us to avoid duplicating the sample information				
SOURCE	Image (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE52024				
	Check in GEO experiment file if the sample is followed by the section of superseries. If superseries section is not found there, the current experiment is individual experiment and not a superseries. This SuperSeries is composed of the following SubSeries:				
	GSE52022 Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease (mRNA)				
	GSE52023 Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease (microRNA)				



superseries_id					
Is this current experiment a part of superseries, then we need to map it to its superseries ID. If the experiment itself is the superseries or individual experiment then the column should be left empty					
For GSE52022 experiment we include GSE52024 as superseries_id					
Direct reference to the experiment superseries via a unique ID					
Image on Left (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE52024					
Check in GEO experiment file if the samples are followed by the section of superseries information. If superseries section is not found there, the current experiment is individual experiment and has no superseries.					
This SuperSeries is composed of the following SubSeries:					
GSE52022 Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease (mRNA)					
GSE52023 Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease (microRNA)					
If this section is found, the series ID above would be superseries ID.					
Series GSE52024 Query DataSets for GSE52024					
Status Public on Dec 01, 2013 Title Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease					
Organism Mus musculus					
Experiment type Expression profiling by array Non-coding RNA profiling by high throughput sequencing					
Summary This SuperSeries is composed of the SubSeries listed below.					
Overall design Refer to individual Series					



Field	disease_type			
Description	Type of AD			
Examples	sporadic AD, familial			
GUIDELINE	Gives information to the cause of disease, if its genetically inherited or caused by other factors e.g. environment.			
SOURCE	Image (Research Paper): http://hmg.oxfordjournals.org/content/19/20/3959.long Image on Right (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-36232/samples/ Disease type information can mainly be found from research paper of individual experiment. Finally, we explored the hypothesis that loss of miRNA function could contribute to tau pathology in humans. To this end, we performed miRNA qRT-PCR from control and sporadic AD cases. Previous studies have shown that miR-16 is stably expressed in human tissues, including AD brain (25-27). Using this miRNA as internal control, we observed a significant decrease in miR-15a expression levels in AD brain when compared with healthy controls (Fig. 4C). There was no significant change in the expression of other miR-15 family members, including miR-15b and miR-195. We used as additional control the ubiquitously expressed let-7a, which is not affected in both groups. These results, together with previously published miRNA microarray data (27,28), strongly suggest that specific miR-15 family members, such as miR-15a, are affected in compromised brain displaying tau hyperphosphorylation.			



Field	rganism				
Description	Sample organism				
Examples	Mus musculus				
GUIDELINE	To obtain information about the species from where the sample collected				
SOURCE	Image on Top (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE20447 Image on Bottom (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-52024/				
	Organism information can be easily found in experiment GEO/ AE files.				
	Status Public on May 30, 2010				
	Title Neuronal microRNA response to Alzheimer's disease amyloid beta				
	Platform organisms Homo sapiens; Mus musculus; Rattus norvegicus				
	Sample organism Mus musculus				
	Experiment type Other				
	E-GEOD-52024 - Genome wide analysis of transcriptome and microRNAs in early stage of Alzheimer's disease				
	Status Released on 1 December 2013, last updated on 3 April 2014				
	Organism Mus musculus				
	Samples (16) Click for detailed sample information and links to data L found inside Mus musculus				



Field	rat_weight			
Description	Weight of rats			
Examples	00-150g			
GUIDELINE	To determine the metabolic changes			
SOURCE	Rat weight information can either be explicitly mentioned in AE/GEO or have to be searched from research paper linked to experiment. Gender information is usually found in Materials and Methods section of paper, figure or supplementary files provided in research paper.			

Field	platform_id					
Description	ID of the platform used for the experiment					
Examples	GPL339, GPL163	354				
GUIDELINE	Quick reference t	to the platform information of the platform	used. Support	s in probe annotation information		
SOURCE	Image (GEO): htt	Image (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE47036				
	Image on bottom	(AE): https://www.ebi.ac.uk/arrayexpres	s/experiments	s/E-GEOD-13691/		
	Platform ID information can be found in GEO experiment file. There can be different platforms used for difference Below screenshot shows information of 2 platforms for a single experiment. Each sample is mapped to the respondence platforms. E-GEOD-13691 - Long-term proteasomal inhibition in transgenic mic control reminiscent of brainstem neuropathology in Alzheimer patients and the state of the platforms of th					
	ZIP/Postal code 089	08	Samples (23)	Click for detailed sample information and links to data		
	Country Spai	in	Arrays (2)	A-AFFY-23 - Affymetrix GeneChip Mouse Expression Array MOE430A [MOE430A] A-AFFY-24 - Affymetrix GeneChip Mouse Expression Array MOE430B [MOE430B]		
	Platforms (1) GPL	.16354 Custom Illumina GoldenGate DNA methylation Beadchip	Protocols (9)			



Field	type_of_treatment (Animal models only)				
Description	Information about the type of treatment used on mice/rat				
Examples	MPTP treated, untreated (if the sample is not treated), treated with Kainic acid, treated with saline or vehicle, rasagiline, given				
	- 1	ΓP, water treated (control), PBS mixed with 18 mix of ethanol (Sigma-Aldrich, St Louis, MO, USA) and			
	neobee® M-5 oil (Stepan Company, Northfield, IL, USA) as vehicle.				
GUIDELINE		tion of correlation between type of treatment used to induce disease in mice/rat, or treatment of			
	induced disease				
SOURCE	Image (GEO): http:/	/www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM862523			
		ly read especially the supplementary files to find exact treatment type information. Sometimes, it can GEO (in treatment protocol section of sample file in GEO) and AE experiment files. Query DataSets for GSE35138			
	Status	Public on Jan 18, 2012			
	Title	Gene expression data from thalamic regions of MPTP-intoxicated mouse brain by acupuncture			
	Organism	Mus musculus			
	Experiment type	Expression profiling by array			
	Summary	Acupuncture stimulations at GB34 and LR3 inhibit the reduction of tyrosine hydroxylase in the nigrostriatal dopaminergic neurons in the parkinsonism animal models. Especially, behavioral tests showed that acupuncture			



Field	physical_injury (Animal models only)					
Description	Information ab	Information about the physical injury done on mice/rat (either for generation of mice/rat model or to assess injury as treatment				
Examples	acupoints acup	acupoints acupuncture-treated, ischemia/reperfusion (I/R) injury 2h, Sham control, nonacupoints acupuncture-treated				
GUIDELINE	Identification	of the process	s used to induce disease in	mice/rat through injury		
SOURCE	Image on Left (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSE35138					
	Image on Righ	nt (AE): https:/	//www.ebi.ac.uk/arrayex	oress/experiments/E-GEOD-23162/samples/		
	Information about type of physical injury is mostly found in materials and methods section of paper. However, entire should be thoroughly read especially the supplementary files to find exact injury method information. Sometimes, it also be found from GEO (in treatment protocol section of sample file in GEO) and AE experiment files.					
		,				
	2h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	2h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	2h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	2h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	8h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	8h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	8h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	8h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	24h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	24h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	24h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	24h	Gpx1-/-	ischemia/reperfusion (I/R) injury			
	Ctrl	Gpx1-/-	sham			



Field	dosage (Animal models only)				
Description	Information about the amount of treatment, at what interval and for what interval done on mice/rat				
Examples	MPTP treated: intraperitoneally injected with saline 0.9% (100 µl) once daily for four weeks				
	Saline treated: 15mg/kg in 0.2 ml volume Saline treated_control				
	Water: recieved Water for 14 days orally				
GUIDELINE	Relevant to obtain the amout of certain substance used to induce or treat the disease in animal models				
SOURCE	Image on Left (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM26703				
	Image on Right (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-36232/samples/				
	Information about dosage is mostly found in materials and methods section of paper. However, entire paper should be thoroughly read especially the supplementary files to find exact dosage information. Sometimes, it can also be found from GEO (in treatment protocol section of sample file in GEO) and AE experiment files.				
	Treatment protocol While mice in the control group (n=9) were intraperitoneally injected with saline 0.9% (100 µl) once daily for four weeks, the mice in the MPTP group (n=9) were intraperitoneally injected with MPTP-HCl (20 mg/kg of free base) dissolved in saline 0.9% (100 µl) at twenty-four hr intervals for four weeks to produce the sustained chronic model of parkinsonism. Acupuncture was performed by hand two hr after the first MPTP injection and then at forty-eight hr intervals (fourteen total sessions). The acupuncture procedure (acupoint GB34 (Yanglingquan) and acupoint LR3 (Taichong) for acupoints; both sides of the hips for non-acupoints) was performed as previously reported. Mice in the acupoints group were immobilized by hand two hr after MPTP administration. Acupuncture needles were inserted bilaterally to depths of 1 mm at acupoint LR3 and 3 mm at acupoint GB34, and then turned at a rate of two spins per sec for fifteen sec as reported in a previous study. For the non-acupoints group, the needles were inserted to depths of 3 mm at both sides of the hips rather than at the acupoints GB34 and LR3 for the acupoints group, and then the same procedures were performed as with the acupoints.				



Field	functional_effect			
Description	Functional effects observed in microarray, sequencing or other experiment as a result of physical injury or chemical treatment			
Examples	mouse affected by MPTP and acupuncture only at the acupoints, are responsible for exerting in the striatal regions the inhibitory			
	effect of acupuncture at the acupoints on MPTP-induced striatal degeneration.: Acupuncture at acupoints GB34 and LR3 has been			
	reported to inhibit nigrostriatal degeneration in parkinsonism models, yet the genes related to this preventive effect of acupuncture			
	on the nigrostriatal dopaminergic system remain elusive. SMARCA4 gene was downregulated in mTLE patients.			
GUIDELINE	Correlation between treatment and gene expression changes seen in experiment by the authors			
SOURCE	Functional effects is a free text information field where different functional effects observed in sample before and after			
	treatment observed is included.			
	Functional effect can be found from result, discussion, conclusion or other analytical sections of research paper.			
	Different functional effect are separated by a ":" (as seen in example above)			
	Functional effects field can also include external links or supplementary files if information about gene expression is large.			



Field	behaviour				
Description	Physical traits seen in mice/rat before and/or after treatment. Behavioural changes that can be notified in diseased patient during				
	the experiment				
Examples	behavioral traits in the mice/rat are akin to PD patients, deficits in fear conditioning, central breathing dysfunction, altered breathing patterns, high respiratory rate, increased duration of inspiration, results of Active Avoidance Learning test, results of novel object recognition test, results of the accelerating rotarod test, results of the grip strength test, results of the test of hanging from an inverted cage lid, overall activity in the novel open field arena, Y maze test, PA(Passive avoidance paradigm) assessment, put NA if not available. Tremor (shaking of hands)				
GUIDELINE	LINE Gives correlation between behaviour change and gene expression changes.				
SOURCE	Image (Reference paper): http://www.ncbi.nlm.nih.gov/pmc/articles/PMC3319057/				
	Behaviour information would rarely be explicitly mentioned in AE/GEO so it has to be searched from research paper linked				
	to experiment. Behaviour information is usually found supplementary tables provided with the research paper. It would be				
	difficult to find behaviour information and demands reading research paper thoroughly.				
	two different toxicological mouse models of PD created using 1-methyl-4-phenyl-				
	1,2,3,6-tetrahydropyridine (MPTP) and methamphetamine (METH). MPTP has been				
	shown to induce Parkinsonism in mice with recapitulation of the hallmark cellular				
	pathology, death of dopaminergic neurons in the SN, which decreases DA input to the				
	target organ, the striatum. 12,13 MPTP probably acts through the inhibition of oxidative				
	phosphorylation in dopaminergic neurons. ¹⁴ METH-treated mice also exhibit cell				
	death of dopaminergic neurons in the SN, and although the parallels to the human				
	disease are less strong, the behavioral traits in the mice are akin to PD patients. 15,16 METH stimulates catecholamine release, although the mechanisms by which large				
	doses result in a PD model are unclear. 17,18				
	doses result in a 1 D model are unclear.				
	1				



Field	mouse_name or rat_name				
Description	Vendor/ commercial name of mouse used				
Examples	C57/Bl6, C57BL/6-129, crossed Tg2576 APP transgenic mice/rat with PS1 M146V homozygous (PS1M146V/M146V) mice/rat: Alzheimers Disease, B6/SJL, B6/SJL (for hemizygous 5xFAD mice/rat) and C57BL/6J (for eIF2α+/S51A mice/rat), FVB/N mice/rat, Sprague-Dawley rat. Put NA if not available				
GUIDELINE	Relates the mouse used with gene expression changes. This could support in understanding the effects specific to a mouse model				
SOURCE	Image on Left (AE): https://www.ebi.ac.uk/arrayexpress/experiments/E-GEOD-47036/samples/ Image on Right (GEO): http://www.ncbi.nlm.nih.gov/geo/query/acc.cgi?acc=GSM1165975				
	thoroughly	read espec wth protoc	ially the supple	ound in materials and methods section of paper. However, entire paper should be ementary files to find exact mouse nformation. Sometimes, it can also be found from ample file in GEO) and AE experiment files. Mouse breeding information is also	
	C57BL/6-129 C57BL/6-129 C57BL/6-129	3TG 3TG 3TG	Status Title Sample type	Public on Sep 19, 2013 APP/PS1M146V/+	
	C57BL/6-129 C57BL/6-129	3TG APP/PSEN1		APP/PS1M146V/+ Mus musculus disease state: alzheimer's disease tissue: bilateral hippocampus genotype/variation: APP/PS1M146V/+ No treatment APP/PS1M146V/+ mice were generated by crossed Tg2576 APP transgenic mice with PS1 M146V homozygous (PS1M146V/M146V) mice. All mice were feed and housed according to standard procedures. total RNA	
	C57BL/6-129 C57BL/6-129	APP/PSEN1	Source name Organism Characteristics Treatment protocol Growth protocol Extracted molecule		
	C57BL/6-129 C57BL/6-129	APP/PSEN1 APP/PSEN1			
	C57BL/6	WT WT			
	C57BL/6 C57BL/6	WT WT			
	C57BL/6	WT			