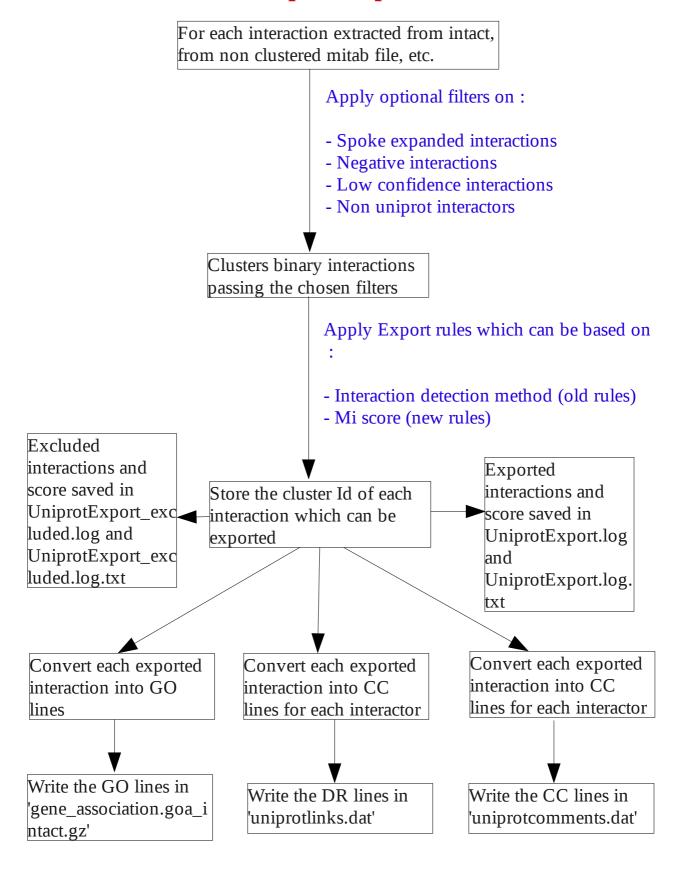
General Uniprot Export Data flow



DR line export specifications

Each DR line must have the following format:

DR IntAct; UNIPROT; INTERACTIONS.

Where:

UNIPROT: is the uniprot AC of a protein in Intact. Isoforms and feature chain ids are not accepted and must be remapped to the parent uniprot AC. It is a mandatory field.

INTERACTIONS: is the number of distinct PPI interactions in IntAct involving this protein. This number must be strictly superior to 0.

- Old Uniprot Export specifications: Only true positive binary interactions involving uniprot proteins must be taken into account. The low confidence interactions must be ignored (interactions with author confidence not matching the 'uniprot-dr-export' annotation of the experiment or interactions with 'uniprot-dr-export' = 'no').
- New Uniprot Export specifications: Spoke expanded binary interactions involving uniprot proteins must be taken into account as well as true binary interactions. Negative interactions are accepted. The low confidence interactions must be ignored (interactions with author confidence not matching the 'uniprot-dr-export' annotation of the experiment or interactions with 'uniprot-dr-export' = 'no').

CC line export specifications

1) Old Uniprot export specifications

Each CC line must have the following format:

For each uniprot protein:

```
AC UNIPROT_MASTER
CC -!- INTERACTION
```

For each binary interaction involving the protein:

- If self interactions (two participants having the same uniprot ac and stoichiometry = 0 or 1 for each participant OR one participant with stoichiometry = 2) and same organisms :

```
CC Self; NbExp=INTERACTIONS; IntAct=FIRST_INTACT, SECOND_INTACT;
```

- If self interactions and different organisms :

```
CC Self (xeno); NbExp=INTERACTIONS; IntAct=FIRST_INTACT, SECOND INTACT;
```

- If two different interactors and same organism :

```
CC SECOND_UNIPROT:GENE; NbExp=INTERACTIONS; IntAct=FIRST_INTACT, SECOND_INTACT;
```

- If two different interactors and different organisms :

```
CC SECOND_UNIPROT:GENE (xeno); NbExp=INTERACTIONS; IntAct=FIRST_INTACT, SECOND_INTACT;
```

//

Where:

UNIPROT_MASTER is the uniprot AC of a protein in Intact. Isoforms and feature chain ids are not accepted and must be remapped to the parent uniprot AC. It is a mandatory field and must be unique in a same file.

SECOND_UNIPROT is the uniprot AC of a protein in Intact interacting with the UNIPROT_MASTER. Isoforms and feature chain ids are accepted. It is a mandatory field and must be unique in a same CC line.

GENE is the gene name associated with the SECOND_UNIPROT. It is a mandatory field. If no gene names are available, first look at the locus name, then at the ORF name.

INTERACTIONS is the number of positive interaction evidences in IntAct involving this proteins. This number must be strictly superior to 0. Only evidences of true binary interactions involving uniprot proteins must be taken into account. The low confidence interactions must be ignored (interactions with author confidence not matching the 'uniprot-dr-export' annotation of the experiment or interactions with 'uniprot-dr-export' = 'no'). All evidences must have interaction detection methods passing export rules (described later).

FIRST_INTACT is the intact AC of the protein in Intact which is associated with the UNIPROT_MASTER. It is a mandatory field.

SECOND_INTACT is the intact AC of the protein in Intact which is associated with the SECOND_UNIPROT. It is a mandatory field.

2) New Uniprot export specifications (copy of the document from Uniprot)

The CC line topic INTERACTION conveys information about binary protein-protein interactions which is derived from the IntAct database.

A description of its current format is available in the UniProtKB User Manual.

This format will be modified to include further supplementary experimental information which will allow the user to more easily gauge the level of support or confidence for any given protein interaction. The new format will also allow the representation of isoform- and chain-specific interactions (such as those involving specific portions of viral polyproteins) as well as negative interactions.

Each binary interaction will be represented by a block of several lines:

- * The first line describes whether the experimental evidence indicates that the two proteins interact or do not interact and provides cross-reference(s) to other databases storing the experimental data.
- * The second line is an optional comment about the interaction.

- * The third and fourth lines identify the interacting proteins.
- * The fifth and following line(s) (where present) contain supplementary experimental information:
 - the type of interaction that was inferred from the experiment(s)
 - the experimental method used to demonstrate the interaction
 - the source of the data (literature reference)

During data import from IntAct, experiments which share identical interaction type and experimental method are condensed into one line (see examples).

The new format of INTERACTION in the flat file is:

Where:

- * Interact= indicates if there is experimental evidence that the two proteins interact (Yes) or do not interact (No) under the experimental conditions described in the cited publication(s).
- * <u>db:db id</u>: Database cross-reference, consisting of database name and unique database identifier.
 - Xref= contains currently only one cross-reference to the IntAct database. May contain cross-references to other protein-protein interactions databases in the future. It can be composed of Intact Ac1, Intact Ac2 which is the identifier of the binary interaction.
 - Source= contains currently only cross-references to the PubMed database. May contain cross-references using DOIs in the future.
- * Comment= contains additional information concerning the interaction (like PTMs, subcellular location, etc.).

- * Protein1= and Protein2= describe the two interacting proteins:
- label: Biologically meaningful label or name for the protein, or chain thereof. This field will generally take the value of the gene name (as shown in the GN line Name= field of the UniProtKB entry). When no gene name is available then the value of the corresponding OrderedLocusNames= or ORFNames= field of the GN line of the UniProtKB entry may be used. In the absence of any available gene designation, this field will be filled with a dash "-".
 If the protein contains multiple chains (e.g. a viral polyprotein), and the interaction involves only one of these chains, then the chain name (from the FT line) is displayed and the identifier which follows in square brackets will contain the FTId (unique feature identifier) for that chain.
- id: UniProtKB accession number or isoform identifier (IsoId= field).
- ft_id: UniProtKB feature identifier (FTId= field).
- ft_id: UniProtKB feature identifier (FTId= field). If Protein2 is derived from a distinct species, that is indicated in the Organism= field:
 - tax_name: Scientific name of the organism.
 - tax_id: NCBI taxonomy database identifier of the organism.
- * InteractionType= describes the interaction type:
- type: PSI-MI controlled vocabulary for interaction type.
- * Method= describes the interaction detection method:
- method: PSI-MI controlled vocabulary for interaction detection method.

Note: Perl-style multipliers indicate whether a pattern (as delimited by parentheses) is optional (?), may occur 0 or more times (*), or 1 or more times (+). Alternative values are separated by a pipe symbol (|).

Examples:

Experimental information deriving from different publications is condensed:

CC -!- INTERACTION:
 CC Interact=Yes; Xref=IntAct:EBI-359343,EBI-79792;
 CC Comment=FANCD2 is ubiquitinated (PubMed:15199141).
 CC Protein1=FANCD2 [Q9BXW9];
 CC Protein2=BRCA2 [P51587];
 CC InteractionType=association; Method=anti bait coimmunoprecipitation;
 Source=PubMed:18212739;

- CC InteractionType=physical association; Method=chromatography technology; Source=PubMed:15115758;
- CC InteractionType=physical association; Method=coimmunoprecipitation; Source=PubMed:15115758, PubMed:15199141;

Positive and negative isoform-specific interactions:

- CC -!- INTERACTION:
- CC Interact=Yes; Xref=IntAct:EBI-1018629,EBI-1569435;
- CC Protein1=PIM1 [P11309-1];
- CC Protein2=ABCG2 [Q9UNQ0];
- CC InteractionType=physical association; Method=anti bait coimmunoprecipitation; Source=PubMed:18056989;
- CC InteractionType=physical association; Method=anti tag coimmunoprecipitation; Source=PubMed:18056989;
- CC InteractionType=physical association; Method=pull down; Source=PubMed:18056989;
- CC InteractionType=physical association; Method=two hybrid;
- Source=PubMed:18056989;
- CC Interact=No; Xref=IntAct:EBI-1018633,EBI-1569435;
- CC Protein1=PIM1 [P11309-2];
- CC Protein2=ABCG2 [Q9UNQ0];
- CC InteractionType=physical association; Method=pull down; Source=PubMed:18056989;

Non-physiological heterologous interaction between rat and human proteins:

- CC -!- INTERACTION:
- CC Interact=Yes; Xref=IntAct:EBI-296087,EBI-1636616;
- CC Protein1=AKT1 [P31749];
- CC Protein2=Arrb2 [P29067]; Organism=Rattus norvegicus

[NCBI_TaxID:10116];

- CC InteractionType=physical association; Method=anti bait coimmunoprecipitation; Source=PubMed:18191226;
- CC InteractionType=physical association; Method=pull down; Source=PubMed:18191226;

Physiological heterologous chain-specific virus-host interaction: virus entry

- CC -!- INTERACTION:
- CC Interact=Yes; Xref=IntAct:EBI-710918,EBI-515315;
- CC Protein1=Non-structural protein 5A [Q9WMX2:PRO_0000037551];
- CC Protein2=FYN [P06241]; Organism=Homo sapiens [NCBI_TaxID=9606];
- CC InteractionType=physical association; Method=anti bait coimmunoprecipitation; Source=PubMed:14993658;

Physiological heterologous chain-specific host-virus interaction: host entry

- CC -!- INTERACTION:
- CC Interact=Yes; Xref=IntAct:EBI-515315,EBI-710918;
- CC Protein1=FYN [P06241];
- CC Protein2=Non-structural protein 5A [Q9WMX2:PRO_0000037551];

Organism=Hepatitis C virus genotype 1b (isolate Con1) [NCBI_TaxID=333284];

CC InteractionType=physical association; Method=anti bait coimmunoprecipitation; Source=PubMed:14993658;

GO line export specifications

Each binary interaction exported in the CC lines will be exported in the GO lines.

Each GO line must have the following format:

If two different interactors :

UniProt **FIRST_UNIPROT** GO:0005515 PMID:**Id1**|**Id2**| **Id3**|... IPI UniProt:**SECOND_UNIPROT**IntAct

– If self interaction :

UniProt FIRST_UNIPROT GO:0042802 PMID:Id1|Id2|
Id3|.. IPI UniProt: SECOND_UNIPROT
IntAct

Where:

FIRST_UNIPROT is the uniprot AC of a protein in Intact. Feature chain ids are not accepted and must be remapped to the parent uniprot AC. It is a mandatory field.

SECOND_UNIPROT is the uniprot AC of a protein in Intact interacting with the FIRST_UNIPROT. Feature chain ids are not accepted and must be remapped to the parent uniprot AC. It is a mandatory field.

SECOND_UNIPROT is the uniprot AC of a protein in Intact interacting with the FIRST_UNIPROT. Feature chain ids are not accepted and must be remapped to the parent uniprot AC. It is a mandatory field.

Id1|**Id2**|**Id3**|... are the pubmed Ids which are referring to the interaction. It is a mandatory field and must not be empty.

Export rules based on the detection method

If the binary interaction has at least one detection method with an 'uniprot-dr-export' annotation = 'yes', the binary interaction is exported.

If the binary interaction has only detection method with an 'uniprot-dr-export' annotation = 'no', the binary interaction is NOT exported.

If the binary interaction has only detection method with an 'uniprot-dr-export' annotation = 'CONDITION', the binary interaction is exported only if the condition is respected for at least one detection method (Ex 2 hybrid method has a condition of 2 to be exported. If two '2-hybrid' methods are found for this interactions, the condition is respected and the interaction can be exported).

List of Detection methods

term: sandwich immunoassay

yes

term: potassium permanganate footprinting

yes

term: luminescence based mammalian interactome mapping

yes

term: lambda repressor two hybrid

yes

term: tandem affinity purification

export: yes

term: 2h fragment pooling: two hybrid fragment pooling approach

id: MI:0399 export: 2

term: adenylate cyclase: adenylate cyclase complementation

id: MI:0014 export: 2

term: affinity chrom: affinity chromatography technologies

id: MI:0004 export: no

term: affinity techniques: affinity technologies

id: MI:0400 export: no

term: anti bait coip: anti bait coimmunoprecipitation

id: MI:0006 export: yes

term: anti tag coip: anti tag coimmunoprecipitation

id: MI:0007 export: yes

term: array technologies

id: MI:0008 export: no

term: bacterial display

id: MI:0009 export: yes

term: beta galactosidase: beta galactosidase complementation

id: MI:0010 export: 2

term: beta lactamase: beta lactamase complementation

id: MI:0011 export: 2

term: biochemical

id: MI:0401 export: no

term: biophysical id: MI:0013 export: no

term: bret: bioluminescence resonance energy transfer

id: MI:0012 export: yes

term: cd: circular dichroism

id: MI:0016 export: no

term: chromatography: chromatography technologies

id: MI:0091 export: no

term: coip: coimmunoprecipitation

id: MI:0019 export: yes

term: collagen film assay

id: MI:0513 export: yes

term: coloc fluoresc probe: colocalization by fluorescent probes cloning

id: MI:0021 export: no

term: comigration in gel: comigration in non denaturing gel electrophoresis

id: MI:0404 export: no

term: competition binding

id: MI:0405 export: yes

term: complementation: protein complementation assay

id: MI:0090

export: 2

term: cosedimentation

id: MI:0027 export: yes

term: crosslink: cross-linking studies

id: MI:0030 export: yes

term: cytoplasmic compl: cytoplasmic complementation assay

id: MI:0228 export: 2

term: deacetylase assay

id: MI:0406 export: yes

term: density sedimentatio: cosedimentation through density gradients

id: MI:0029 export: yes

term: dhfr reconstruction: dihydrofolate reductase reconstruction

id: MI:0111 export: 2

term: display technologies

id: MI:0034 export: no

term: dls: dynamic light scattering

id: MI:0038 export: no

term: electron microscopy

id: MI:0040 export: yes

term: electron resonance

id: MI:0043 export: yes

term: elisa: enzyme linked immunosorbent assay

id: MI:0411 export: yes

term: endor: electron nuclear double resonance

id: MI:0041 export: yes

term: enzymatic studies

id: MI:0415 export: no

term: epr: electron paramagnetic resonance

id: MI:0042 export: yes

term: experimental

id: MI:0045 export: no

term: facs: fluorescence-activated cell sorting

id: MI:0054 export: no

term: far western blotting

id: MI:0047 export: yes

term: fcs: fluorescence correlation spectroscopy

id: MI:0052 export: yes

term: filamentous phage: filamentous phage display

id: MI:0048 export: yes

term: filter binding

id: MI:0049 export: yes

term: fluorescence: fluorescence technologies

id: MI:0051 export: no

term: fluorescence imaging: fluorescence microscopy

id: MI:0416 export: no

term: fluorescence spectr: classical fluorescence spectroscopy

id: MI:0017 export: yes

term: footprinting id: MI:0417 export: yes

term: fps: fluorescence polarization spectroscopy

id: MI:0053 export: yes

term: fret: fluorescent resonance energy transfer

id: MI:0055 export: yes

term: gallex: lex-a dimerization assay

id: MI:0369 export: 2

term: gfp complementation: green fluorescence protein complementation assay

id: MI:0229 export: 2

term: gtpase assay id: MI:0419

export: yes

term: htrf: homogeneous time resolved fluorescence

id: MI:0510 export: yes

term: imaging techniques

id: MI:0428 export: no

term: in gel kinase assay: in-gel kinase assay

id: MI:0423 export: yes

term: in gel phosphatase: in gel phosphatase assay

id: MI:0514 export: yes term: inference id: MI:0362 export: no

term: inferred by author

id: MI:0363 export: no

term: inferred by curator

id: MI:0364 export: no

term: ion exchange chrom: ion exchange chromatography

id: MI:0226 export: no

term: itc: isothermal titration calorimetry

id: MI:0065 export: yes

term: kinase htrf: kinase homogeneous time resolved fluorescence

id: MI:0420 export: yes

term: kinase spa: kinase scintillation proximity assay

id: MI:0425 export: yes

term: lambda phage: lambda phage display

id: MI:0066 export: yes

term: light microscopy

id: MI:0426 export: no

term: light scattering

id: MI:0067 export: no

term: mappit: mammalian protein protein interaction trap

id: MI:0231 export: 2

term: membrane compl: membrane bound complementation assay

id: MI:0230 export: 2

term: methyltransferase as: methyltransferase assay

id: MI:0515

export: yes

term: molecular sieving

id: MI:0071 export: no

term: mrna display

id: MI:0073 export: yes

term: ms of complexes: mass spectrometry studies of complexes

id: MI:0069 export: yes

term: nmr: nuclear magnetic resonance

id: MI:0077 export: yes

term: peptide array

id: MI:0081 export: yes

term: phage display

id: MI:0084 export: yes

term: phosphatase assay

id: MI:0434 export: yes

term: phosphatase htrf: phosphatase homogeneous time resolved fluorescence

id: MI:0509 export: yes

term: pisa: protein in situ array

id: MI:0092 export: yes

term: protease assay

id: MI:0435 export: yes

term: protease htrf: protease homogeneous time resolved fluorescence

id: MI:0511 export: yes

term: protein array

id: MI:0089 export: yes

term: protein crosslink: protein cross-linking with a bifunctional reagent

id: MI:0031 export: yes

term: protein kinase assay

id: MI:0424 export: yes

term: pull down id: MI:0096 export: yes

term: radiolabeled acetate: deacetylase radiometric assay

id: MI:0508 export: yes

term: radiolabeled methyl: methyltransferase radiometric assay

id: MI:0516 export: yes

term: reverse phase chrom: reverse phase chromatography

id: MI:0227 export: no

term: reverse rrs: reverse ras recruitment system

id: MI:0097 export: 2

term: ribosome display

id: MI:0098 export: yes

term: saturation binding

id: MI:0440 export: yes

term: seldi chip: proteinchip(r) on a surface-enhanced laser desorption/ionization

id: MI:0095 export: no

term: sem: electron tomography

id: MI:0410 export: no

term: sls: static light scattering

id: MI:0104 export: no

term: solution sedimentati: cosedimentation in solution

id: MI:0028 export: yes

term: spa: scintillation proximity assay

id: MI:0099 export: yes

term: spr: surface plasmon resonance

id: MI:0107 export: yes

term: t7 phage: t7 phage display

id: MI:0108 export: yes

term: tem: transmission electron microscopy

id: MI:0020 export: yes

term: toxcat: tox-r dimerization assay

id: MI:0370 export: 2

term: transcription compl: transcriptional complementation assay

id: MI:0232 export: 2

term: two hybrid id: MI:0018 export: 2

term: two hybrid array

id: MI:0397 export: 2

term: two hybrid pooling: two hybrid pooling approach

id: MI:0398 export: 2

term: ub reconstruction: ubiquitin reconstruction

id: MI:0112 export: 2

term: x-ray: x-ray crystallography

id: MI:0114 export: yes

term: yeast display

id: MI:0115 export: yes

term: zymography

id: MI:0512 export: yes

Export rules based on the mi score

The binary interaction must have a score superior to a Threshold value (0.43 or 0.40 for the moment).

The binary interaction must have at least one evidence of true binary interaction which is not COLOCALIZATION.