# Rfam 15: RNA families database in 2025

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### **Abstract**

The Rfam database, a widely-used repository of non-coding RNA (ncRNA) families, has undergone significant updates in release 15.0. This paper introduces major improvements, including the expansion of Rfamseq to 26,106 genomes, a 76% increase, incorporating the latest UniProt reference proteomes and additional viral genomes. Sixty-five RNA families were enhanced using experimentally determined 3D structures, improving the accuracy of consensus secondary structures and annotations. R-scape covariation analysis was used to refine structural predictions in 26 families. Gene Ontology and Sequence Ontology annotations were comprehensively updated, increasing GO term coverage to 75% of families. The release adds 14 new Hepatitis C Virus RNA families and completes microRNA family synchronisation with miRBase, resulting in 1,603 microRNA families. New data types, including FULL alignments, have been implemented. Integration with APICURON for improved curator attribution and multiple website enhancements further improve user experience. These updates significantly expand Rfam's coverage and improve annotation quality, reinforcing its critical role in RNA research, genome annotation, and the development of machine learning models. Rfam is freely available at <a href="https://rfam.org">https://rfam.org</a>.

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#### Introduction

 The Rfam database was established in 2002 (1) in order to provide a central repository of non-coding RNA (ncRNA) families for genomic annotations. Each family is represented by three key components: (i) a multiple sequence alignment called the SEED alignment that contains aligned sequences of homologous RNA sequences that share a consensus secondary structure, (ii) a covariance model (CM) built using the Infernal software (2) that was trained on the SEED alignment, and (ii) a set of matches that were found using the covariance model, called the set of FULL hits. The families are built manually by Rfam curators using scientific literature or alignments submitted by the community.

For each model the curators search for homologs in a sequence database, Rfamseq, and select a bit score threshold, called the gathering threshold, that separates homologous sequences from unrelated or more distantly related sequences, taking into account phylogenetic distribution of hits. The cutoff can be used by Infernal to report only sequences that score higher than this threshold when annotating genomes or searching in sequence databases.

Each major release of the Rfam database, such as 14.0 or 15.0, corresponds to a new version of Rfamseq. Prior to Rfam 13.0, Rfamseq was based on a set of sequences from the ENA database (3). However, due to the explosive growth in ENA, in Rfam 13.0 Rfamseq transitioned to a representative and reduced redundancy set of genomes produced by the UniProt team (4) for the Reference Proteomes dataset (5). The representative proteomes from UniProt are mapped to their corresponding genomes and the genomic sequence is analysed by Rfam (5). This provides users with a representative collection of genomes which have comprehensive ncRNA and protein annotations.

Rfam has been widely adopted by various scientific communities, with uses ranging from the annotation of small non-coding RNAs (ncRNAs) in genomic resources like Ensembl (6), to serving as a dataset for training machine learning models such as AlphaFold 3 (7). Additionally, Rfam is frequently used as a key reference for known non-coding RNAs. To ensure it remains current and valuable to the scientific community, we continuously review, update, and enhance the Rfam families.

In release 15.0 we have updated and expanded Rfamseq, improved families using 3D structures and R-scape (8), improved Gene Ontology (9) and Sequence Ontology (10) annotations and completed the synchronisation of miRBase (11) microRNAs into Rfam.

# New Rfamseq database

Rfam 15 is based on the 2024\_03 release of UniProt reference proteomes that includes 23,158 genomes (see Figure 1 showing the taxonomic distribution). We attempted to download all components of the genomes used to build the proteome set. In some cases, the specified genome was replaced by a newer version, which was used instead. In a very few cases, less than 50, all versions of the genome were either outdated or removed. In those cases we did not fetch any genome.

In addition to the genomes in the UniProt set, we also fetched all viral genomes from the protein information resource (12) at 75% percent identity for an additional 2,985 viral genomes. This set was added because during our viral work, described below, we found that the UniProt reference proteome work under reports the diversity of viral genomes. This has led to some viral Rfam families having very few or no FULL hits. Rfam would prefer that all families have at least one match outside of the SEED sequences. While using an additional set of viral genomes leads to a large increase in the number of genomes, the total size of Rfamseq is dominated by cellular organisms as <1% of the nucleotides come from a viral genome. Table 1 shows the comparison of Rfamseq in the last major release Rfam 14.0 vs Rfam 15.0.

Table 1. The number of genomes in Rfamseq corresponding to release 14.0 and 15.0. The size on disk is the uncompressed size of all sequences in Rfamseq.

Kingdom	Rfamseq v14	Rfamseq v15
Total sequence length	3.68e11	1.46e12
Number of species	14,774	26,106
Eukaryotes	1,057	2,613
Archaea	418	370
Bacteria	7,808	9,601
Viruses	5,491	13,552
Number of Rfam hits	2,897,296	10,736,534
Size on disk	170 GB	772 GB

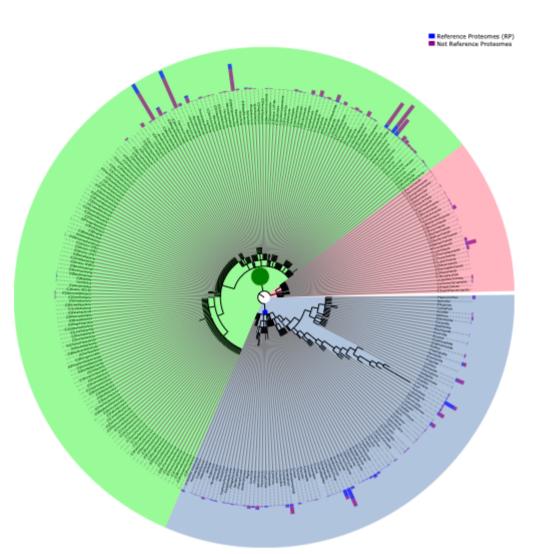


Figure 1. The taxonomic distribution of cellular organisms in Rfamseq 15 organised by the phylogenetic kingdom. Shown on the outside is the number of reference and non-reference proteomes in each lineage. Each coloured section indicates the kingdom with bacteria (green), archaea (pink) and eukaryotes (blue).

Once Rfamseq was updated all families were searched against the updated database and the matches were collected. Many older or larger families have not been rescanned in years due to technical pipeline limitations, for example, the 5S rRNA (RF00001) has not had its matches updated since 2014.

Once all families were updated, we compared the matches of families in 14.3, which was the last published version, and 15.0 to determine how Rfam has changed. Of the 3,431 families in common between 14.3 and 15.0, 98 had no FULL hits in 14.3 and when moving to 15.0, 23 of these families gained at least one hit while 21 lost all hits, leading to 96 families in 15.0 without FULL hits. Families without FULL hits occur because either the curator-selected gathering threshold of the family is too high, or if Rfamseq does not contain any similar sequences. Future work will look to see if these families require updating the threshold or if additional genomes need to be added to Rfamseq.

Of the 3,302 families with hits in both releases, the average family grew by 166%, with 2,335 gaining at least one hit, 547 losing hits and 420 with no change in the number of hits.

Examining the families with an increase shows that there are 26 families with a greater than 10x growth. These families are primarily microRNA families matching to plant genomes. While, as described below, these families have been synchronised with miRBase, we are revisiting these families to see if they need additional future updates in the light of these additional hits, or if the genomes are poorly assembled and should be excluded from Rfamseq.

Overall the update to Rfamseq has led to a considerable change in all Rfam families. This will provide the community with a consistent and up-to-date dataset to reuse. Future updates to Rfamseq will seek to maximise the coverage of known organisms.

# Improving existing Rfam families

#### Using RNA 3D structures to revise Rfam secondary structures

Rfam families are based on multiple sequence alignments annotated with consensus secondary structures that indicate base pairing. The accuracy and completeness of the consensus secondary structure is of critical importance as it guides the SEED alignment, informs the Infernal CM, and is used for training and benchmarking of software for RNA 2D and 3D structure prediction, including AlphaFold 3 (7) and R-scape (8).

Most of the Rfam families with known 3D structure were created prior to the 3D structure determination using less accurate, predicted secondary structures. For example, while the FMN riboswitch family (RF00050) correctly captured 5 helices of the FMN riboswitch, it did not include 1 helix, 2 pseudoknots, and several key base pairings (Figure 2B). As such, one major goal of this project was to include the pseudoknots which are observed in 3D structures into the Rfam families.

To aid the Rfam curators with updating Rfam alignments an automated pipeline was developed to map RNA sequences and secondary structures of experimentally determined RNA 3D structures to Rfam SEED alignments. Every week Rfam families are mapped to the latest set of RNA chains from PDB (13) using the Infernal *cmscan* program. For each Rfam family with one or more matching structures, the sequences and secondary structures are iteratively added to the alignment in the Stockholm format using Infernal's *cmalign* program. The base pairing information, as determined by FR3D (14), is also included in the alignment as an additional GR annotation line (one per 3D structure). The pipeline is automatically executed weekly and the Rfam curators receive notifications of newly mapped structures (Figure 2A).

The resulting alignments undergo manual review to determine whether the family consensus secondary structure is consistent with the base pair information from PDB structures. The structure and annotated based pairs are always kept in the alignment, even in cases where the structure is not used to update the consensus secondary structure, e.g. low resolution structure or non-canonical base pairs. Additional annotations are also included in the Stockholm files, such as RNA structural elements. For example P1, P2, and P3 domains are included in the Pistol ribozyme RF02679, while the SAM riboswitch (RF00162) now includes a kink-turn annotation.

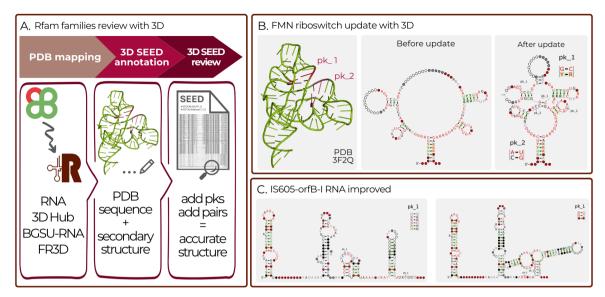


Figure 2. A) Pipeline for reviewing Rfam families using experimentally determined RNA 3D structures from PDB. B) Example of the FMN riboswitch (RF00050) before and after improvements with a 3D structure. Improvements included adding 2 pseudoknots and creating a missing helicees. C) A summary of how the IS605-orfB-I RNA (RF03065) family was reviewed and improved using R-scape.

Our pipeline detects 143 Rfam families which match at least one 3D structure. We began updating families with 3D structures in release 14.5 and since then Rfam has updated 65 families, with 298 chains from 3D structures. This includes well known families such as the SAM riboswitch (RF00162), the FMN riboswitch (RF00050) and microRNA 16 precursor (RF00254). The updates included the addition of pseudoknots (PK) in 42 of the 65 families, of which 6 had 2 PK, and 1 had 3. These features include ligand binding sites for riboswitches (28 annotations), locations of well known motifs such kink-turns (8 annotations), and structural regions such as helices and domains (60 annotations). A detailed table showing which families were updated with which 3D structures and can also be found online at <a href="https://rfam.org/3d">https://rfam.org/3d</a>. Rfam curators work to keep families as up-to-date with the latest structures as possible, which has led to 11 families being updated more than once since release 14.5.

The review of structures is part of the ongoing curation process, and additional families will be improved in future releases. The pipeline is implemented in Python and is available at <a href="https://github.com/Rfam/rfam-3d-seed-alignments">https://github.com/Rfam/rfam-3d-seed-alignments</a> and the weekly updates are found in the file pdb\_full\_region.txt.gz located in the preview section of the FTP archive (<a href="http://ftp.ebi.ac.uk/pub/databases/Rfam/.preview/">http://ftp.ebi.ac.uk/pub/databases/Rfam/.preview/</a>).

# Using R-scape to improve Rfam consensus secondary structures

Since version 13.0 (5) the Rfam website included the results of the R-scape analysis (8) to help users evaluate consensus secondary structures for each Rfam family, as well as alternative secondary structures generated using the R-scape CaCoFold algorithm. CaCoFold evaluates all possible consensus secondary structures that are consistent with multiple

sequence alignment and can propose a structure maximising the number of base pairs with statistically significant covariation (15).

In release 14.9, the R-scape software was used to systematically review all Rfam families and identify those that could be enhanced using R-scape CaCoFold structures. We examined all families with an increased number of covarying base pairs and selected 26 families for updates. These families are listed in Table 2. Shown in Figure 2C is an example of a family, IS605-orfB-I, before and after improvement.

Table 2: Summary of the R-scape based improvements. This shows the families which were improved by using R-scape CaCoFold structures.

Family	Additional Covarying Basepairs
RF02033 (HEARO)	24
RF03065 (IS605-orfB-I)	14
RF03068 (RT-3)	8
RF03072 (raiA)	5
RF02969 (DUF3800-I)	4
RF01688 (Actino-pnp)	3
RF02004 (group-II-D1D4-5)	3
RF02005 (group-II-D1D4-6)	3
RF02913 (pemK)	3
RF03077 (RT-2)	3
RF03135 (L4-Archaeoglobi)	3
RF03144 (eL15-	
Euryarchaeota)	3
RF00062 (HgcC)	2
RF01731 (TwoAYGGAY)	2
RF01794 (sok)	2
RF02221 (sRNA-Xcc1)	2
RF02947 (cow-rumen-2)	2
RF03000 (LOOT)	2
RF03158 (L31-Actinobacteria)	2
RF01864	
(plasmodium_snoR21)	1
RF01867 (CC2171)	1
RF02944 (c4-2)	1
RF02968 (DUF3800-IX)	1

RF02987 (GA-cis)	1
RF03019 (RT-16)	1
RF03046 (Pseudomonadales-1)	1

# Gene and Sequence Ontology annotations

Rfam is commonly used as a source of functional information for non-coding RNAs. This takes several forms, from users reading our curated descriptions and Wikipedia articles to understand the role of an ncRNA, to using the Gene Ontology (GO) and Sequence Ontology (SO) annotations that Rfam curators provide to understand the role of an ncRNA. Rfam is the largest source of GO annotations with over 10 million sequences having an Rfam based GO annotation. In addition to providing information to human users, Rfam is also used in training Large Language Models (LLM). LLMs such as ChatGPT and Claude have clearly been trained on Rfam data and are able to output entries in the formats, Stockholm and DESC, that Rfam uses. Thus providing a completely new way for scientists to access the information found in Rfam.

The Gene Ontology is a resource to classify and provide functional information for biomolecules using structured annotations and ontologies (9). Similarly the Sequence Ontology provides an ontology and structured annotations for the types of biomolecules (10). Both resources are continuously updated to better reflect scientific knowledge. For example, since the last publication of Rfam the SO gained several terms specific to the location of rRNA, e.g. cytosolic\_rRNA (SO:0002343) and obsoleted the more generic rRNA terms.

Rfam provides GO and SO annotations for families, however, these annotations are not regularly reviewed to stay up-to-date with the latest changes in the ontologies. As an example, prior to 15.0 the rRNA families used obsoleted SO terms. For 15.0 we manually reviewed all annotations and updated them to better reflect changes in the GO and SO, as well as improved the specificity of annotations where possible.

We ensured that all families had at least one up-to-date SO term and the annotation was as specific as possible. For GO terms, we ensured the terms were up-to-date and added related terms where possible. A few examples of the changes we made were to 1) ensure all snoRNAs had a snoRNA specific SO term and included an RNA processing GO term; 2) add the sodium ion binding term to the NA+ riboswitch; and 3) updated families with the generic mature\_transcript SO term to the more specific ncRNA SO term. Of the 3,431 common families between 14.3 and 15.0 2,084 had a GO term in 14.3 while 2,426 did in 15.0. For all families in 15.0 we now have at least one GO term for 3,157, which covers 75% of all families, an increase from 60% in 14.3. These changes increased the total number of GO terms from 3,752 to 4,446. As these updates propagate through the community, we expect these changes to lead to millions more sequences annotated with functional information.

#### Viral RNAs

Starting with Rfam 14.3 we collaborated with the Marz group to develop a new workflow for viral RNA families (16). This workflow is based upon whole genome alignments of viruses and led to the creation of *Flaviviridae* and *Coronaviridae* families as described previously (16). Since the last publication we have continued to use this workflow and have added *Hepatitis C Virus* (HCV) (17) families. We included 14 new families and removed 4 outdated families. A schematic of the new families can be seen in Figure 3. These families include structures found in both the non-coding and coding regions of the viral alignments. All viral families organised by viral clade can be seen at <a href="https://rfam.org/viruses">https://rfam.org/viruses</a>. We plan to continue to use the viral pipeline and improve the representation of viral families in Rfam and expect to focus on HDV and pestivirus families in the near future.

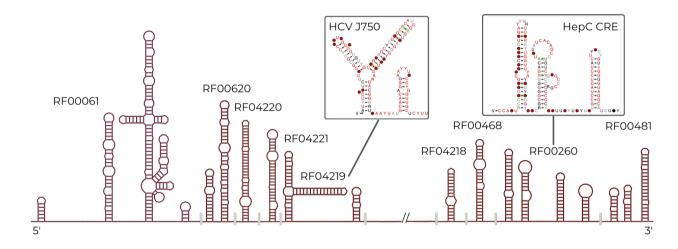


Figure 3: A schematic of the HCV viral families Rfam now contains. These families were built from whole genome alignments, as described in (17). The figure shows their relative locations in the HCV genomic alignment and shown in the inset are two of the new HCV families.

#### **MicroRNAs**

MicroRNAs are an important class of ncRNA that regulate gene expression in animals and plants, with many microRNAs being implicated in disease. For example, the mir-17~92 cluster (18) and mir-155 (19) are amongst a number of microRNAs which act as oncogenes. Understanding the evolutionary and family relationships of microRNAs across species allows the transfer of annotation, for example from model organisms to humans and vice versa.

Since early releases, Rfam has included microRNA families, e.g let-7 (20) was deposited in 2002 for Rfam 1.1. However, they were not subject to regular review and were not coordinated with miRBase (11), an authoritative resource for microRNA annotation that assigns identifiers for microRNA genes and sequences. As of Rfam 13.0, out of 1,983 microRNA families found in miRBase v21, only 28% matched one or more of the 529 Rfam microRNA families.

Starting in Rfam 14, we began systematically reviewing microRNA families with the goal to synchronise Rfam microRNA families with miRBase. To this end, we worked to ensure that

the SEED alignments of all microRNA families in Rfam were built from microRNA sequences that are tracked in the miRBase database. Multiple sequence alignments of microRNA sequences were extracted from miRBase and mapped to Rfam accessions. Each sequence was assigned a RNAcentral Unique RNA Sequence (URS) (21) identifier to remove sequence redundancy and represent only distinct sequences for each species. For each alignment a covariance model was built using Infernal and used to search the Rfamseq database. Bit score thresholds for each model (known as gathering thresholds (16)) were manually curated. These thresholds enable automatic and accurate genome annotation using Rfam microRNA families with Infernal.

Pre-existing Rfam microRNA families were reviewed and replaced with miRBase-derived SEED alignments where possible. New families were created when Rfam did not already represent the microRNA sequences. MicroRNA families found in Rfam that did not match miRBase were reviewed and updated or deleted. In cases where the miRBase alignment matched several Rfam families we manually inspected the alignment and families and split and merged families and built clans of families as appropriate.

Rfam release 15.0 marks the completion of the synchronisation process, as we have processed all microRNA alignments from miRBase. For roughly 200 miRBase-derived alignments, we determined that the family was not suitable for inclusion into Rfam. In most cases this was because there was a single unique sequence in the miRBase alignment. A small fraction of the alignments which have not been added may be removed from miRBase in the future. Table 3 shows the summary of changes since release 14.3, and a list of all microRNA families is found in Supplementary information in Table S1. All miRBase microRNA sequences that are not represented in updated Rfam families will be periodically and systematically reviewed in a cycle of improvements and curation of both Rfam and miRBase. Rfam and miRBase will continue to synchronise our resources as new microRNAs and new microRNA alignments are available, with miRBase acting as the repository of microRNA sequences, and Rfam as the authority on the grouping of those sequences into families. Rfam microRNA family classifications will be available through both resources.

Table 3: A summary of the changes in Rfam microRNA families since last publication (16) with release 14.3.

	Changes for Rfam 15.0
Total number of microRNA families	1,603
New families	722
Updated families	881
Deleted families	8

#### New families

While the primary focus of Rfam has been to complete the microRNA project, and improve existing families with 3D structures, we have continued to create new families. Since Rfam 14.3 (16), Rfam has created 16 new non-microRNA or viral RNA families. The families cover

a range of phylogenetic and functional types. A few examples include Bacteroidales small SRP (RF04183), the signal recognition particle RNA of Bacteroidetes (22), Hairpin-meta1, a virus-like ribozyme reported in RNA satellites of plant viruses (RF04190) (23), Hovlinc ribozyme (RF04188) (24), a newly discovered type of self-cleaving ribozymes found in human and other hominids and RF04222 PLRV xrRNA, a exoribonuclease-resistant RNA detected in Flavivirus Potato virus (25), RF04247 bZIP family which is a non canonical Hac1/Xbp1 intron found in *Metazoa* (26). Shown in Figure 4 are the secondary structures of selected families.

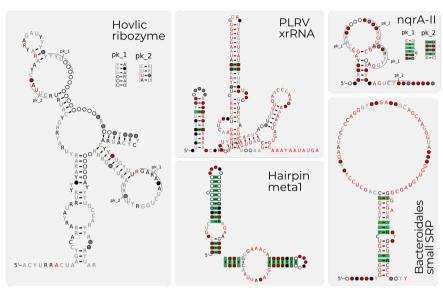


Figure 4: A selection of the new families for Rfam 15.0. The five families shown Hovlinc ribozyme (RF04188), the PLRV xrRNA (RF04222), the hairpin meta ribozyme (RF04190), nqrA-II ncRNA motif (RF04310) and Bacteriodales small SRP (RF04183) are several of the new families created for Rfam 15.0.

These families are curated through a combination of literature reviews and community submissions. We actively encourage groups with candidate RNAs to contribute their alignments and secondary structure data to Rfam. However, it is important to note that Rfam requires all sequences in SEED alignments to be traceable to public databases such as GenBank (27) or RNAcentral (21). With the increasing ease of sequencing, many laboratories maintain private sequence databases for constructing alignments. As a result, many of these new alignments cannot be incorporated into Rfam due to the lack of traceable public records. We strongly recommend that the scientific community submit their sequences to publicly accessible databases, ensuring they are available for reuse and analysis by the broader research community. Furthermore, when publishing alignments, using public accessions will facilitate faster and more efficient integration into resources like Rfam.

### Other improvements

#### Updating FULL alignments

Rfam provides several data types for interested users to download from our FTP site. These include the SEED alignments, CMs and FULL sequences. Older versions of Rfam included

FULL alignments, which are an alignment of all matching sequences to the CM. Previously, these were no longer produced because of technical limitations, however, with improvements to infernal and growth in available compute power it is now possible to create these alignments again. We now produce full alignments for all Rfam families. These are available for each model in the 'full\_alignments' section on the FTP site (https://ftp.ebi.ac.uk/pub/databases/Rfam/CURRENT/full\_alignments).

#### **APICURON** integration

APICURON (<a href="https://apicuron.org/">https://apicuron.org/</a>) is a database to track and credit biocurators for the work they do (28). The database allows for databases to credit curators for each activity they perform and provides overall statistics for each database including Rfam. In Rfam, it is common for a curator to perform many small updates and fixes to families to produce a release. Currently, this work goes unacknowledged except during publication, which are generally separated by several years. By integrating with APICURON we are able to display all the changes curators perform as part of their duties on a more regular schedule. Additionally, APICURON updates ORCID records (<a href="https://orcid.org/">https://orcid.org/</a>) to help credit curators for their activities. We update our APICURON records with each release and our APICURON page is available at: <a href="https://apicuron.org/databases/rfam">https://apicuron.org/databases/rfam</a>. APICURON requires that each resource track the activities of each curator, unfortunately some historical Rfam activity does not have the required specificity, leading to some older curators lacking attributions. This is being worked on and we hope to properly credit all curators for their contributions to Rfam.

#### Website improvements

The Rfam website (<a href="http://rfam.org">http://rfam.org</a>) has undergone continuous development since our last publication. We have added several new features, such as dedicated landing pages for each project, e.g. viral families can easily be browsed at <a href="http://rfam.org/viruses">http://rfam.org/viruses</a>. These pages make it simpler for users interested in a particular project, e.g. the 3D structure alignments, to see our progress and fetch all data related to the project.

We have recently updated some terminology used for riboswitch families in Rfam. We have found that some users are confused that Rfam riboswitch families generally include just the aptamer domains. This is because aptamers align well, while it is difficult to build an alignment of a whole riboswitch. We have added a short note to the header of each Rfam riboswitch page to indicate that this family is only the aptamer domain and a short explanation of what that means. Additionally, the descriptions of families now include the term 'aptamer' to clarify this point. We have not updated the identifiers or accessions of any families.

Finally, we have integrated the RNAcentral LitScan widget from RNAcentral. RNAcentral has developed a pipeline, LitScan, which identifies and extracts mentions of any non-coding RNA in all open access literature (29). The result of this pipeline can be reused by embedding a simple HTML widget. The widget allows users to browse and search the related literature in a convenient way. We have embedded this widget on all Rfam family pages under the new 'Publications' tab. The results are updated with each RNAcentral release, roughly every 4 months. An example of the widget for the Glutamine riboswitch family page is shown in Figure 5.

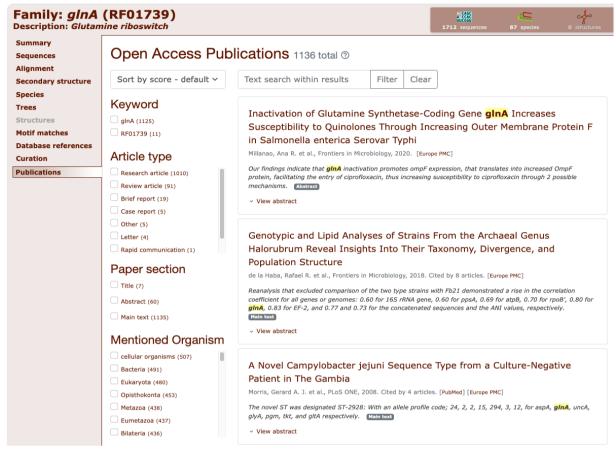


Figure 5: The LitScan widget embedded in the Glutamine riboswitch family page. This widget shows users which open access articles mention the family. Families are searched using their Rfam ID and Rfam accession in all open access literature. It allows users to search the matched articles by several types of metadata such as the article type (research, review, report, etc), the section of the paper which mentions the family and organisms mentioned in the paper. Additionally, users can perform a text search within the sentences which mention the Rfam family.

#### Conclusions

After more than 20 years of work (1), Rfam has grown to over 4,000 families and continues to be a key resource in RNA science. We continue to provide a large centralised collection of ncRNA families, which are used in many ways. Rfam was originally created to annotate genomes, but we have grown far beyond that use case. We would like to take this opportunity to thank everyone who has worked on or collaborated with Rfam over the past 20 years. Anyone interested in learning about the history and changes in Rfam since its inception are encouraged to visit <a href="https://rfam.org/rfam20">https://rfam.org/rfam20</a> to find interviews with many former and current staff and users.

Since our last publication we have focused on completing our synchronisation with miRBase and improving existing families using R-scape and 3D structures. This has led to a large increase in the number and quality of families. Rfam plans to stay synchronised with miRBase and complete the 3D structure improvements by connecting each family to at least one

structure where possible. Outside of those projects, we will continue to import viral RNAs and return to capturing more novel families discovered by the community. Finally, we will explore ways to expand Rfamseg to better improve our coverage of the phylogenetic space.

It is essential to note that there has been a major shift in bioinformatics with the publication of AlphaFold and its effect on protein structure prediction. We expect that as the field of RNA structure prediction matures, Rfam will continue to play a key role as a source of ground truth. In order to best serve this new use case, we will begin exploring ways to grow faster than before, while maintaining the high standards the community has come to expect of Rfam alignments. This will be essential to providing the test and training sets required for deep learning based approaches.

Finally, we encourage any interested community members to reach out for collaboration or to provide data. As discussed here, much of our major work is carried out with interested community members and can lead to new directions for the resource. We invite new data submissions and feedback at <a href="https://docs.rfam.org/page/contact-us.html">https://docs.rfam.org/page/contact-us.html</a>.

# Data availability

All Rfam data are released under the Creative Commons Zero (CC0) licence at <a href="https://rfam.org">https://rfam.org</a>. The data can be accessed via an API, a public MySQL database, and the FTP archive. The Rfam documentation (<a href="https://docs.rfam.org">https://docs.rfam.org</a>) and (30) contain detailed instructions. All code is available on GitHub under the Apache 2.0 licence at <a href="https://github.com/Rfam">https://github.com/Rfam</a>.

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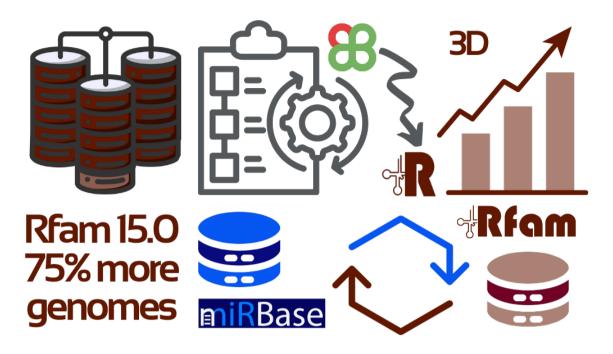
#### Conflict of interest

AB is a member of the Nucleic Acids Research Editorial Board.

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# **Graphical Abstract**



Rfam has undergone a major update with the release of 15.0. We have increased the number of genomes in our sequence database Rfamseq by 75%, completed the synchronisation with miRBase and improved 65 families using 3D structures.

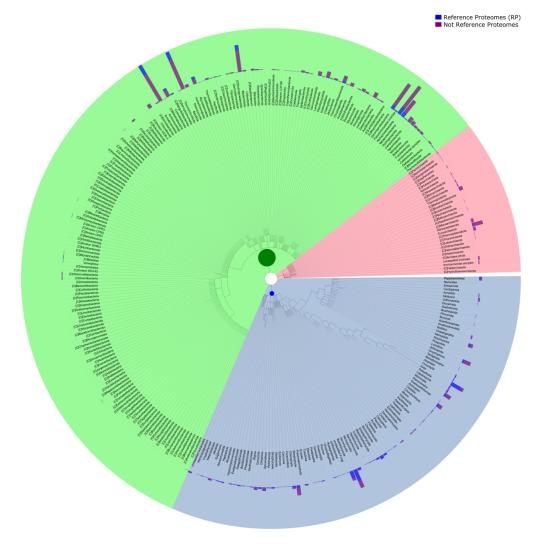
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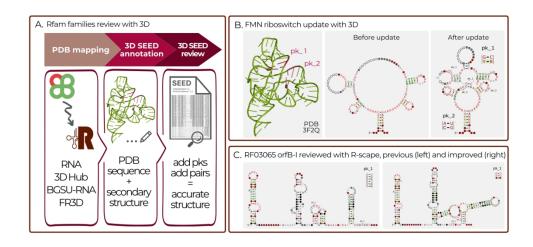
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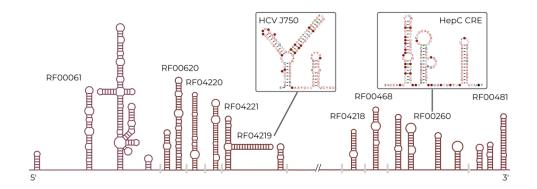


The taxonomic distribution of cellular organisms in Rfamseq 15 organised by the phylogenetic kingdom. Shown on the outside is the number of reference and non-reference proteomes in each lineage. Each coloured section indicates the kingdom with bacteria (green), archaea (pink) and eukaryotes (blue).

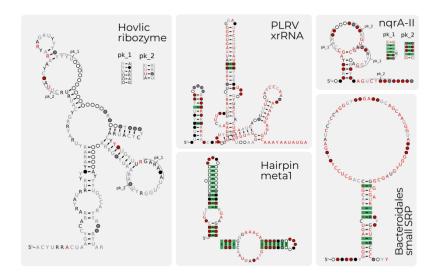
1291x1329mm (118 x 118 DPI)



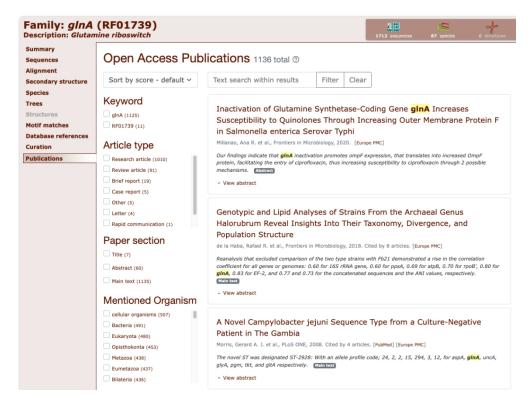
A) Pipeline for reviewing Rfam families using experimentally determined RNA 3D structures from PDB. B) Example of the FMN riboswitch (RF00050) before and after improvements with a 3D structure. Improvements included adding 2 pseudoknots and creating a missing helicees. C) A summary of how the IS605-orfB-I RNA (RF03065) family was reviewed and improved using R-scape.



A schematic of the HCV viral families Rfam now contains. These families were built from whole genome alignments, as described in (17). The figure shows their relative locations in the HCV genomic alignment and shown in the inset are two of the new HCV families.

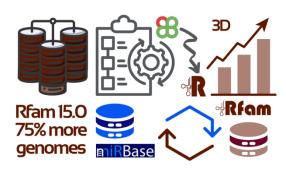


A selection of the new families for Rfam 15.0. The five families shown Hovlinc ribozyme (RF04188), the PLRV xrRNA (RF04222), the hairpin meta ribozyme (RF04190), nqrA-II ncRNA motif (RF04310) and Bacteriodales small SRP (RF04183) are several of the new families created for Rfam 15.0.



The LitScan widget embedded in the Glutamine riboswitch family page. This widget shows users which open access articles mention the family. Families are searched using their Rfam ID and Rfam accession in all open access literature. It allows users to search the matched articles by several types of metadata such as the article type (research, review, report, etc), the section of the paper which mentions the family and organisms mentioned in the paper. Additionally, users can perform a text search within the sentences which mention the Rfam family.

424x318mm (72 x 72 DPI)



Rfam has undergone a major update with the release of 15.0. We have increased the number of genomes in our sequence database Rfamseq by 75%, completed the synchronisation with miRBase and improved 65 families using 3D structures.

Kingdom	Rfamseq v14	Rfamseq v15
Total sequence length	3.68e11	1.46e12
Number of species	14,774	26,106
Eukaryotes	1,057	2,613
Archaea	418	370
Bacteria	7,808	9,601
Viruses	5,491	13,552
Number of Rfam hits	2,897,296	10,736,534
Size on disk	170 GB	772 GB

Family	Additional Covarying Basepairs
RF02033 (HEARO)	24
RF03065 (IS605-orfB-I)	14
RF03068 (RT-3)	8
RF03072 (raiA)	5
RF02969 (DUF3800-I)	4
RF01688 (Actino-pnp)	3
RF02004 (group-II-D1D4-5)	3
RF02005 (group-II-D1D4-6)	3
RF02913 (pemK)	3
RF03077 (RT-2)	3
RF03135 (L4-Archaeoglobi)	3
RF03144 (eL15- Euryarchaeota)	3
RF00062 (HgcC)	2
RF01731 (TwoAYGGAY)	2
RF01794 (sok)	2
RF02221 (sRNA-Xcc1)	2
RF02947 (cow-rumen-2)	2
RF03000 (LOOT)	2
RF03158 (L31-Actinobacteria)	2
RF01864 (plasmodium_snoR21)	1
RF01867 (CC2171)	1
RF02944 (c4-2)	1
RF02968 (DUF3800-IX)	1
RF02987 (GA-cis)	1
RF03019 (RT-16)	1
RF03046 (Pseudomonadales-1)	1

	Changes for Rfam 15.0
Total number of microRNA families	1,603
New families	722
Updated families	881
Deleted families	8

Rfam Accession	Rfam ID	miRBase Family Accession
RF00027	let-7	MIPF0000002
RF00047	mir-2	MIPF0000049
RF00051	mir-17	MIPF0000001
RF00052	lin-4	MIPF0000303
RF00053	mir-7	MIPF0000022
RF00073	mir-156	MIPF0000008
RF00074	mir-29	MIPF0000009
RF00075	mir-166	MIPF0000004
RF00076	mir-181	MIPF0000007
RF00103	mir-1	MIPF0000038
RF00104	mir-10	MIPF0000033
RF00104	mir-10	MIPF0000025
RF00104	mir-10	MIPF0000268
RF00104	mir-10	MIPF0000271
RF00129	mir-103	MIPF0000024
RF00130	mir-192	MIPF0000063
RF00131	mir-30	MIPF0000005
RF00143	mir-6	MIPF0000119
RF00144	mir-199	MIPF0000040
RF00237	mir-9	MIPF0000014
RF00239	mir-124	MIPF0000021
RF00241	mir-8	MIPF0000019
RF00244	mir-26	MIPF0000043
RF00245	mir-19	MIPF0000011
RF00246	mir-135	MIPF0000028
RF00247	mir-160	MIPF0000032
RF00251	mir-219	MIPF0000044
RF00253	mir-101	MIPF0000046
RF00254	mir-16	MIPF0000006
RF00255	mir-218	MIPF0000026
RF00256	mir-196	MIPF0000031

RF00257	mir-194	MIPF0000055
RF00258	mir-130	MIPF0000034
RF00363	mir-BART1	MIPF0000325
RF00365	mir-BHRF1-1	MIPF0000331
RF00366	mir-BHRF1-2	MIPF0000332
RF00446	mir-133	MIPF0000029
RF00455	mir-15	MIPF000006
RF00637	mir-276	MIPF0000124
RF00638	MIR159	MIPF0000010
RF00639	mir-515	MIPF0000020
RF00640	MIR167	MIPF0000023
RF00641	mir-154	MIPF0000018
RF00642	mir-23	MIPF0000027
RF00643	MIR171_1	MIPF0000030
RF00644	mir-27	MIPF0000036
RF00645	MIR169_2	MIPF0000037
RF00646	mir-204	MIPF0000042
RF00647	MIR164	MIPF0001794
RF00648	MIR396	MIPF0000047
RF00649	mir-128	MIPF0000048
RF00650	mir-153	MIPF0000050
RF00651	mir-221	MIPF0000051
RF00652	MIR478	MIPF0000052
RF00653	mir-22	MIPF0000053
RF00654	mir-216	MIPF0000054
RF00655	mir-28	MIPF0000057
RF00656	mir-205	MIPF0000058
RF00657	mir-184	MIPF0000059
RF00658	mir-21	MIPF0000060
RF00659	mir-365	MIPF0000061
RF00660	mir-214	MIPF0000062
RF00661	mir-31	MIPF0000064
RF00662	mir-132	MIPF0000065

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RF00664	mir-223	MIPF0000067
RF00665	mir-290	MIPF0000068
RF00666	mir-32	MIPF0000069
RF00667	mir-33	MIPF0000070
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RF00679	mir-210	MIPF0000086
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RF00683	mir-143	MIPF0000094
RF00684	mir-122	MIPF0000095
RF00685	mir-36	MIPF0000096
RF00686	mir-338	MIPF0000097
RF00687	mir-136	MIPF0000099
RF00688	MIR394	MIPF0000100
RF00689	MIR390	MIPF0000101
RF00690	MIR408	MIPF0000102
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RF00701	mir-126	MIPF0000115
RF00702	mir-182	MIPF0000116
RF00703	mir-139	MIPF0000117
RF00704	MIR397	MIPF0000120
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RF00706	mir-263	MIPF0000122
RF00707	mir-197	MIPF0000123
RF00708	mir-450	MIPF0000128
RF00709	mir-455	MIPF0000129
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RF00714	MIR535	MIPF0000136
RF00715	mir-383	MIPF0000137
RF00716	mir-3	MIPF0000140
RF00717	mir-315	MIPF0000141
RF00718	mir-431	MIPF0000142
RF00719	mir-326	MIPF0000143
RF00720	mir-317	MIPF0000144
RF00721	MIR475	MIPF0000145
RF00722	mir-451	MIPF0000148
RF00723	mir-448	MIPF0000149
RF00724	mir-282	MIPF0000150
RF00725	mir-iab-4	MIPF0000151
RF00726	mir-87	MIPF0000152
RF00727	bantam	MIPF0000153
RF00728	mir-81	MIPF0000154

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RF00731	mir-155	MIPF0000157
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RF03456	mir-724	MIPF0001616
RF03457	MIR173	MIPF0001175
RF03458	mir-766	MIPF0000677
RF03459	mir-1890	MIPF0000848
RF03460	mir-561	MIPF0000623
RF03461	MIR842	MIPF0001158
RF03462	mir-4677	MIPF0001848
RF03463	mir-3036	MIPF0001098
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RF03465	MIR9666	MIPF0001930

RF03466	mir-1184	MIPF0000592
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RF03468	mir-2830	MIPF0001052
RF03469	mir-1179	MIPF0000632
RF03470	mir-571	MIPF0000757
RF03471	mir-745	MIPF0001267
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RF03476	mir-7360	MIPF0001649
RF03477	MIR862_2	MIPF0001295
RF03478	mir-630	MIPF0000653
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RF03480	mir-1284	MIPF0000604
RF03481	mir-9014	MIPF0002031
RF03482	mir-4774	MIPF0001578
RF03483	MIR319	MIPF0001104
RF03484	mir-K12-10	MIPF0000261
RF03485	mir-4000	MIPF0000867
RF03486	mir-3940	MIPF0001599
RF03487	mir-4423	MIPF0001587
RF03488	mir-2366	MIPF0001403
RF03489	mir-4796	MIPF0001402
RF03490	mir-1426	MIPF0000692
RF03491	mir-3146	MIPF0001626
RF03492	mir-878	MIPF0000434
RF03493	mir-1281	MIPF0000695
RF03494	mir-4680	MIPF0001988
RF03495	mir-968	MIPF0000865
RF03496	MIR2086	MIPF0001302
RF03497	MIR5038	MIPF0001368
RF03498	mir-3145	MIPF0001427

RF03499	mir-702	MIPF0001163
RF03500	mir-1262	MIPF0000595
RF03501	mir-3140	MIPF0001615
RF03502	mir-1914	MIPF0001040
RF03503	mir-547	MIPF0000779
RF03504	mir-9461	MIPF0002000
RF03505	mir-9128	MIPF0002044
RF03506	mir-587	MIPF0000563
RF03507	mir-4429	MIPF0001474
RF03508	mir-970	MIPF0000700
RF03509	mir-4660	MIPF0001460
RF03510	mir-3660	MIPF0001397
RF03511	mir-5444	MIPF0001358
RF03512	mir-3479	MIPF0001629
RF03514	mir-6128	MIPF0001442
RF03516	MIR5229	MIPF0001323
RF03517	MIR5225	MIPF0001388
RF03518	mir-1893	MIPF0001812
RF03519	mir-238	MIPF0000862
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RF03521	mir-23c	MIPF0001429
RF03522	mir-4850	MIPF0002086
RF03523	mir-4703	MIPF0001759
RF03524	mir-3174	MIPF0001569
RF03525	mir-1343	MIPF0001206
RF03526	mir-7177	MIPF0002020
RF03527	mir-3072	MIPF0001410
RF03528	mir-1264	MIPF0000098
RF03548	mir-302_2	MIPF0000658
RF03549	mir-15_2	MIPF0001828
RF03550	mir-31_2	MIPF0001803
RF03552	mir-43_2	MIPF0001285
RF03553	mir-190_2	MIPF0001853

RF03554         mir-92_2         MIPF0001099           RF03555         mir-2_2         MIPF0001838           RF03556         mir-559         MIPF0000656           RF03557         MIR6019         MIPF0001486           RF03558         mir-629         MIPF0001555           RF03559         mir-8303         MIPF0002121           RF03560         mir-3926         MIPF0002121           RF03561         mir-3533         MIPF000263           RF03562         mir-2004         MIPF0002042           RF03563         mir-BART16         MIPF0002042           RF03564         mir-9416         MIPF0000991           RF03565         mir-1282         MIPF0000587           RF03566         mir-1282         MIPF0001222           RF03567         MIR8024         MIPF0001789           RF03568         mir-3204         MIPF0000198           RF03569         mir-7973         MIPF0001964           RF03570         mir-8286         MIPF0001543           RF03572         MIR2109         MIPF0001275           RF03573         mir-1954         MIPF0001275           RF03574         mir-4847         MIPF00001220           RF03578         mir-4435	-		
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RF03571 mir-734 MIPF0001543 RF03572 MIR2109 MIPF0001000 RF03573 mir-1954 MIPF0001275 RF03574 mir-4847 MIPF0002100 RF03575 mir-rL1-14 MIPF0000993 RF03576 mir-4435 MIPF0001220 RF03577 mir-9193 MIPF0002089 RF03578 mir-3910 MIPF0001148 RF03579 mir-2058 MIPF0001793 RF03580 mir-928 MIPF0000950 RF03581 mir-4878 MIPF0000950 RF03582 mir-m22-1 MIPF0000451 RF03583 MIR781 MIPF0001137 RF03584 mir-2290 MIPF0001973 RF03585 mir-3938 MIPF0001520	RF03569	mir-7973	MIPF0001693
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RF03573 mir-1954 MIPF0001275 RF03574 mir-4847 MIPF0002100 RF03575 mir-rL1-14 MIPF0000993 RF03576 mir-4435 MIPF0001220 RF03577 mir-9193 MIPF0002089 RF03578 mir-3910 MIPF0001148 RF03579 mir-2058 MIPF0001793 RF03580 mir-928 MIPF0000950 RF03581 mir-4878 MIPF0000950 RF03582 mir-m22-1 MIPF0000451 RF03583 MIR781 MIPF0001137 RF03584 mir-2290 MIPF0001973 RF03585 mir-3938 MIPF0001520	RF03571	mir-734	MIPF0001543
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RF03590	mir-2009	MIPF0001923
RF03591	mir-3117	MIPF0001506
RF03592	mir-2567	MIPF0000934
RF03593	mir-2331	MIPF0001951
RF03594	mir-3817	MIPF0001217
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RF03596	mir-5880	MIPF0001483
RF03597	MIR6438	MIPF0001538
RF03598	MIR6024	MIPF0001459
RF03599	MIR1883	MIPF0000683
RF03600	MIR3627	MIPF0001471
RF03601	MIR7993	MIPF0001663
RF03602	mir-8250	MIPF0002067
RF03603	mir-4899	MIPF0001837
RF03604	mir-5728	MIPF0001558
RF03605	MIR4221	MIPF0001128
RF03606	mir-2071	MIPF0001681
RF03607	mir-9447	MIPF0001994
RF03608	mir-9440	MIPF0001915
RF03609	mir-5470	MIPF0001967
RF03610	mir-H7	MIPF0000929
RF03611	mir-7147	MIPF0001705
RF03612	mir-608	MIPF0000972
RF03613	MIR6234	MIPF0001642
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RF03988	mir-6301	MIPF0001556
RF03989	mir-7950	MIPF0001678
RF03990	mir-4888	MIPF0001724
RF03991	mir-9209	MIPF0001927
RF03992	mir-6302	MIPF0001398
RF03993	mir-8536	MIPF0002030
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RF03995	mir-7957	MIPF0001668
RF03996	mir-3012	MIPF0001018
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RF03998	mir-2160	MIPF0000769
RF03999	mir-7883	MIPF0001849
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RF04005	mir-4599	MIPF0001252
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RF04010	mir-3003	MIPF0000818
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RF04015	MIR946	MIPF0000427
RF04016	mir-4873	MIPF0001806
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RF04021	mir-3492	MIPF0001622
RF04022	mir-4863	MIPF0001778
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RF04045	mir-2147	MIPF0000751
RF04046	mir-2154	MIPF0000781
RF04047	mir-9228	MIPF0001876
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RF04054	mir-2944	MIPF0001335
RF04055	mir-1261	MIPF0000802
RF04056	mir-982	MIPF0000843
RF04057	MIR1432	MIPF0001063
RF04059	MIR5020	MIPF0001226
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RF04064	MIR5267	MIPF0001276
RF04065	mir-7563	MIPF0001709
RF04066	mir-1256	MIPF0000594
RF04067	mir-4520	MIPF0001272
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RF04075	mir-1545	MIPF0001711
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RF04078	mir-1905	MIPF0000753
RF04079	MIR161	MIPF0000455
RF04080	mir-762	MIPF0000624
RF04081	mir-1992	MIPF0000705
RF04082	mir-54	MIPF0000874
RF04083	MIR528	MIPF0000868
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RF04085	mir-1271	MIPF0000483

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RF04094         MIR537         MIPF0000207           RF04095         MIR9746         MIPF0001884           RF04096         mir-4433         MIPF0001826           RF04097         MIR2630         MIPF0000814           RF04098         MIR9471         MIPF0002045           RF04099         MIR837         MIPF0001155           RF04100         mir-493         MIPF0000230           RF04101         mir-8864         MIPF0001919           RF04102         MIR6457         MIPF0001534           RF04103         mir-8356         MIPF0001997           RF04104         MIR918         MIPF0000374           RF04105         mir-3165         MIPF0000608           RF04106         mir-1303         MIPF0000608           RF04107         mir-5391         MIPF0001944           RF04108         mir-2513         MIPF0000980           RF04109         mir-1305         MIPF0000965           RF04110         MIR5084         MIPF0001706           RF04111         MIR6274         MIPF0001478           RF04113         MIR1523         MIPF0001327           RF04114         MIR4374         MIPF0001413           RF04115         mir-6130         MIPF000	RF04092	MIR1508	MIPF0000719
RF04095 MIR9746 MIPF0001884 RF04096 mir-4433 MIPF0001826 RF04097 MIR2630 MIPF0000814 RF04098 MIR9471 MIPF0002045 RF04099 MIR837 MIPF0001155 RF04100 mir-493 MIPF0001919 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0001706 RF04111 MIR6274 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001944 RF04116 MIR9672 MIPF0001904 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04093	mir-1784	MIPF0001873
RF04096 mir-4433 MIPF0001826 RF04097 MIR2630 MIPF0000814 RF04098 MIR9471 MIPF0002045 RF04099 MIR837 MIPF0001155 RF04100 mir-493 MIPF0000230 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0001997 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000985 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04094	MIR537	MIPF0000207
RF04097 MIR2630 MIPF0000814 RF04098 MIR9471 MIPF0002045 RF04099 MIR837 MIPF0001155 RF04100 mir-493 MIPF0000230 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0001635 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000980 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04095	MIR9746	MIPF0001884
RF04098 MIR9471 MIPF0002045 RF04099 MIR837 MIPF0001155 RF04100 mir-493 MIPF0000230 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000980 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04096	mir-4433	MIPF0001826
RF04099 MIR837 MIPF0001155 RF04100 mir-493 MIPF0000230 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04097	MIR2630	MIPF0000814
RF04100 mir-493 MIPF0000230 RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001478 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04098	MIR9471	MIPF0002045
RF04101 mir-8864 MIPF0001919 RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001327 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04099	MIR837	MIPF0001155
RF04102 MIR6457 MIPF0001534 RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04100	mir-493	MIPF0000230
RF04103 mir-8356 MIPF0001997 RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001942 RF04116 MIR9672 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04101	mir-8864	MIPF0001919
RF04104 MIR918 MIPF0000374 RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04102	MIR6457	MIPF0001534
RF04105 mir-3165 MIPF0001635 RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04103	mir-8356	MIPF0001997
RF04106 mir-1303 MIPF0000608 RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04104	MIR918	MIPF0000374
RF04107 mir-5391 MIPF0001944 RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04105	mir-3165	MIPF0001635
RF04108 mir-2513 MIPF0000980 RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04106	mir-1303	MIPF0000608
RF04109 mir-1305 MIPF0000965 RF04110 MIR5084 MIPF0002026 RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04107	mir-5391	MIPF0001944
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RF04111 MIR6274 MIPF0001706 RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04109	mir-1305	MIPF0000965
RF04112 MIR5512 MIPF0001478 RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04110	MIR5084	MIPF0002026
RF04113 MIR1523 MIPF0001327 RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04111	MIR6274	MIPF0001706
RF04114 MIR4374 MIPF0001152 RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04112	MIR5512	MIPF0001478
RF04115 mir-6130 MIPF0001413 RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04113	MIR1523	MIPF0001327
RF04116 MIR9672 MIPF0001942 RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04114	MIR4374	MIPF0001152
RF04117 MIR5200 MIPF0001904 RF04118 MIR9481 MIPF0002096	RF04115	mir-6130	MIPF0001413
RF04118 MIR9481 MIPF0002096	RF04116	MIR9672	MIPF0001942
	RF04117	MIR5200	MIPF0001904
RF04119 MIR1858 MIPF0000650	RF04118	MIR9481	MIPF0002096
	RF04119	MIR1858	MIPF0000650

RF04120	MIR5998	MIPF0001613
RF04121	mir-1289	MIPF0000626
RF04122	mir-1204	MIPF0000671
RF04123	mir-3127	MIPF0001439
RF04124	MIR8565	MIPF0001889
RF04125	MIR9408	MIPF0002027
RF04126	mir-9198	MIPF0001887
RF04127	MIR8007	MIPF0001839
RF04128	MIR8562	MIPF0001861
RF04129	MIR4240	MIPF0001198
RF04130	MIR7996	MIPF0001691
RF04131	mir-2032	MIPF0000736
RF04132	mir-4436	MIPF0001236
RF04133	MIR2646	MIPF0001010
RF04134	MIR7504	MIPF0001798
RF04135	MIR5062	MIPF0001922
RF04136	MIR1862	MIPF0000580
RF04137	MIR834	MIPF0001183
RF04138	mir-9243	MIPF0001926
RF04139	MIR841	MIPF0001112
RF04140	mir-3118	MIPF0001928
RF04141	MIR8622	MIPF0001931
RF04142	mir-1299	MIPF0000625
RF04143	MIR869	MIPF0001167
RF04144	MIR9783	MIPF0001947
RF04145	MIR8706	MIPF0001953
RF04146	mir-4742	MIPF0001552
RF04147	MIR6224	MIPF0001435
RF04148	MIR844	MIPF0001197
RF04149	MIR447	MIPF0000170
RF04150	MIR2655	MIPF0000813
RF04151	MIR4387	MIPF0001109
RF04152	mir-3618	MIPF0001710

RF04153	MIR856	MIPF0001162
RF04154	mir-1277	MIPF0001937
RF04155	mir-9236	MIPF0001905
RF04156	MIR2670	MIPF0000855
RF04157	MIR9486	MIPF0002054
RF04158	mir-568	MIPF0000408
RF04159	MIR1437	MIPF0001762
RF04160	MIR868	MIPF0001173
RF04161	mir-3688	MIPF0001263
RF04162	MIR9662	MIPF0002119
RF04163	mir-8364	MIPF0001877
RF04164	MIR1507	MIPF0000699
RF04165	mir-2483	MIPF0001532
RF04166	mir-2278	MIPF0001522
RF04167	mir-1972	MIPF0001025
RF04168	MIR839	MIPF0001143
RF04169	mir-2235	MIPF0000825
RF04170	MIR7723	MIPF0001739
RF04171	MIR1509	MIPF0000771
RF04172	MIR8675	MIPF0002106
RF04173	MIR847	MIPF0001171
RF04174	MIR4208	MIPF0001108
RF04175	mir-1260b	MIPF0001381
RF04176	mir-3149	MIPF0001935
RF04185	MIR2118_2	MIPF0001409
RF04186	mir-278_2	MIPF0000728
RF04187	MIR862	MIPF0001145
RF04193	mir-51	MIPF0000268
RF04194	mir-57	MIPF0000271
RF04195	MIR6217	MIPF0001550
RF04196	MIR5638	MIPF0001350
RF04197	mir-9457	MIPF0002122
RF04198	MIR8670	MIPF0002041

RF04199	MIR6027	MIPF0001424
RF04200	mir-9437	MIPF0001943
RF04201	MIR7510	MIPF0001684
RF04202	MIR5380	MIPF0001331
RF04203	MIR1144	MIPF0000432
RF04204	mir-3473	MIPF0001230
RF04205	MIR2606	MIPF0000888
RF04206	MIR9555	MIPF0002081
RF04207	MIR5048	MIPF0001453
RF04208	MIR8742	MIPF0002016
RF04209	MIR7533	MIPF0001834
RF04210	MIR8643	MIPF0002008
RF04211	MIR5185	MIPF0001246
RF04212	mir-8904	MIPF0001977
RF04213	MIR7807	MIPF0002091
RF04214	MIR7526	MIPF0001651
RF04215	mir-9214	MIPF0002108
RF04216	mir-509	MIPF0000130
RF04217	mir-297	MIPF0000204
RF04223	MIR2619	MIPF0001294
RF04224	mir-9229	MIPF0001875
RF04225	MIR7502	MIPF0001869
RF04226	mir-9186	MIPF0001868
RF04227	mir-9215	MIPF0001893
RF04228	MIR6140	MIPF0001666
RF04229	mir-9279	MIPF0001911
RF04230	mir-9261	MIPF0001938
RF04231	mir-9191	MIPF0001987
RF04232	mir-9318	MIPF0002056
RF04233	mir-680	MIPF0000338
RF04234	mir-1421	MIPF0000564
RF04235	mir-242_2	MIPF0000731
RF04236	mir-1285	MIPF0000559

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RF04237	mir-1490	MIPF0000525
RF04238	mir-4012	MIPF0000964
RF04239	mir-4679	MIPF0001228
RF04240	mir-4716	MIPF0001476
RF04241	mir-5014	MIPF0001634
RF04242	mir-5595	MIPF0001590
RF04243	MIR169_3	MIPF0000253
RF04244	MIR169_6	MIPF0001058
RF04245	MIR169_7	MIPF0001690
RF04246	MIR169_8	MIPF0001712
RF04248	MIR7486	MIPF0001682
RF04249	mir-310	MIPF0000566
RF04250	mir-236	MIPF0000232
RF04251	MIR5070	MIPF0002013
RF04252	mir-8186	MIPF0002035
RF04254	mir-1677	MIPF0000849
RF04255	MIR8001	MIPF0001852
RF04257	mir-9230	MIPF0002035
RF04258	mir-994	MIPF0001045
RF04259	mir-2003	MIPF0002049
RF04260	mir-2005	MIPF0001956
RF04261	MIR1861	MIPF0000567
RF04262	MIR1319	MIPF0001448
RF04263	MIR2593	MIPF0000733
RF04264	mir-5408	MIPF0001313
RF04265	mir-4650	MIPF00001234
RF04266	mir-9412	MIPF0001891
RF04267	mir-362	MIPF0000209
RF04268	mir-512	MIPF0000518
RF04269	mir-373	MIPF0000500
RF04270	mir-743	MIPF0000386
RF04271	mir-1244	MIPF0000569
RF04272	mir-8908	MIPF0001890

MIR827_2	MIPF0001769
mir-9223	MIPF0001885
mir-2733	MIPF0000766
MIR4371	MIPF0001141
mir-1268	MIPF0000946
MIR4372	MIPF0001257
mir-H20	MIPF0001760
mir-890	MIPF0000386
mir-8791	MIPF0001882
mir-507	MIPF0000130
mir-3135	MIPF0001219
mir-532	MIPF0000113
mir-660	MIPF0000113
mir-513	MIPF0000130
mir-9195	MIPF0001939
mir-9201	MIPF0001898
mir-3596	MIPF0001194
mir-1197	MIPF0000126
mir-368	MIPF0000091
mir-379	MIPF0000126
mir-889	MIPF0000514
mir-3578	MIPF0001166
mir-329	MIPF0000110
mir-485	MIPF0000201
mir-35_2	MIPF0001648
mir-36_2	MIPF0001664
MIR814	MIPF0000351
mir-39	MIPF0000304
mir-200	MIPF0000491
mir-506	MIPF0000130
MIR162_2	MIPF0000169
	mir-9223 mir-2733 MIR4371 mir-1268 MIR4372 mir-H20 mir-890 mir-8791 mir-507 mir-3135 mir-532 mir-660 mir-513 mir-9195 mir-9201 mir-3596 mir-1197 mir-368 mir-379 mir-368 mir-379 mir-389 mir-3578 mir-329 mir-485 mir-35_2 mir-36_2 MIR814 mir-39 mir-200 mir-506

Table S1: The mapping between Rfam accessions, ids and miRBase family accessions.