

Structural Bioinformatics

De novo SVM classification of precursor microRNAs from genomic pseudo hairpins using global and intrinsic folding measures

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1 BIOGENESIS OF MATURE MICRORNAS

In the prevailing biogenesis model of miRNAs maturation (Bartel 2004; Kim 2005), >1000-nts primary transcripts (*pri-miRs*) originate from the intergenic or intronic regions of annotated (non-)protein-coding transcription units (Rodriguez *et al.*, 2004). The *pri-miRs* are cleaved by the nuclear RNase III endonuclease Drosha/Pasha complex, yielding ~70–120-nts precursor transcripts (*pre-miRs*) with 5' phosphate and ~2-nts 3' overhang. These *pre-miRs* exhibiting characteristic imperfect and extended RNA stem-loop structures are actively exported by the cargo transporter Exportin-5 in a Ran-GTP dependent manner into the cytoplasm (Bohnsack *et al.*, 2004; Yi *et al.*, 2003; Zeng and Cullen 2004). The *pre-miRs* are further excised by another RNase III endonuclease Dicer into an intermediate duplex *miR:miR**, a ~21–23-nts asymmetric mature miRNA duplex. The *miR:miR** is recruited by a ribonucleoprotein RNA-Induced Silencing Complex (RISC) (Gregory *et al.*, 2005; Maniataki and Mourelatos 2005; Rivas *et al.*, 2005; Tang 2005). The strand *miR* with weaker hydrogen binding survives as the mature miRNA, which is preferentially loaded into RISC. miRNA-directed posttranscriptional silencing of target genes occurs by mRNA degradation (Brennecke *et al.*, 2005), or translational arrest of protein synthesis (Doench and Sharp 2004), or mRNA deadenylation (Wu *et al.*, 2006).

2 MATERIALS AND METHODS

2.1 Biologically relevant datasets

8,494 pseudo hairpin sequences. We analyzed 8,494 pseudo hairpin sequences that were extracted from the protein-coding regions (CDSs) according to the UCSC refGene annotation tables (Karolchik *et al.*, 2003) and human RefSeq genes (Pruitt and Maglott 2001). As wrongly assumed 'negative samples' can distort the decision boundary of SVM in an unpredictable and/or significant manner, special requirements were imposed on the selection of genomic inverted repeats. First, they must originate from genomic regions that do not undergo any known experimentally validated alternative splicing (AS) events, as described previously (Xue *et al.*, 2005). This criterion ensures that they do not encode genuine human *pre-miRs*. Second, they are analogous to genuine human *pre-miRs* by displaying similar distribution in terms of their length ~90-nts, hairpin structures with stem ≥ 8 -bps including the GU wobble pairs, and MFEs ≤ -15 kcal/mol. In addition, they fold without multiple loops in their RNA structures as verified by the RNAfold program (Hofacker 2003).

2,241 non-redundant *pre-miR* sequences. We retrieved 4,028 annotated *pre-miR* sequences spanning across 45 species from miRBase Registry Database release 8.2, July 2006 (Griffiths-Jones *et al.*, 2006). As strong sequence homologies existed among *pre-miRs* both within a single and between different species, the homologs of the training *pre-miRs* must be excluded from the testing and inde-

pendent evaluation sets. The original dataset was filtered to 90% identity using a greedy incremental clustering algorithm (Li and Godzik 2006). Briefly, all the sequences were first sorted in order of decreasing length and the longest one became the representative of the first cluster. Each remaining sequence was compared with the existing representatives and grouped into their cluster if the similarity with any representative was above a given threshold (default value is 0.9), else that sequence became the representative of a new cluster. Consequently, 2,241 non-redundant *pre-miRs* spanning 41 species (categorized into arthropoda, nematoda, vertebrata, viridiplantae, and viruses) served as the reference positive set; none of the sequences from *Gorilla gorilla*, *Macaca nemestrina*, *Pan paniscus*, and *Pongo pygmaeus* was retained.

12,387 functional prokaryotic and eukaryotic ncRNA sequences. We retrieved all curated seed ncRNA sequences from Rfam repository release 7.0, March 2005 (Griffiths-Jones *et al.*, 2005). After removing 46 types of *pre-miRs*, 12,387 functional prokaryotic and eukaryotic ncRNAs spanning 457 types (categorized into 16 classes) served as the negative non *pre-miR* dataset. Functional ncRNAs have similar length distribution to the known *pre-miRs*, and can fold with hairpin(s) or stem-loop(s) (Eddy 2001; Storz 2002; Svoboda and Cara 2006).

31 mRNA sequences. We investigated 31 mRNA sequences that tend to fold into complex RNA structures with extremely negative MFEs (Freyhult *et al.*, 2005). They were randomly selected from GeneBank DNA database (Benson *et al.*, 2005) with the following GenBank accession numbers: NM_001005151.1, NM_001003967.1, NM_177233.4, AY675236.1, NM_001004202.1, NM_178539.2, AB164385.1, AY555511.1, AB189435.1, NM_178307.2, NM_001003966.1, NM_205498.1, NM_013564.3, Z81556.1, NM_131070.2, X56279.1, AK045412.1, AF452886.1, BC049701.1, BC050086.1, NM_172343.1, AY182163.1, BC072691.1, CV127341.1, NC_004671.1, X00910.1, AY226143.1, AJ621386, CV122154.1, X68284, and CV199185.1.

2.2 Feature vector

Adjusted base pairing propensity, *dP* measures the total number of base pairs present in the RNA secondary structure *S* (Schultes *et al.*, 1999) divided by the length *L* in nucleotides. It removes the bias that a long sequence tends to have more base pairs. *dP* ranges [0.0, 0.5], 0.0 for no base pair interactions and 0.5 for maximum of *L/2* base pairs.

Adjusted minimum free energy of folding, *dG* measures the thermodynamic stability of RNA structure *S* i.e., the lowest MFE for the most favorable conformation, divided by the length *L* in nucleotides (Freyhult *et al.*, 2005). It removes the bias that a long sequence tends to have lower negative MFE (Seffens and Digby 1999).

MFE Index 1, *MFEI* is the ratio of *dG* and %G+C content (Zhang *et al.*, 2006).

Adjusted shannon entropy, *dQ* in Eq. (1), characterizes the base pairing probability distribution (BPPD) in a RNA structure *S* as a chaotic dynamical system (Freyhult *et al.*, 2005; Huynen *et al.*, 1997; Schultes *et al.*, 1999). Low values of *dQ* correspond to BPPD that are dominated by single, a few, or by the absence of base pairings. These bases are better predicted than those having multiple alternative states.

$$dQ = -\frac{1}{L} \sum_{i < j} p_{ij} \log_2(p_{ij}), \quad p_{ij} = \sum_{S_a \in S(\mathbf{x})} P(S_a) d_{ij}^a. \quad (1)$$

Here, p_{ij} denotes the probability of bases i and j pair, computed from the McCaskill's algorithm (J.S.McCaskill 1990); $d_{ij}^a = 1$ if i and j pair, 0 otherwise. RNA molecules exist *in vivo* as an ensemble of secondary structures $S_a \in S(\mathbf{x})$ with the Boltzmann distribution probability $P(S_a)$ (Mathews 2004).

Adjusted base pair distance, dD in Eq. (2), is the base pair distance for all pairs of structures S_a and S_b inferred from sequence \mathbf{s} (Freyhult *et al.*, 2005; Moulton *et al.*, 2000).

$$dD = \frac{1}{2L} \sum_{S_a, S_b \in S(\mathbf{s})} P(S_a)P(S_b) d_{BP}(S_a, S_b) = \frac{1}{L} \sum_{i < j} p_{ij} (1 - p_{ij}). \quad (2)$$

Here, the number of base pairs not shared by them is given by $d_{BP}(S_a, S_b) = |S_a \cup S_b| - |S_a \cap S_b| = \sum_{i < j} (d_{ij}^a + d_{ij}^b - 2d_{ij}^a d_{ij}^b)$. The number of base pairs in S_a is $|S_a| = \sum_{i < j} d_{ij}^a$. Definitions of p_{ij} and d_{ij}^a follow those of dQ in Eq. (1).

Second (or the Fiedler) eigenvalue, dF in Eq. (3), measures the compactness of a tree-graph $G = (V, E)$ (Fera *et al.*, 2004; Gan *et al.*, 2004). At the coarsest scale, each vertex $v \in V$ represents a bulge loop, hairpin loop, internal loop, the 5' and 3' unpaired termini, or the multi-branch loop; each edge $e \in E$ denotes a RNA stem. dF is computed from the Laplacian matrix $\mathbf{L}(G)$, a mathematical representation of the tree-graph G . dF can be used as a similarity measure among a collection of RNA secondary structures.

$$\mathbf{L}(G)\mathbf{X} = \mathbf{I}\mathbf{X} \Leftrightarrow dF = \text{FidlerEigen}[\mathbf{L}(G)]. \quad (3)$$

MFE Index 2, $MFEI_2$ is the ratio of dG and the number of stems S , which are structural motifs containing more than three contiguous base pairs.

Normalized feature vectors. The Z-score $Z(\mathbf{s}_n)$ in Eq. (4) normalizes the feature $S(\mathbf{s}_n)$ of n^{th} native RNA sequence \mathbf{s}_n in terms of the number of standard deviations by which $S(\mathbf{s}_n)$ differs from the mean of inferred $R = 10^4$ random RNA sequences \mathbf{r}_n . $S(\mathbf{s}_n)$ can be dG , dP , dQ , dD , and dF ; corresponding Z-scores are denoted as zG , zP , zQ , zD , and zF .

$$Z(\mathbf{s}_n) = \frac{S(\mathbf{s}_n) - \bar{m}_n}{s_n}, \quad s_n^2 = \frac{1}{R-1} \sum_{i=1}^R [S_i(\mathbf{r}_n) - \bar{m}_n]^2. \quad (4)$$

Here, $S(\mathbf{r}_n)$ is the computed feature for the i^{th} random sequence of \mathbf{r}_n , \bar{m}_n and s_n are the sample mean and the standard deviation of the feature $S(\mathbf{s}_n)$. The entire set of R random sequences \mathbf{r}_n is synthesized by the "Altschul-Erikson algorithm" (Altschul and Erickson 1985), a form of dinucleotide shuffling. Briefly, it shuffles \mathbf{s}_n while preserving exactly both the mono- and di-nucleotide frequencies. The \mathbf{r}_n shares the same first and last nucleotides as \mathbf{s}_n . The order of the shuffled nucleotides is 'less random' due to fewer possible dinucleotide-preserving permutations.

2.3 Statistical tests and performance evaluation

F-scores of features. The "quality" of the i^{th} feature is described commonly by the F-scores $F1$ (Dror *et al.*, 2005) and $F2$ (Chen and Lin 2006) in Eq. (5). The larger their values for the i^{th} feature, the more likely this feature possesses discriminative importance/power.

$$F1 = \frac{|m_+^* - m_-^*|}{|s_+^* + s_-^*|}, \quad F2 = \frac{(m_+^* - \bar{m})^2 + (m_-^* - \bar{m})^2}{(s_+^*)^2 + (s_-^*)^2}. \quad (5)$$

Here m_+^*/m_-^* and s_+^*/s_-^* denote the means and standard deviations of the positive (+) and negative (-) training datasets, correspondingly. The numerator and denominator describe the discrimination between the two classes, and that within each of the two classes.

Performance measures. SE (Sensitivity or recall), SP (Specificity), ACC (Accuracy), Fm (F-measure) (Liu *et al.*, 2006), and MCC (Matthew's Correlation Coefficient) (Bhasin *et al.*, 2006) are defined in Eq. (6). All metrics (except MCC) range [0.0, 1.0]; closer to 1.0 implies better scores, and *vice-versa*. MCC ranges [-1.0, 1.0]; -1.0, 0.0, and 1.0 indicate worst possible, perfectly random, and best possible classification, respectively. Unlike ACC , Fm and MCC account for the unbalanced datasets.

$$SE = \frac{TP}{TP + FN}, \quad SP = \frac{TN}{TN + FP}, \quad ACC = \frac{TP + TN}{TP + TN + FP + FN},$$

$$Fm = \frac{2(SP \cdot PPV)}{SP + PPV} \quad \text{where } PPV \text{ (Positive Predictive Value)} = \frac{TP}{TP + FP}, \quad (6)$$

$$MCC = \frac{TP \cdot TN - FN \cdot FP}{\sqrt{(TP + FN)(TN + FP)(TP + FP)(TN + FN)}}.$$

Here TP , FN , FP , and TN denote the number of true/false *pre-miRs* detected/missed, correspondingly. The "quality" of a binary classification, denoted simply as ROC score, is measured by the normalized area under the Receiver Operating Characteristic curve (ROC) that plots the trade-off between the SE and FPR (False-positive rate = $1 - SP$) across all possible classification thresholds (Lasko *et al.*, 2005). ROC score ranges [0.5, 1.0]; closer to 0.5 (about the upward diagonal) and to 1.0 (along the left-top boundary) signify a totally random and a perfect classifier (Lasko *et al.*, 2005). In this work, the ROC score was computed by the trapezoidal rule (Kestler 2001).

Benchmarking miPred. Both $3SVM$ (Xue *et al.*, 2005) and Naïve Bayesian Classifier (NBC) served as independent baseline models to benchmark the performance improvements or deterioration (if any) of *miPred*. The original $3SVM$ was previously trained on 163 human *pre-miRs* and 168 pseudo hairpins using the older libSVM 2.36 with the "-b 1" option disabled. Here, $3SVM$ was trained on randomly selected 200 human *pre-miRs* and 400 pseudo hairpins using the latest libSVM 2.82 (the "-b 1" option is enabled) and the optimal hyperparameter pair (C, γ) . $3SVM$ was applied to the testing and independent evaluation datasets with "svm-predict -b 1". The Bayes Classifier Induction (*bci*) version 2.14, a free implementation of NBC (<http://fuzzy.cs.unimagdeburg.de/~borgelt/bayes.html>), was used for training and testing with the exact samples and attributes employed by $3SVM$ and *miPred*; denoted as $3SVM-NBC$ and *miPred-NBC*. For training, "*bci* -L1" yielded better classification results than the default "-LO". In theory, NBC seeks to maximize the probability $P(X|C) = P(f_1, f_2, \dots, f_n|C)$ such that the sample X belongs to one of the binary classes $C = (T, F)$.

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Table S1. The prediction performances of *miPred*, *miPred-NBC*, *3SVM*, and *3SVM-NBC* evaluated on the *pre-miR* datasets TR-H (200 human *pre-miR*s and 400 pseudo hairpins), TE-H (remaining 123 human *pre-miR*s and 246 pseudo hairpins), and IE-NH (1,918 *pre-miR*s across 40 non-human species and 3,836 pseudo hairpins).*miPred*

Species	Genus	TP	FN	P	FP	TN	N	%SE	%SP	%FPR	%ACC
<i>Homo sapiens</i>	Vertebrata	176	24	200	10	390	400	88.00	97.50	2.50	94.33
<i>Homo sapiens</i>	Vertebrata	104	19	123	5	241	246	84.55	97.97	2.03	93.50
<i>Anopheles gambiae</i>	Arthropoda	37	1	38	1	75	76	97.37	98.68	1.32	98.25
<i>Apis mellifera</i>	Arthropoda	25	0	25	2	48	50	100	96	4	97.33
<i>Arabidopsis thaliana</i>	Viridiplantae	101	7	108	2	214	216	93.52	99.07	0.93	97.22
<i>Ateles geoffroyi</i>	Vertebrata	2	0	2	0	4	4	100	100	0	100
<i>Bos taurus</i>	Vertebrata	8	3	11	2	20	22	72.73	90.91	9.09	84.85
<i>Caenorhabditis briggsae</i>	Nematoda	72	4	76	1	151	152	94.74	99.34	0.66	97.81
<i>Caenorhabditis elegans</i>	Nematoda	96	17	113	7	219	226	84.96	96.9	3.1	92.92
<i>Canis familiaris</i>	Vertebrata	3	0	3	0	6	6	100	100	0	100
<i>Danio rerio</i>	Vertebrata	235	11	246	19	473	492	95.53	96.14	3.86	95.94
<i>Drosophila melanogaster</i>	Arthropoda	67	6	73	4	142	146	91.78	97.26	2.74	95.43
<i>Drosophila pseudoobscura</i>	Arthropoda	32	3	35	1	69	70	91.43	98.57	1.43	96.19
Epstein Barr (EBV)	Viruses	22	0	22	2	42	44	100	95.45	4.55	96.97
<i>Fugu rubripes</i>	Vertebrata	68	2	70	2	138	140	97.14	98.57	1.43	98.1
<i>Gallus gallus</i>	Vertebrata	87	5	92	4	180	184	94.57	97.83	2.17	96.74
<i>Glycine max</i>	Viridiplantae	20	1	21	0	42	42	95.24	100	0	98.41
Herpes Simplex (HSV)	Viruses	1	0	1	0	2	2	100	100	0	100
Human cytomegalovirus (HCMV)	Viruses	11	0	11	1	21	22	100	95.45	4.55	96.97
Kaposi sarcoma-associated herpesvirus (KSHV)	Viruses	11	1	12	0	24	24	91.67	100	0	97.22
<i>Lagothrix lagotricha</i>	Vertebrata	1	1	2	0	4	4	50	100	0	83.33
<i>Lemur catta</i>	Vertebrata	2	1	3	0	6	6	66.67	100	0	88.89
<i>Macaca mulatta</i>	Vertebrata	1	1	2	0	4	4	50	100	0	83.33
<i>Medicago truncatula</i>	Viridiplantae	17	1	18	0	36	36	94.44	100	0	98.15
Mouse γ -herpesvirus (MGHV68)	Viruses	8	1	9	1	17	18	88.89	94.44	5.56	92.59
<i>Mus musculus</i>	Vertebrata	166	33	199	9	389	398	83.42	97.74	2.26	92.96
<i>Oryza sativa</i>	Viridiplantae	140	12	152	4	300	304	92.11	98.68	1.32	96.49
<i>Ovis aries</i>	Vertebrata	2	0	2	0	4	4	100	100	0	100
<i>Pan troglodytes</i>	Vertebrata	2	1	3	1	5	6	66.67	83.33	16.67	77.78
<i>Physcomitrella patens</i>	Viridiplantae	17	0	17	0	34	34	100	100	0	100
<i>Populus trichocarpa</i>	Viridiplantae	144	13	157	13	301	314	91.72	95.86	4.14	94.48
<i>Rattus norvegicus</i>	Vertebrata	56	12	68	10	126	136	82.35	92.65	7.35	89.22
Rhesus lymphocryptovirus	Viruses	16	0	16	2	30	32	100	93.75	6.25	95.83
<i>Saccharum officinarum</i>	Viridiplantae	3	1	4	0	8	8	75	100	0	91.67
<i>Saguinus labiatus</i>	Vertebrata	1	1	2	0	4	4	50	100	0	83.33
Simian virus (SV40)	Viruses	1	0	1	0	2	2	100	100	0	100
<i>Sorghum bicolor</i>	Viridiplantae	48	2	50	2	98	100	96	98	2	97.33
<i>Sus scrofa</i>	Vertebrata	1	1	2	0	4	4	50	100	0	83.33
<i>Tetraodon nigroviridis</i>	Vertebrata	40	3	43	0	86	86	93.02	100	0	97.67
<i>Xenopus laevis</i>	Vertebrata	4	1	5	0	10	10	80	100	0	93.33
<i>Xenopus tropicalis</i>	Vertebrata	119	6	125	4	246	250	95.2	98.4	1.6	97.33
<i>Zea mays</i>	Viridiplantae	79	0	79	5	153	158	100	96.84	3.16	97.89
Total samples		2046	195	2241	114	4368	4482				

(Species) Row 1 (TR-H), row 2 (TE-H), and the remaining rows 3–43 (IE-NH). TP (real *pre-miR*s detected), FN (real *pre-miR*s missed), P (real *pre-miR*s), FP (pseudo hairpins detected), TN (pseudo hairpins missed), N (pseudo hairpins), %SE (Sensitivity), %SP (Specificity), %FPR (False-positive rate), and %ACC (Accuracy).

miPred-NBC

Species	Genus	TP	FN	P	FP	TN	N	%SE	%SP	%FPR	%ACC
Homo sapiens	Vertebrata	200	0	200	0	400	400	100.00	100.00	0.00	100.00
Homo sapiens	Vertebrata	46	77	123	36	210	246	37.40	85.37	14.63	69.38
<i>Anopheles gambiae</i>	Arthropoda	12	26	38	7	69	76	31.58	90.79	9.21	71.05
<i>Apis mellifera</i>	Arthropoda	6	19	25	6	44	50	24.00	88.00	12.00	66.67
<i>Arabidopsis thaliana</i>	Viridiplantae	20	88	108	27	189	216	18.52	87.50	12.50	64.51
<i>Ateles geoffroyi</i>	Vertebrata	0	2	2	1	3	4	0.00	75.00	25.00	50.00
<i>Bos taurus</i>	Vertebrata	1	10	11	2	20	22	9.09	90.91	9.09	63.64
<i>Caenorhabditis briggsae</i>	Nematoda	27	49	76	20	132	152	35.53	86.84	13.16	69.74
<i>Caenorhabditis elegans</i>	Nematoda	51	62	113	26	200	226	45.13	88.50	11.50	74.04
<i>Canis familiaris</i>	Vertebrata	0	3	3	1	5	6	0.00	83.33	16.67	55.56
<i>Danio rerio</i>	Vertebrata	71	175	246	62	430	492	28.86	87.40	12.60	67.89
<i>Drosophila melanogaster</i>	Arthropoda	21	52	73	17	129	146	28.77	88.36	11.64	68.49
<i>Drosophila pseudoobscura</i>	Arthropoda	12	23	35	5	65	70	34.29	92.86	7.14	73.33
Epstein Barr (EBV)	Viruses	6	16	22	4	40	44	27.27	90.91	9.09	69.70
<i>Fugu rubripes</i>	Vertebrata	10	60	70	18	122	140	14.29	87.14	12.86	62.86
<i>Gallus gallus</i>	Vertebrata	24	68	92	22	162	184	26.09	88.04	11.96	67.39
<i>Glycine max</i>	Viridiplantae	2	19	21	3	39	42	9.52	92.86	7.14	65.08
Herpes Simplex (HSV)	Viruses	0	1	1	1	1	2	0.00	50.00	50.00	33.33
Human cytomegalovirus (HCMV)	Viruses	0	11	11	5	17	22	0.00	77.27	22.73	51.52
Kaposi sarcoma-associated herpesvirus (KSHV)	Viruses	1	11	12	5	19	24	8.33	79.17	20.83	55.56
<i>Lagothrix lagotricha</i>	Vertebrata	0	2	2	0	4	4	0.00	100.00	0.00	66.67
<i>Lemur catta</i>	Vertebrata	0	3	3	2	4	6	0.00	66.67	33.33	44.44
<i>Macaca mulatta</i>	Vertebrata	0	2	2	0	4	4	0.00	100.00	0.00	66.67
<i>Medicago truncatula</i>	Viridiplantae	4	14	18	4	32	36	22.22	88.89	11.11	66.67
Mouse γ -herpesvirus (MGHV68)	Viruses	2	7	9	3	15	18	22.22	83.33	16.67	62.96
<i>Mus musculus</i>	Vertebrata	37	162	199	52	346	398	18.59	86.93	13.07	64.15
<i>Oryza sativa</i>	Viridiplantae	35	117	152	37	267	304	23.03	87.83	12.17	66.23
<i>Ovis aries</i>	Vertebrata	0	2	2	0	4	4	0.00	100.00	0.00	66.67
<i>Pan troglodytes</i>	Vertebrata	2	1	3	1	5	6	66.67	83.33	16.67	77.78
<i>Physcomitrella patens</i>	Viridiplantae	3	14	17	3	31	34	17.65	91.18	8.82	66.67
<i>Populus trichocarpa</i>	Viridiplantae	33	124	157	41	273	314	21.02	86.94	13.06	64.97
<i>Rattus norvegicus</i>	Vertebrata	23	45	68	11	125	136	33.82	91.91	8.09	72.55
Rhesus lymphocryptovirus	Viruses	5	11	16	2	30	32	31.25	93.75	6.25	72.92
<i>Saccharum officinarum</i>	Viridiplantae	1	3	4	0	8	8	25.00	100.00	0.00	75.00
<i>Saguinus labiatus</i>	Vertebrata	0	2	2	1	3	4	0.00	75.00	25.00	50.00
Simian virus (SV40)	Viruses	0	1	1	1	1	2	0.00	50.00	50.00	33.33
<i>Sorghum bicolor</i>	Viridiplantae	7	43	50	13	87	100	14.00	87.00	13.00	62.67
<i>Sus scrofa</i>	Vertebrata	0	2	2	1	3	4	0.00	75.00	25.00	50.00
<i>Tetraodon nigroviridis</i>	Vertebrata	9	34	43	9	77	86	20.93	89.53	10.47	66.67
<i>Xenopus laevis</i>	Vertebrata	2	3	5	4	6	10	40.00	60.00	40.00	53.33
<i>Xenopus tropicalis</i>	Vertebrata	35	90	125	33	217	250	28.00	86.80	13.20	67.20
<i>Zea mays</i>	Viridiplantae	16	63	79	18	140	158	20.25	88.61	11.39	65.82
Total samples		724	1517	2241	504	3978	4482				

(Species) Row 1 (TR-H), row 2 (TE-H), and the remaining rows 3–43 (IE-NH). TP (real pre-miRs detected), FN (real pre-miRs missed), P (real pre-miRs), FP (pseudo hairpins detected), TN (pseudo hairpins missed), N (pseudo hairpins), %SE (Sensitivity), %SP (Specificity), %FPR (False-positive rate), and %ACC (Accuracy).

3SVM[†]

Species	Genus	TP	FN	P	FP	TN	N	%SE	%SP	%FPR	%ACC
<i>Homo sapiens</i>	Vertebrata	172	28	200	12	388	400	86.00	97.00	3.00	93.33
<i>Homo sapiens</i>	Vertebrata	79	29	108	10	206	216	73.15	95.37	4.63	87.96
<i>Anopheles gambiae</i>	Arthropoda	33	4	37	1	73	74	89.19	98.65	1.35	95.50
<i>Apis mellifera</i>	Arthropoda	23	2	25	1	49	50	92.00	98.00	2.00	96.00
<i>Arabidopsis thaliana</i>	Viridiplantae	69	2	71	5	137	142	97.18	96.48	3.52	96.71
<i>Ateles geoffroyi</i>	Vertebrata	2	0	2	0	4	4	100.00	100.00	0.00	100.00
<i>Bos taurus</i>	Vertebrata	7	1	8	3	13	16	87.50	81.25	18.75	83.33
<i>Caenorhabditis briggsae</i>	Nematoda	68	2	70	6	134	140	97.14	95.71	4.29	96.19
<i>Caenorhabditis elegans</i>	Nematoda	94	13	107	4	210	214	87.85	98.13	1.87	94.70
<i>Canis familiaris</i>	Vertebrata	3	0	3	1	5	6	100.00	83.33	16.67	88.89
<i>Danio rerio</i>	Vertebrata	201	32	233	30	436	466	86.27	93.56	6.44	91.13
<i>Drosophila melanogaster</i>	Arthropoda	57	9	66	7	125	132	86.36	94.70	5.30	91.92
<i>Drosophila pseudoobscura</i>	Arthropoda	28	7	35	1	69	70	80.00	98.57	1.43	92.38
Epstein Barr (EBV)	Viruses	19	3	22	0	44	44	86.36	100.00	0.00	95.45
<i>Fugu rubripes</i>	Vertebrata	48	16	64	5	123	128	75.00	96.09	3.91	89.06
<i>Gallus gallus</i>	Vertebrata	73	14	87	4	170	174	83.91	97.70	2.30	93.10
<i>Glycine max</i>	Viridiplantae	16	0	16	0	32	32	100.00	100.00	0.00	100.00
Herpes Simplex (HSV)	Viruses	0	1	1	0	2	2	0.00	100.00	0.00	66.67
Human cytomegalovirus (HCMV)	Viruses	8	3	11	0	22	22	72.73	100.00	0.00	90.91
Kaposi sarcoma-associated herpesvirus (KSHV)	Viruses	4	8	12	0	24	24	33.33	100.00	0.00	77.78
<i>Lagothrix lagotricha</i>	Vertebrata	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Lemur catta</i>	Vertebrata	2	0	2	0	4	4	100.00	100.00	0.00	100.00
<i>Macaca mulatta</i>	Vertebrata	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Medicago truncatula</i>	Viridiplantae	15	0	15	2	28	30	100.00	93.33	6.67	95.56
Mouse γ -herpesvirus (MGHV68)	Viruses	5	4	9	1	17	18	55.56	94.44	5.56	81.48
<i>Mus musculus</i>	Vertebrata	145	41	186	5	367	372	77.96	98.66	1.34	91.76
<i>Oryza sativa</i>	Viridiplantae	106	9	115	11	219	230	92.17	95.22	4.78	94.20
<i>Ovis aries</i>	Vertebrata	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Pan troglodytes</i>	Vertebrata	2	1	3	0	6	6	66.67	100.00	0.00	88.89
<i>Physcomitrella patens</i>	Viridiplantae	14	0	14	0	28	28	100.00	100.00	0.00	100.00
<i>Populus trichocarpa</i>	Viridiplantae	106	15	121	12	230	242	87.60	95.04	4.96	92.56
<i>Rattus norvegicus</i>	Vertebrata	50	12	62	5	119	124	80.65	95.97	4.03	90.86
<i>Rhesus lymphocryptovirus</i>	Viruses	16	0	16	1	31	32	100.00	96.88	3.13	97.92
<i>Saccharum officinarum</i>	Viridiplantae	0	0	0	0	0	0	NaN	NaN	NaN	NaN
<i>Saguinus labiatus</i>	Vertebrata	0	1	1	0	2	2	0.00	100.00	0.00	66.67
Simian virus (SV40)	Viruses	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Sorghum bicolor</i>	Viridiplantae	33	2	35	2	68	70	94.29	97.14	2.86	96.19
<i>Sus scrofa</i>	Vertebrata	0	2	2	0	4	4	0.00	100.00	0.00	66.67
<i>Tetraodon nigroviridis</i>	Vertebrata	39	2	41	3	79	82	95.12	96.34	3.66	95.94
<i>Xenopus laevis</i>	Vertebrata	2	3	5	1	9	10	40.00	90.00	10.00	73.33
<i>Xenopus tropicalis</i>	Vertebrata	101	21	122	7	237	244	82.79	97.13	2.87	92.35
<i>Zea mays</i>	Viridiplantae	50	2	52	7	97	104	96.15	93.27	6.73	94.23
Total samples		1694	289	1983	147	3819	3966				

[†], 3SVM model was trained on 200 human *pre-miRs* and 400 pseudo hairpins randomly selected using the latest libSVM 2.82 (the "-b 1" option was enabled) and the optimal hyperparameter pair (C , γ). (*Species*) Row 1 (TR-H), row 2 (TE-H), and the remaining rows 3–43 (IE-NH). *TP* (real *pre-miRs* detected), *FN* (real *pre-miRs* missed), *P* (real *pre-miRs*), *FP* (pseudo hairpins detected), *TN* (pseudo hairpins missed), *N* (pseudo hairpins), *%SE* (Sensitivity), *%SP* (Specificity), *%FPR* (False-positive rate), and *%ACC* (Accuracy).

3SVM-NBC

Species	Genus	TP	FN	P	FP	TN	N	%SE	%SP	%FPR	%ACC
Homo sapiens	Vertebrata	196	4	200	13	387	400	98.00	96.75	3.25	97.17
Homo sapiens	Vertebrata	71	37	108	51	165	216	65.74	76.39	23.61	72.84
<i>Anopheles gambiae</i>	Arthropoda	27	10	37	18	56	74	72.97	75.68	24.32	74.77
<i>Apis mellifera</i>	Arthropoda	20	5	25	9	41	50	80.00	82.00	18.00	81.33
<i>Arabidopsis thaliana</i>	Viridiplantae	44	27	71	30	112	142	61.97	78.87	21.13	73.24
<i>Ateles geoffroyi</i>	Vertebrata	1	1	2	1	3	4	50.00	75.00	25.00	66.67
<i>Bos taurus</i>	Vertebrata	4	4	8	4	12	16	50.00	75.00	25.00	66.67
<i>Caenorhabditis briggsae</i>	Nematoda	52	18	70	23	117	140	74.29	83.57	16.43	80.48
<i>Caenorhabditis elegans</i>	Nematoda	87	20	107	39	175	214	81.31	81.78	18.22	81.62
<i>Canis familiaris</i>	Vertebrata	3	0	3	2	4	6	100.00	66.67	33.33	77.78
<i>Danio rerio</i>	Vertebrata	140	93	233	112	354	466	60.09	75.97	24.03	70.67
<i>Drosophila melanogaster</i>	Arthropoda	38	28	66	31	101	132	57.58	76.52	23.48	70.20
<i>Drosophila pseudoobscura</i>	Arthropoda	20	15	35	15	55	70	57.14	78.57	21.43	71.43
Epstein Barr (EBV)	Viruses	12	10	22	9	35	44	54.55	79.55	20.45	71.21
<i>Fugu rubripes</i>	Vertebrata	31	33	64	33	95	128	48.44	74.22	25.78	65.63
<i>Gallus gallus</i>	Vertebrata	48	39	87	44	130	174	55.17	74.71	25.29	68.20
<i>Glycine max</i>	Viridiplantae	5	11	16	5	27	32	31.25	84.38	15.63	66.67
Herpes Simplex (HSV)	Viruses	0	1	1	0	2	2	0.00	100.00	0.00	66.67
Human cytomegalovirus (HCMV)	Viruses	3	8	11	1	21	22	27.27	95.45	4.55	72.73
Kaposi sarcoma-associated herpesvirus (KSHV)	Viruses	2	10	12	5	19	24	16.67	79.17	20.83	58.33
<i>Lagothrix lagotricha</i>	Vertebrata	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Lemur catta</i>	Vertebrata	2	0	2	0	4	4	100.00	100.00	0.00	100.00
<i>Macaca mulatta</i>	Vertebrata	1	0	1	1	1	2	100.00	50.00	50.00	66.67
<i>Medicago truncatula</i>	Viridiplantae	8	7	15	6	24	30	53.33	80.00	20.00	71.11
Mouse γ -herpesvirus (MGHV68)	Viruses	4	5	9	7	11	18	44.44	61.11	38.89	55.56
<i>Mus musculus</i>	Vertebrata	110	76	186	83	289	372	59.14	77.69	22.31	71.51
<i>Oryza sativa</i>	Viridiplantae	73	42	115	56	174	230	63.48	75.65	24.35	71.59
<i>Ovis aries</i>	Vertebrata	1	0	1	1	1	2	100.00	50.00	50.00	66.67
<i>Pan troglodytes</i>	Vertebrata	2	1	3	1	5	6	66.67	83.33	16.67	77.78
<i>Physcomitrella patens</i>	Viridiplantae	7	7	14	3	25	28	50.00	89.29	10.71	76.19
<i>Populus trichocarpa</i>	Viridiplantae	73	48	121	52	190	242	60.33	78.51	21.49	72.45
<i>Rattus norvegicus</i>	Vertebrata	39	23	62	24	100	124	62.90	80.65	19.35	74.73
Rhesus lymphocryptovirus	Viruses	10	6	16	6	26	32	62.50	81.25	18.75	75.00
<i>Saccharum officinarum</i>	Viridiplantae	0	0	0	0	0	0	NaN	NaN	NaN	NaN
<i>Saguinus labiatus</i>	Vertebrata	0	1	1	0	2	2	0.00	100.00	0.00	66.67
Simian virus (SV40)	Viruses	1	0	1	0	2	2	100.00	100.00	0.00	100.00
<i>Sorghum bicolor</i>	Viridiplantae	19	16	35	14	56	70	54.29	80.00	20.00	71.43
<i>Sus scrofa</i>	Vertebrata	1	1	2	0	4	4	50.00	100.00	0.00	83.33
<i>Tetraodon nigroviridis</i>	Vertebrata	18	23	41	17	65	82	43.90	79.27	20.73	67.48
<i>Xenopus laevis</i>	Vertebrata	0	5	5	1	9	10	0.00	90.00	10.00	60.00
<i>Xenopus tropicalis</i>	Vertebrata	68	54	122	49	195	244	55.74	79.92	20.08	71.86
<i>Zea mays</i>	Viridiplantae	18	34	52	30	74	104	34.62	71.15	28.85	58.97
Total samples		1260	723	1983	796	3170	3966				

(Species) Row 1 (TR-H), row 2 (TE-H), and the remaining rows 3–43 (IE-NH). TP (real pre-miRs detected), FN (real pre-miRs missed), P (real pre-miRs), FP (pseudo hairpins detected), TN (pseudo hairpins missed), N (pseudo hairpins), %SE (Sensitivity), %SP (Specificity), %FPR (False-positive rate), and %ACC (Accuracy).

Table S2. The mean sensitivity and specificity of *miPred*, *miPred-NBC*, *3SVM*, and *3SVM-NBC* evaluated on the non-human *pre-miR* dataset IE-NH (1,918 *pre-miRs* across 40 non-human species and 3,836 pseudo hairpins) categorized by genus of *pre-miRs*.

Genus	No. of species	<i>miPred</i>		<i>miPred-NBC</i>		No. of excluded species	<i>3SVM</i> [‡]		<i>3SVM-NBC</i>	
		%SE	%SP	%SE	%SP		%SE	%SP	%SE	%SP
Arthropoda	4	95.14 ± 2.11	97.63 ± 0.63	29.66 ± 2.20	90.00 ± 1.14	0	86.89 ± 2.57	97.48 ± 0.94	66.92 ± 5.71	78.19 ± 1.41
Viridiplantae	9	93.11 ± 2.47	98.72 ± 0.51	19.02 ± 1.60	90.09 ± 1.40	1	95.92 ± 1.57	96.31 ± 0.93	51.16 ± 4.31	79.73 ± 1.92
Vertebrata [†]	18	79.29 ± 4.56	97.53 ± 1.05	15.91 ± 4.43	84.83 ± 2.60	0	76.44 ± 7.48	96.11 ± 1.35	61.23 ± 7.22	79.58 ± 3.53
Nematoda	2	89.85 ± 4.89	98.12 ± 1.22	40.33 ± 4.80	87.67 ± 0.83	0	92.50 ± 4.65	96.92 ± 1.21	77.80 ± 3.51	82.68 ± 0.90
Viruses	7	97.22 ± 1.81	97.01 ± 1.08	12.72 ± 5.23	74.92 ± 6.81	0	64.00 ± 14.04	98.76 ± 0.84	43.63 ± 12.49	85.22 ± 5.36

[†], *Homo sapiens* is excluded. [‡], *3SVM* model was trained on 200 human *pre-miRs* and 400 pseudo hairpins randomly selected using the latest libSVM 2.82 (the "-b 1" option was enabled) and the optimal hyperparameter pair (C, γ). %SE (Sensitivity) and %SP (Specificity). Values are expressed as mean ± standard error.

Table S3. The prediction performances of *miPred*, *miPred-NBC*, *3SVM*, and *3SVM-NBC* evaluated on the non *pre-miR* datasets IE-NC (12,387 functional ncRNAs) and IE-M (31 mRNAs).

Accession	Type ^f	Class	<i>miPred</i> <i>miPred-NBC</i>					<i>3SVM</i> ^g <i>3SVM-NBC</i>				
			<i>N</i>	<i>TN</i>	% <i>SP</i>	<i>TN</i>	% <i>SP</i>	<i>N</i>	<i>TN</i>	% <i>SP</i>	<i>TN</i>	% <i>SP</i>
RF00001	5S ribosomal RNA	Gene/rRNA	589	409	69.44	517	87.78	2	2	100.00	1	50.00
RF00002	5.8S ribosomal RNA	Gene/rRNA	63	59	93.65	59	93.65	1	1	100.00	1	100.00
RF00003	U1 spliceosomal RNA	Gene/snRNA/splicing	54	38	70.37	45	83.33	0	0	NaN	0	NaN
RF00004	U2 spliceosomal RNA	Gene/snRNA/splicing	73	8	10.96	53	72.60	0	0	NaN	0	NaN
RF00005	tRNA	Gene/tRNA	1114	953	85.55	969	86.98	158	150	94.94	142	89.87
RF00006	Vault RNA	Gene	9	5	55.56	8	88.89	3	3	100.00	1	33.33
RF00007	U12 minor spliceosomal RNA	Gene/snRNA/splicing	7	4	57.14	7	100.00	0	0	NaN	0	NaN
RF00008	Hammerhead ribozyme (type III)	Gene/ribozyme	84	61	72.62	68	80.95	1	1	100.00	1	100.00
RF00009	Nuclear RNase P	Gene/ribozyme	53	16	30.19	50	94.34	0	0	NaN	0	NaN
RF00010	Bacterial RNase P class A	Gene/ribozyme	236	77	32.63	203	86.02	0	0	NaN	0	NaN
RF00011	Bacterial RNase P class B	Gene/ribozyme	30	12	40.00	28	93.33	0	0	NaN	0	NaN
RF00012	U3 small nucleolar RNA	Gene/snRNA/guide/C/D-box	21	10	47.62	18	85.71	0	0	NaN	0	NaN
RF00013	6S / SsrS RNA	Gene	7	1	14.29	6	85.71	2	0	0.00	1	50.00
RF00014	DsrA RNA	Gene/sRNA	3	0	0.00	2	66.67	0	0	NaN	0	NaN
RF00015	U4 spliceosomal RNA	Gene/snRNA/splicing	25	21	84.00	25	100.00	1	1	100.00	1	100.00
RF00016	U14 small nucleolar RNA	Gene/snRNA/guide/C/D-box	18	17	94.44	16	88.89	2	2	100.00	2	100.00
RF00017	Eukaryotic type signal recognition particle RNA	Gene	70	3	4.29	61	87.14	0	0	NaN	0	NaN
RF00018	CsrB/RsmB RNA family	Gene/sRNA	9	9	100.00	8	88.89	0	0	NaN	0	NaN
RF00019	Y RNA	Gene	15	9	60.00	12	80.00	5	5	100.00	2	40.00
RF00020	U5 spliceosomal RNA	Gene/snRNA/splicing	32	12	37.50	26	81.25	0	0	NaN	0	NaN
RF00021	Spot 42 RNA	Gene/sRNA	8	0	0.00	8	100.00	0	0	NaN	0	NaN
RF00022	GcvB RNA	Gene/sRNA	5	3	60.00	5	100.00	0	0	NaN	0	NaN
RF00023	tmRNA	Gene	87	53	60.92	79	90.80	0	0	NaN	0	NaN
RF00024	Vertebrate telomerase RNA	Gene	35	10	28.57	31	88.57	0	0	NaN	0	NaN
RF00025	Ciliate telomerase RNA	Gene	16	13	81.25	12	75.00	0	0	NaN	0	NaN
RF00026	U6 spliceosomal RNA	Gene/snRNA/splicing	53	52	98.11	48	90.57	0	0	NaN	0	NaN
RF00028	Group I catalytic intron	Intron	30	15	50.00	29	96.67	0	0	NaN	0	NaN
RF00029	Group II catalytic intron	Intron	116	37	31.90	89	76.72	0	0	NaN	0	NaN
RF00030	RNase MRP	Gene/ribozyme	26	9	34.62	25	96.15	0	0	NaN	0	NaN
RF00031	Selenocysteine insertion sequence	Cis-reg	64	52	81.25	50	78.13	56	56	100.00	50	89.29
RF00032	Histone 3' UTR stem-loop	Cis-reg	64	64	100.00	57	89.06	26	26	100.00	26	100.00
RF00033	MicF RNA	Gene/antisense	9	8	88.89	6	66.67	0	0	NaN	0	NaN
RF00034	RprA RNA	Gene/sRNA	9	7	77.78	9	100.00	0	0	NaN	0	NaN
RF00035	OxyS RNA	Gene/sRNA	6	4	66.67	6	100.00	0	0	NaN	0	NaN
RF00036	HIV Rev response element	Cis-reg	65	0	0.00	39	60.00	0	0	NaN	0	NaN
RF00037	Iron response element	Cis-reg	39	39	100.00	33	84.62	0	0	NaN	0	NaN
RF00038	PrfA thermoregulator UTR	Cis-reg/thermoregulator	11	11	100.00	11	100.00	5	5	100.00	5	100.00
RF00039	DicF RNA	Gene/antisense	5	5	100.00	5	100.00	2	2	100.00	2	100.00
RF00040	RNase E 5' UTR element	Cis-reg	7	5	71.43	7	100.00	0	0	NaN	0	NaN
RF00041	Enteroviral 3' UTR element	Cis-reg	60	49	81.67	45	75.00	0	0	NaN	0	NaN
RF00042	CopA-like RNA	Gene/antisense	17	0	0.00	11	64.71	0	0	NaN	0	NaN
RF00043	R1162-like plasmid antisense RNA	Gene/antisense	6	6	100.00	5	83.33	0	0	NaN	0	NaN
RF00044	Bacteriophage pRNA	Gene	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00045	U17/E1 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	23	16	69.57	18	78.26	0	0	NaN	0	NaN
RF00046	Small nucleolar RNA R30/Z108	Gene/snRNA/guide/C/D-box	6	6	100.00	2	33.33	0	0	NaN	0	NaN
RF00048	Enterovirus cis-acting replication element	Cis-reg	56	31	55.36	35	62.50	56	30	53.57	23	41.07
RF00049	U36/R47/Z100 small nucleolar RNA	Gene/snRNA/guide/C/D-box	20	20	100.00	19	95.00	3	3	100.00	2	66.67
RF00050	FMN riboswitch (RFN element)	Cis-reg/riboswitch	48	41	85.42	45	93.75	0	0	NaN	0	NaN
RF00054	U25 small nucleolar RNA	Gene/snRNA/guide/C/D-box	8	8	100.00	7	87.50	2	2	100.00	1	50.00
RF00055	Small nucleolar RNA Z37	Gene/snRNA/guide/C/D-box	8	8	100.00	5	62.50	0	0	NaN	0	NaN
RF00056	U71 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	15	10	66.67	11	73.33	0	0	NaN	0	NaN
RF00057	RyhB RNA	Gene/sRNA	9	9	100.00	6	66.67	0	0	NaN	0	NaN
RF00058	HgcF RNA	Gene	4	0	0.00	4	100.00	0	0	NaN	0	NaN
RF00059	TPP riboswitch (THI element)	Cis-reg/riboswitch	236	223	94.49	201	85.17	4	4	100.00	4	100.00

Accession	Type [†]	Class	miPred					3SVM [‡]				
			N	TN	%SP	TN	%SP	N	TN	%SP	TN	%SP
RF00060	HgcE RNA	Gene	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00061	Hepatitis C virus IRES	Cis-reg/IRES	786	658	83.72	674	85.75	1	0	0.00	0	0.00
RF00062	HgcC family RNA	Gene	22	7	31.82	22	100.00	0	0	NaN	0	NaN
RF00063	SscA RNA	Gene	5	5	100.00	3	60.00	0	0	NaN	0	NaN
RF00064	HgcG RNA	Gene	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00065	snoR9 / snoR19 family	Gene/snRNA/guide/C/D-box	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00066	U7 small nuclear RNA	Gene/snRNA	28	24	85.71	24	85.71	7	7	100.00	7	100.00
RF00067	U15 small nucleolar RNA	Gene/snRNA/guide/C/D-box	18	16	88.89	15	83.33	2	1	50.00	2	100.00
RF00068	U21 small nucleolar RNA	Gene/snRNA/guide/C/D-box	5	5	100.00	4	80.00	3	3	100.00	1	33.33
RF00069	U24/Z20/U76 small nucleolar RNA	Gene/snRNA/guide/C/D-box	14	14	100.00	10	71.43	3	3	100.00	3	100.00
RF00070	Small nucleolar RNA U29	Gene/snRNA/guide/C/D-box	10	10	100.00	6	60.00	2	2	100.00	2	100.00
RF00071	U73 small nucleolar RNA	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	2	2	100.00	2	100.00
RF00072	U23 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	6	2	33.33	3	50.00	0	0	NaN	0	NaN
RF00077	SraB RNA	Gene/sRNA	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00078	SraD RNA	Gene/sRNA	5	5	100.00	4	80.00	0	0	NaN	0	NaN
RF00079	SraE/RygA/RygB family RNA	Gene/sRNA	6	5	83.33	4	66.67	0	0	NaN	0	NaN
RF00080	yybP-ykoY element	Cis-reg/riboswitch	74	52	70.27	70	94.59	2	2	100.00	1	50.00
RF00081	SraH RNA	Gene/sRNA	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00082	SraG RNA	Gene/sRNA	5	4	80.00	5	100.00	0	0	NaN	0	NaN
RF00083	SraJ RNA	Gene/sRNA	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00084	CsrC RNA family	Gene/sRNA	5	1	20.00	4	80.00	0	0	NaN	0	NaN
RF00085	U28 small nucleolar RNA	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	2	2	100.00	1	50.00
RF00086	U27/Z191/snR74/Z4 small nucleolar RNA	Gene/snRNA/guide/C/D-box	10	10	100.00	7	70.00	0	0	NaN	0	NaN
RF00087	U26 small nucleolar RNA	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	2	0	0.00	2	100.00
RF00088	U30 small nucleolar RNA	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	2	1	50.00	1	50.00
RF00089	U31 small nucleolar RNA	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	4	4	100.00	2	50.00
RF00090	U19 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	3	0	0.00	2	66.67	0	0	NaN	0	NaN
RF00091	Small nucleolar RNA E2/ACA6/M2/MBI-136	Gene/snRNA/guide/H/ACA-box	10	2	20.00	8	80.00	0	0	NaN	0	NaN
RF00092	E3 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	9	4	44.44	9	100.00	0	0	NaN	0	NaN
RF00093	U18 small nucleolar RNA	Gene/snRNA/guide/C/D-box	16	16	100.00	14	87.50	10	9	90.00	8	80.00
RF00094	Hepatitis delta virus ribozyme	Gene/ribozyme	15	14	93.33	15	100.00	0	0	NaN	0	NaN
RF00095	Pyrococcus C/D box small nucleolar RNA	Gene/snRNA/guide/C/D-box	38	38	100.00	37	97.37	18	18	100.00	17	94.44
RF00096	U8 small nucleolar RNA	Gene/snRNA/guide/C/D-box	5	2	40.00	3	60.00	0	0	NaN	0	NaN
RF00097	Plant small nucleolar RNA R71	Gene/snRNA/guide/C/D-box	21	18	85.71	21	100.00	0	0	NaN	0	NaN
RF00098	Snake H/ACA box small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	22	22	100.00	20	90.91	0	0	NaN	0	NaN
RF00099	U22 small nucleolar RNA	Gene/snRNA/guide/C/D-box	3	2	66.67	2	66.67	0	0	NaN	0	NaN
RF00100	7SK RNA	Gene	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00101	SraC/RyeA RNA	Gene/sRNA	7	3	42.86	7	100.00	0	0	NaN	0	NaN
RF00102	VA RNA	Gene	23	0	0.00	22	95.65	0	0	NaN	0	NaN
RF00105	HBII-52 small nucleolar RNA	Gene/snRNA/guide/C/D-box	23	23	100.00	14	60.87	1	1	100.00	1	100.00
RF00106	RNAI	Gene/antisense	10	0	0.00	6	60.00	0	0	NaN	0	NaN
RF00107	FinP	Gene	6	0	0.00	6	100.00	0	0	NaN	0	NaN
RF00108	HBII-85 small nucleolar RNA	Gene/snRNA/guide/C/D-box	7	7	100.00	7	100.00	0	0	NaN	0	NaN
RF00109	Vimentin 3' UTR protein-binding region	Cis-reg	12	12	100.00	11	91.67	2	2	100.00	2	100.00
RF00110	RybB RNA	Gene/sRNA	4	2	50.00	4	100.00	2	2	100.00	0	0.00
RF00111	RyeB RNA	Gene/sRNA	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00112	RyeE RNA	Gene/sRNA	3	3	100.00	2	66.67	0	0	NaN	0	NaN
RF00113	QUAD RNA	Gene/sRNA	15	6	40.00	15	100.00	0	0	NaN	0	NaN
RF00114	Ribosomal S15 leader	Cis-reg	11	11	100.00	7	63.64	0	0	NaN	0	NaN
RF00115	IS061 RNA	Gene/sRNA	5	5	100.00	2	40.00	0	0	NaN	0	NaN
RF00116	C0465 RNA	Gene/sRNA	3	3	100.00	2	66.67	0	0	NaN	0	NaN
RF00117	C0719 RNA	Gene/sRNA	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00118	rydB RNA	Gene/sRNA	5	5	100.00	4	80.00	5	5	100.00	0	0.00
RF00119	C0299 RNA	Gene/sRNA	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00120	C0343 RNA	Gene/sRNA	4	4	100.00	1	25.00	0	0	NaN	0	NaN
RF00121	MicC RNA	Gene/sRNA	4	3	75.00	3	75.00	0	0	NaN	0	NaN
RF00122	GadY	Gene/sRNA	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00124	IS102 RNA	Gene/sRNA	8	1	12.50	8	100.00	0	0	NaN	0	NaN
RF00125	IS128 RNA	Gene/sRNA	3	1	33.33	3	100.00	0	0	NaN	0	NaN

Accession	Type [†]	Class	miPred					3SVM [‡]				
			N	TN	%SP	TN	%SP	N	TN	%SP	TN	%SP
RF00191	U99 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00192	Bovine leukaemia virus RNA packaging signal	Cis-reg	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00193	Citrus tristeza virus replication signal	Cis-reg	9	9	100.00	9	100.00	0	0	NaN	0	NaN
RF00194	Rubella virus 3' cis-acting element	Cis-reg	9	9	100.00	9	100.00	0	0	NaN	0	NaN
RF00195	RsmY RNA family	Gene/snRNA	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00196	Alfalfa mosaic virus RNA 1 5' UTR stem-loop	Cis-reg	4	2	50.00	0	0.00	2	2	100.00	0	0.00
RF00197	rbcL 5' UTR RNA stabilising element	Cis-reg	3	2	66.67	3	100.00	0	0	NaN	0	NaN
RF00198	SL1 RNA	Gene	28	0	0.00	24	85.71	0	0	NaN	0	NaN
RF00199	SL2 RNA	Gene	32	10	31.25	24	75.00	0	0	NaN	0	NaN
RF00200	Small nucleolar RNA Z199	Gene/snRNA/guide/C/D-box	8	8	100.00	7	87.50	6	6	100.00	4	66.67
RF00201	Small nucleolar RNA Z278	Gene/snRNA/guide/C/D-box	7	5	71.43	7	100.00	7	7	100.00	5	71.43
RF00202	Small nucleolar RNA R66	Gene/snRNA/guide/C/D-box	6	6	100.00	6	100.00	1	1	100.00	1	100.00
RF00203	Small nucleolar RNA R160	Gene/snRNA/guide/C/D-box	9	9	100.00	9	100.00	4	4	100.00	4	100.00
RF00204	Small nucleolar RNA R12	Gene/snRNA/guide/C/D-box	9	9	100.00	8	88.89	2	2	100.00	2	100.00
RF00205	Small nucleolar RNA R41	Gene/snRNA/guide/C/D-box	7	7	100.00	6	85.71	7	7	100.00	1	14.29
RF00206	Small nucleolar RNA U54	Gene/snRNA/guide/C/D-box	13	13	100.00	11	84.62	1	1	100.00	1	100.00
RF00207	K10 transport/localisation element (TLS)	Cis-reg	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00208	Small nucleolar RNA R72	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00209	Pestivirus IRES	Cis-reg/IRES	25	1	4.00	20	80.00	0	0	NaN	0	NaN
RF00210	Aphovirus IRES	Cis-reg/IRES	32	2	6.25	29	90.63	0	0	NaN	0	NaN
RF00211	Small nucleolar RNA U35	Gene/snRNA/guide/C/D-box	8	8	100.00	5	62.50	1	1	100.00	0	0.00
RF00212	U38 small nucleolar RNA	Gene/snRNA/guide/C/D-box	7	7	100.00	6	85.71	3	3	100.00	2	66.67
RF00213	Small nucleolar RNA R38	Gene/snRNA/guide/C/D-box	12	10	83.33	11	91.67	6	6	100.00	3	50.00
RF00214	Retrovirus direct repeat 1 (dr1)	Cis-reg	25	24	96.00	21	84.00	1	0	0.00	1	100.00
RF00215	Tombus virus defective interfering (DI) RNA region 3	Cis-reg	48	48	100.00	34	70.83	6	6	100.00	6	100.00
RF00216	c-myc IRES	Cis-reg/IRES	23	23	100.00	21	91.30	0	0	NaN	0	NaN
RF00217	Small nucleolar RNA U20	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	4	4	100.00	3	75.00
RF00218	Small nucleolar RNA U40	Gene/snRNA/guide/C/D-box	9	9	100.00	9	100.00	8	8	100.00	4	50.00
RF00219	Small nucleolar RNA U32	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00220	Human rhinovirus internal cis-acting regulatory element	Cis-reg	12	12	100.00	12	100.00	10	10	100.00	10	100.00
RF00221	Small nucleolar RNA U43	Gene/snRNA/guide/C/D-box	6	5	83.33	3	50.00	3	2	66.67	3	100.00
RF00222	Bag-1 IRES	Cis-reg/IRES	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00223	bip IRES	Cis-reg/IRES	4	4	100.00	4	100.00	2	2	100.00	2	100.00
RF00224	FGF-2 IRES	Cis-reg/IRES	3	3	100.00	2	66.67	0	0	NaN	0	NaN
RF00225	Tobamovirus IRES	Cis-reg/IRES	7	7	100.00	7	100.00	0	0	NaN	0	NaN
RF00226	n-myc IRES	Cis-reg/IRES	6	6	100.00	6	100.00	0	0	NaN	0	NaN
RF00227	FIE3 (fitz instability element 3') element	Cis-reg	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00228	Hepatitis A virus IRES	Cis-reg/IRES	23	9	39.13	22	95.65	0	0	NaN	0	NaN
RF00229	Picornavirus IRES	Cis-reg/IRES	195	96	49.23	180	92.31	0	0	NaN	0	NaN
RF00230	T-box leader	Cis-reg	66	28	42.42	60	90.91	0	0	NaN	0	NaN
RF00231	U93 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00232	Spi-1 (PU.1) 5' UTR regulatory element	Cis-reg	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00233	Tymovirus/Pomovirus tRNA-like 3' UTR element	Cis-reg	27	27	100.00	23	85.19	0	0	NaN	0	NaN
RF00234	glmS glucosamine-6-phosphate activated ribozyme	Cis-reg/riboswitch	14	10	71.43	11	78.57	0	0	NaN	0	NaN
RF00235	Plasmid RNAlII	Gene	7	0	0.00	7	100.00	0	0	NaN	0	NaN
RF00236	ctRNA	Gene/antisense	17	0	0.00	16	94.12	0	0	NaN	0	NaN
RF00238	ctRNA	Gene/antisense	48	5	10.42	44	91.67	0	0	NaN	0	NaN
RF00240	RNA-OUT	Gene	7	0	0.00	3	42.86	7	2	28.57	3	42.86
RF00242	ctRNA	Gene/antisense	15	6	40.00	10	66.67	0	0	NaN	0	NaN
RF00243	traJ 5' UTR	Cis-reg	6	2	33.33	6	100.00	0	0	NaN	0	NaN
RF00250	Trans-activation response element (TAR)	Cis-reg	416	26	6.25	370	88.94	412	49	11.89	221	53.64
RF00252	Alfalfa mosaic virus coat protein binding (CPB) RNA	Cis-reg	18	2	11.11	18	100.00	0	0	NaN	0	NaN
RF00259	Interferon gamma 5' UTR regulatory element	Cis-reg	5	5	100.00	2	40.00	0	0	NaN	0	NaN
RF00260	Hepatitis C virus (HCV) cis-acting replication element	Cis-reg	52	52	100.00	52	100.00	52	52	100.00	46	88.46
RF00261	L-myc IRES	Cis-reg/IRES	2	2	100.00	2	100.00	0	0	NaN	0	NaN
RF00262	sar RNA	Gene	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00263	U68 small nucleolar RNA	Gene/snRNA/guide/H/ACA-box	4	3	75.00	3	75.00	0	0	NaN	0	NaN
RF00264	Small nucleolar RNA U64	Gene/snRNA/guide/H/ACA-box	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00265	Small nucleolar RNA U69	Gene/snRNA/guide/H/ACA-box	3	1	33.33	2	66.67	0	0	NaN	0	NaN

Accession	Type [†]	Class	miPred					3SVM [‡]				
			N	TN	%SP	TN	%SP	N	TN	%SP	TN	%SP
RF00326	Small nucleolar RNA Z155	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00327	Small nucleolar RNA Z194	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00328	Small nucleolar RNA Z161/Z228	Gene/snRNA/guide/C/D-box	7	7	100.00	5	71.43	2	2	100.00	2	100.00
RF00329	Small nucleolar RNA Z162	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00330	Small nucleolar RNA Z43	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	2	2	100.00	1	50.00
RF00331	Small nucleolar RNA Z169	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	2	2	100.00	2	100.00
RF00332	Small nucleolar RNA Z266	Gene/snRNA/guide/C/D-box	4	4	100.00	2	50.00	2	2	100.00	0	0.00
RF00333	Small nucleolar RNA Z157/R69/R10	Gene/snRNA/guide/C/D-box	10	8	80.00	5	50.00	2	2	100.00	0	0.00
RF00334	Small nucleolar RNA MBI-28	Gene/snRNA/guide/H/ACA-box	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00335	Small nucleolar RNA Z13/snr52	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	3	3	100.00	3	100.00
RF00336	Small nucleolar RNA J26	Gene/snRNA/guide/C/D-box	5	5	100.00	3	60.00	0	0	NaN	0	NaN
RF00337	Small nucleolar RNA Z112	Gene/snRNA/guide/C/D-box	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00338	Small nucleolar RNA snR53	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	1	0	0.00	1	100.00
RF00339	Small nucleolar RNA snoR60	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00340	Small nucleolar RNA snoMBI-87	Gene/snRNA/guide/H/ACA-box	6	0	0.00	4	66.67	0	0	NaN	0	NaN
RF00341	Small nucleolar RNA Z39	Gene/snRNA/guide/C/D-box	5	5	100.00	5	100.00	4	4	100.00	4	100.00
RF00342	Small nucleolar RNA Z40	Gene/snRNA/guide/C/D-box	5	5	100.00	3	60.00	5	5	100.00	5	100.00
RF00343	Small nucleolar RNA Z122	Gene/snRNA/guide/C/D-box	3	3	100.00	0	0.00	1	1	100.00	1	100.00
RF00344	Small nucleolar RNA Z267	Gene/snRNA/guide/C/D-box	5	5	100.00	3	60.00	2	2	100.00	2	100.00
RF00345	Small nucleolar RNA snoR1	Gene/snRNA/guide/C/D-box	7	7	100.00	6	85.71	3	3	100.00	1	33.33
RF00346	Small nucleolar RNA snoZ1	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00347	Small nucleolar RNA Z50	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	1	1	100.00	0	0.00
RF00348	Small nucleolar RNA snoR9	Gene/snRNA/guide/C/D-box	7	7	100.00	6	85.71	3	3	100.00	3	100.00
RF00349	Small nucleolar RNA R11/Z151	Gene/snRNA/guide/C/D-box	5	5	100.00	3	60.00	0	0	NaN	0	NaN
RF00350	Small nucleolar RNA Z152/R70/R12/	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00351	Small nucleolar RNA R20	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	2	2	100.00	0	0.00
RF00352	Small nucleolar RNA R21	Gene/snRNA/guide/C/D-box	4	4	100.00	2	50.00	0	0	NaN	0	NaN
RF00353	Small nucleolar RNA snoR31/Z110/Z27	Gene/snRNA/guide/C/D-box	8	5	62.50	2	25.00	0	0	NaN	0	NaN
RF00355	Small nucleolar RNA snoR28	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00356	Small nucleolar RNA R32/R81/Z41	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	4	4	100.00	1	25.00
RF00357	Small nucleolar RNA R44/J54	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	3	3	100.00	3	100.00
RF00358	Small nucleolar RNA Z101	Gene/snRNA/guide/C/D-box	3	3	100.00	2	66.67	2	2	100.00	0	0.00
RF00359	Small nucleolar RNA Z102/R77	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	2	2	100.00	2	100.00
RF00360	Small nucleolar RNA Z107/R87	Gene/snRNA/guide/C/D-box	6	5	83.33	6	100.00	0	0	NaN	0	NaN
RF00361	Small nucleolar RNA Z119	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00362	Pospiviroid RY motif stem loop	Cis-reg	16	14	87.50	15	93.75	11	11	100.00	3	27.27
RF00368	sroB RNA	Gene/sRNA	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00369	sroC RNA	Gene/sRNA	5	0	0.00	4	80.00	0	0	NaN	0	NaN
RF00370	sroD RNA	Gene/sRNA	3	2	66.67	3	100.00	0	0	NaN	0	NaN
RF00371	sroE RNA	Gene/sRNA	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00372	sroH RNA	Gene/sRNA	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00373	Archaeal RNase P	Gene/ribozyme	40	16	40.00	33	82.50	0	0	NaN	0	NaN
RF00374	Gammaretrovirus core encapsidation signal	Cis-reg	23	11	47.83	23	100.00	0	0	NaN	0	NaN
RF00375	HIV primer binding site (PBS)	Cis-reg	373	265	71.05	334	89.54	0	0	NaN	0	NaN
RF00376	HIV gag stem loop 3 (GSL3)	Cis-reg	1374	1371	99.78	1200	87.34	9	9	100.00	4	44.44
RF00377	Small nucleolar RNA U6-53/MBII-28	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00378	Qrr RNA	Gene/sRNA	14	7	50.00	9	64.29	0	0	NaN	0	NaN
RF00379	ydaO/yuaA element	Cis-reg/riboswitch	35	35	100.00	32	91.43	0	0	NaN	0	NaN
RF00380	ykoK element	Cis-reg/riboswitch	39	25	64.10	32	82.05	0	0	NaN	0	NaN
RF00381	Antizyme RNA frameshifting stimulation element	Cis-reg/frameshift	13	12	92.31	12	92.31	10	10	100.00	7	70.00
RF00382	DnaX ribosomal frameshifting element	Cis-reg/frameshift	3	3	100.00	2	66.67	0	0	NaN	0	NaN
RF00383	Insertion sequence IS1222 ribosomal frameshifting element	Cis-reg/frameshift	6	6	100.00	6	100.00	0	0	NaN	0	NaN
RF00384	Poxvirus AX element late mRNA CE	Cis-reg	7	7	100.00	7	100.00	0	0	NaN	0	NaN
RF00385	Infectious bronchitis virus D-RNA	Cis-reg	10	6	60.00	10	100.00	10	8	80.00	6	60.00
RF00386	Enterovirus 5' cloverleaf cis-acting replication element	Cis-reg	60	5	8.33	52	86.67	0	0	NaN	0	NaN
RF00387	FGF-1 IRES	Cis-reg/IRES	6	6	100.00	6	100.00	0	0	NaN	0	NaN
RF00388	Qa RNA	Gene/antisense	5	2	40.00	3	60.00	0	0	NaN	0	NaN
RF00389	Bamboo mosaic virus satellite RNA CE	Cis-reg	42	42	100.00	41	97.62	0	0	NaN	0	NaN
RF00390	UPSK RNA	Cis-reg	4	4	100.00	4	100.00	0	0	NaN	0	NaN

De novo SVM classification of precursor microRNAs

Accession	Type [†]	Class	miPred					3SVM [‡]				
			N	TN	%SP	TN	%SP	N	TN	%SP	TN	%SP
RF00391	RtT RNA	Cis-reg	19	16	84.21	18	94.74	0	0	NaN	0	NaN
RF00392	Small nucleolar RNA ACA5	Gene/snRNA/guide/H/ACA-box	6	6	100.00	4	66.67	0	0	NaN	0	NaN
RF00393	Small nucleolar RNA ACA8	Gene/snRNA/guide/H/ACA-box	5	4	80.00	4	80.00	0	0	NaN	0	NaN
RF00394	Small nucleolar RNA ACA4	Gene/snRNA/guide/H/ACA-box	7	4	57.14	7	100.00	0	0	NaN	0	NaN
RF00395	Small nucleolar RNA ACA10	Gene/snRNA/guide/H/ACA-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00396	Small nucleolar RNA ACA13	Gene/snRNA/guide/H/ACA-box	3	0	0.00	1	33.33	0	0	NaN	0	NaN
RF00397	Small nucleolar RNA ACA14	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00398	Small nucleolar RNA ACA15	Gene/snRNA/guide/H/ACA-box	4	2	50.00	4	100.00	0	0	NaN	0	NaN
RF00399	Small nucleolar RNA ACA24	Gene/snRNA/guide/H/ACA-box	5	5	100.00	4	80.00	0	0	NaN	0	NaN
RF00400	Small nucleolar RNA ACA28	Gene/snRNA/guide/H/ACA-box	3	2	66.67	3	100.00	0	0	NaN	0	NaN
RF00401	Small nucleolar RNA ACA20	Gene/snRNA/guide/H/ACA-box	17	4	23.53	14	82.35	0	0	NaN	0	NaN
RF00402	Small nucleolar RNA ACA25	Gene/snRNA/guide/H/ACA-box	9	7	77.78	8	88.89	0	0	NaN	0	NaN
RF00403	Small nucleolar RNA ACA41	Gene/snRNA/guide/H/ACA-box	6	1	16.67	6	100.00	0	0	NaN	0	NaN
RF00404	Small nucleolar RNA ACA46	Gene/snRNA/guide/H/ACA-box	3	1	33.33	2	66.67	0	0	NaN	0	NaN
RF00405	Small nucleolar RNA ACA44	Gene/snRNA/guide/H/ACA-box	6	6	100.00	6	100.00	1	1	100.00	1	100.00
RF00406	Small nucleolar RNA ACA42	Gene/snRNA/guide/H/ACA-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00407	Small nucleolar RNA ACA50	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00408	Small nucleolar RNA ACA1	Gene/snRNA/guide/H/ACA-box	6	5	83.33	5	83.33	0	0	NaN	0	NaN
RF00409	Small nucleolar RNA ACA7	Gene/snRNA/guide/H/ACA-box	8	8	100.00	6	75.00	1	1	100.00	1	100.00
RF00410	Small nucleolar RNA ACA2/ACA34	Gene/snRNA/guide/H/ACA-box	18	5	27.78	16	88.89	0	0	NaN	0	NaN
RF00411	Small nucleolar RNA ACA9	Gene/snRNA/guide/H/ACA-box	6	3	50.00	5	83.33	0	0	NaN	0	NaN
RF00412	Small nucleolar RNA ACA21	Gene/snRNA/guide/H/ACA-box	5	1	20.00	3	60.00	0	0	NaN	0	NaN
RF00413	Small nucleolar RNA ACA19	Gene/snRNA/guide/H/ACA-box	4	1	25.00	3	75.00	0	0	NaN	0	NaN
RF00414	Small nucleolar RNA ACA22	Gene/snRNA/guide/H/ACA-box	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00415	Small nucleolar RNA ACA30/ACA37/MBI-26	Gene/snRNA/guide/H/ACA-box	6	6	100.00	6	100.00	0	0	NaN	0	NaN
RF00416	Small nucleolar RNA ACA43	Gene/snRNA/guide/H/ACA-box	7	7	100.00	6	85.71	0	0	NaN	0	NaN
RF00417	Small nucleolar RNA ACA56	Gene/snRNA/guide/H/ACA-box	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00418	Small nucleolar RNA ACA52	Gene/snRNA/guide/H/ACA-box	4	0	0.00	3	75.00	0	0	NaN	0	NaN
RF00419	Small nucleolar RNA ACA52	Gene/snRNA/guide/H/ACA-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00420	Small nucleolar RNA ACA61	Gene/snRNA/guide/H/ACA-box	4	3	75.00	3	75.00	0	0	NaN	0	NaN
RF00421	Small nucleolar RNA ACA32	Gene/snRNA/guide/H/ACA-box	9	6	66.67	6	66.67	0	0	NaN	0	NaN
RF00422	Small nucleolar RNA ACA12	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00423	Small nucleolar RNA ACA26	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00424	Small nucleolar RNA ACA47	Gene/snRNA/guide/H/ACA-box	6	2	33.33	4	66.67	0	0	NaN	0	NaN
RF00425	Small nucleolar RNA ACA18	Gene/snRNA/guide/H/ACA-box	6	3	50.00	3	50.00	0	0	NaN	0	NaN
RF00426	Small nucleolar RNA ACA45	Gene/snRNA/guide/H/ACA-box	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00427	Small nucleolar RNA ACA11	Gene/snRNA/guide/H/ACA-box	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00428	Small nucleolar RNA ACA38	Gene/snRNA/guide/H/ACA-box	5	4	80.00	5	100.00	0	0	NaN	0	NaN
RF00429	Small nucleolar RNA ACA29	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00430	Small nucleolar RNA ACA54	Gene/snRNA/guide/H/ACA-box	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00431	Small nucleolar RNA ACA55	Gene/snRNA/guide/H/ACA-box	3	0	0.00	3	100.00	0	0	NaN	0	NaN
RF00432	Small nucleolar RNA ACA51	Gene/snRNA/guide/H/ACA-box	9	8	88.89	9	100.00	0	0	NaN	0	NaN
RF00433	Hsp90 CE	Cis-reg/thermoregulator	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00434	Luteovirus cap-independent translation element (BTE)	Cis-reg	17	17	100.00	13	76.47	0	0	NaN	0	NaN
RF00435	Repression of heat shock gene expression (ROSE) element	Cis-reg/thermoregulator	3	2	66.67	2	66.67	0	0	NaN	0	NaN
RF00436	UnaL2 line 3' element	Cis-reg	144	141	97.92	113	78.47	50	49	98.00	13	26.00
RF00437	Hairy RNA localisation element (HLE)	Cis-reg	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00438	Small nucleolar RNA ACA33	Gene/snRNA/guide/H/ACA-box	5	5	100.00	4	80.00	0	0	NaN	0	NaN
RF00439	Small nucleolar RNA U87	Gene/snRNA/guide/C/D-box	4	4	100.00	3	75.00	0	0	NaN	0	NaN
RF00440	Small nucleolar RNA U37	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	3	3	100.00	3	100.00
RF00441	Small nucleolar RNA Z242	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00442	ykkC-ykkD element	Cis-reg/riboswitch	16	15	93.75	14	87.50	0	0	NaN	0	NaN
RF00443	Small nucleolar RNA ACA27	Gene/snRNA/guide/H/ACA-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00444	PrrF RNA	Gene/snRNA	7	2	28.57	7	100.00	0	0	NaN	0	NaN
RF00447	Voltage-gated potassium-channel Kv1.4 IRES	Cis-reg/IRES	6	5	83.33	6	100.00	0	0	NaN	0	NaN
RF00448	Epstein-Barr virus nuclear antigen (EBNA) IRES	Cis-reg/IRES	8	8	100.00	8	100.00	0	0	NaN	0	NaN
RF00449	HIF-1 alpha IRES	Cis-reg/IRES	7	7	100.00	7	100.00	0	0	NaN	0	NaN
RF00450	Small nucleolar RNA R105/R108	Gene/snRNA/guide/C/D-box	4	3	75.00	4	100.00	0	0	NaN	0	NaN

Accession	Type [†]	Class	miPred					3SVM [‡]				
			N	TN	%SP	TN	%SP	N	TN	%SP	TN	%SP
RF00453	Cardiovirus cis-acting replication element	Cis-reg	12	11	91.67	9	75.00	2	2	100.00	2	100.00
RF00454	p27 CE	Cis-reg	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00457	Mnt IRES	Cis-reg/IRES	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00458	Cripavirus IRES	Cis-reg/IRES	7	6	85.71	6	85.71	0	0	NaN	0	NaN
RF00459	Mason-Pfizer monkey virus packaging signal	Cis-reg	4	4	100.00	4	100.00	0	0	NaN	0	NaN
RF00460	U1A polyadenylation inhibition element (PIE)	Cis-reg	6	6	100.00	6	100.00	3	3	100.00	3	100.00
RF00461	Vascular endothelial growth factor (VEGF) IRES A	Cis-reg/IRES	7	7	100.00	7	100.00	0	0	NaN	0	NaN
RF00462	APC IRES	Cis-reg/IRES	6	6	100.00	2	33.33	0	0	NaN	0	NaN
RF00463	Apolipoprotein B (apoB) 5' UTR CE	Cis-reg	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00465	Japanese encephalitis virus (JEV) hairpin structure	Cis-reg	20	19	95.00	19	95.00	12	12	100.00	5	41.67
RF00466	Agrobacterium tumefaciens ROSE element	Cis-reg/thermoregulator	3	1	33.33	3	100.00	0	0	NaN	0	NaN
RF00467	Rous sarcoma virus (RSV) primer binding site (PBS)	Cis-reg	23	1	4.35	21	91.30	22	13	59.09	18	81.82
RF00468	Hepatitis C stem-loop VII	Cis-reg	63	9	14.29	32	50.79	63	45	71.43	63	100.00
RF00469	Hepatitis C stem-loop IV	Cis-reg	109	2	1.83	109	100.00	109	109	100.00	61	55.96
RF00470	Togavirus 5' plus strand CE	Cis-reg	32	5	15.63	29	90.63	0	0	NaN	0	NaN
RF00471	Small nucleolar RNA snR48	Gene/snRNA/guide/C/D-box	6	6	100.00	5	83.33	1	1	100.00	0	0.00
RF00472	Small nucleolar RNA snR55/Z10	Gene/snRNA/guide/C/D-box	7	7	100.00	4	57.14	0	0	NaN	0	NaN
RF00473	Small nucleolar RNA snR54	Gene/snRNA/guide/C/D-box	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00474	Small nucleolar RNA snR57	Gene/snRNA/guide/C/D-box	6	6	100.00	5	83.33	2	2	100.00	0	0.00
RF00475	Small nucleolar RNA snR69	Gene/snRNA/guide/C/D-box	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00476	Small nucleolar RNA snR61/Z11	Gene/snRNA/guide/C/D-box	9	9	100.00	8	88.89	0	0	NaN	0	NaN
RF00477	Small nucleolar RNA snR66	Gene/snRNA/guide/C/D-box	5	5	100.00	5	100.00	0	0	NaN	0	NaN
RF00478	Small nucleolar RNA U88	Gene/snRNA/guide/C/D-box	4	0	0.00	3	75.00	0	0	NaN	0	NaN
RF00479	Small nucleolar RNA snR71	Gene/snRNA/guide/C/D-box	5	5	100.00	3	60.00	0	0	NaN	0	NaN
RF00480	HIV Ribosomal frameshift signal	Cis-reg/frameshift	768	152	19.79	704	91.67	765	719	93.99	107	13.99
RF00481	Hepatitis C virus 3'X element	Cis-reg	22	0	0.00	13	59.09	0	0	NaN	0	NaN
RF00482	Small nucleolar RNA F1/F2/snoR5a	Gene/snRNA/guide/H/ACA-box	8	5	62.50	6	75.00	0	0	NaN	0	NaN
RF00483	Insulin-like growth factor II IRES	Cis-reg/IRES	8	8	100.00	7	87.50	0	0	NaN	0	NaN
RF00484	Connexin-32 IRES	Cis-reg/IRES	6	6	100.00	5	83.33	0	0	NaN	0	NaN
RF00485	Potassium channel RNA editing signal	Cis-reg	85	76	89.41	69	81.18	13	10	76.92	7	53.85
RF00487	Connexin-43 IRES	Cis-reg/IRES	13	13	100.00	12	92.31	0	0	NaN	0	NaN
RF00488	Yeast U1 spliceosomal RNA	Gene/snRNA/splicing	6	0	0.00	5	83.33	0	0	NaN	0	NaN
RF00489	ctRNA	Gene/antisense	15	6	40.00	14	93.33	10	8	80.00	7	70.00
RF00490	S-element	Cis-reg	13	13	100.00	9	69.23	3	3	100.00	3	100.00
RF00491	Simian virus 40 late polyadenylation signal (SVLPA)	Cis-reg	3	3	100.00	2	66.67	0	0	NaN	0	NaN
RF00492	Small nucleolar RNA U12-22	Gene/snRNA/guide/C/D-box	7	7	100.00	6	85.71	3	3	100.00	3	100.00
RF00493	Small nucleolar RNA U2-30	Gene/snRNA/guide/C/D-box	3	3	100.00	3	100.00	0	0	NaN	0	NaN
RF00494	Small nucleolar RNA U2-19	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	1	1	100.00	1	100.00
RF00495	Heat shock protein 70 (Hsp70) IRES	Cis-reg/IRES	13	13	100.00	13	100.00	0	0	NaN	0	NaN
RF00496	Coronavirus SL-III cis-acting replication element	Cis-reg	5	5	100.00	5	100.00	3	3	100.00	1	33.33
RF00497	Dengue virus 3'-SL cis-acting replication element	Cis-reg	23	5	21.74	21	91.30	0	0	NaN	0	NaN
RF00498	Equine arteritis virus leader TRS hairpin (LTH)	Cis-reg	4	4	100.00	4	100.00	4	4	100.00	4	100.00
RF00499	Human parechovirus 1 (HPeV1) cis regulatory element	Cis-reg	5	2	40.00	5	100.00	0	0	NaN	0	NaN
RF00500	Turnip crinkle virus (TCV) repressor of minus strand synthesis H5	Cis-reg	3	2	66.67	3	100.00	3	3	100.00	2	66.67
RF00501	Rotavirus cis-acting replication element	Cis-reg	14	14	100.00	8	57.14	4	4	100.00	1	25.00
RF00502	Turnip crinkle virus (TCV) core promoter hairpin (Pr)	Cis-reg	4	4	100.00	4	100.00	4	4	100.00	2	50.00
RF00503	RNAIII	Gene	12	2	16.67	12	100.00	0	0	NaN	0	NaN
RF00504	gcvT element	Cis-reg/riboswitch	117	111	94.87	102	87.18	3	3	100.00	2	66.67
RF00505	RydC RNA	Gene/sRNA	3	3	100.00	3	100.00	2	2	100.00	2	100.00
RF00506	Threonine operon leader	Cis-reg	27	1	3.70	25	92.59	0	0	NaN	0	NaN
RF00507	Coronavirus frameshifting stimulation element	Cis-reg/frameshift	18	12	66.67	15	83.33	0	0	NaN	0	NaN
RF00509	Small nucleolar RNA snR64	Gene/snRNA/guide/C/D-box	4	4	100.00	4	100.00	0	0	NaN	0	NaN
-	mRNAs	-	31	27	87.10	27	87.10	0	0	NaN	0	NaN
Total ncRNA samples (exclude mRNAs)			-	-	-	-	-	2404	1884	-	1199	-

†, cis-regulatory element (CE); internal ribosome entry site (IRES). N (non pre-miRs), TN (non pre-miRs missed), and %SP (Specificity). ‡, 3SVM model was trained on 200 human pre-miRs and 400 pseudo hairpins randomly selected using the latest libSVM 2.82 (the "-b 1" option was enabled) and the optimal hyperparameter pair (C, γ).

Table S4. The mean specificity of *miPred*, *miPred-NBC*, *3SVM*, and *3SVM-NBC* evaluated on the non *pre-miR* dataset IE-NC (12,387 functional ncRNAs) categorized by classes of ncRNAs.

Classes of ncRNAs	No. of types	miPred		3SVM [†]	
		%SP	%SP	No. of excluded types	%SP
<i>Cis-reg</i>	77	74.91 ± 4.03	87.99 ± 2.03	46	83.36 ± 5.60
<i>Cis-reg frameshift</i>	5	75.75 ± 15.27	86.80 ± 5.68	3	96.99 ± 3.01
<i>Cis-reg IRES</i>	24	85.47 ± 6.02	91.02 ± 3.06	22	50.00 ± 50.00
<i>Cis-reg riboswitch</i>	12	82.28 ± 3.96	85.77 ± 2.56	8	100.00 ± 0.00
<i>Cis-reg thermoregulator</i>	4	75.00 ± 15.96	91.67 ± 8.33	3	100.00 ± 0.00
<i>Gene</i>	24	34.73 ± 7.71	86.65 ± 3.03	18	70.57 ± 18.19
<i>Gene antisense</i>	10	41.93 ± 13.01	78.05 ± 5.03	8	90.00 ± 10.00
<i>Gene ribozyme</i>	9	60.08 ± 10.10	91.54 ± 2.36	6	97.44 ± 2.56
<i>Gene rRNA</i>	3	70.66 ± 12.94	90.74 ± 1.70	1	100.00 ± 0.00
<i>Gene snRNA</i>	1	85.71 ± 0.00	85.71 ± 0.00	0	100.00 ± 0.00
<i>Gene snRNA guide C/D-box</i>	165	94.61 ± 1.28	84.59 ± 1.58	72	92.78 ± 2.32
<i>Gene snRNA guide H/ACA-box</i>	71	60.97 ± 4.33	84.97 ± 2.04	68	100.00 ± 0.00
<i>Gene snRNA splicing</i>	7	51.16 ± 13.89	87.30 ± 3.83	6	100.00 ± 0.00
<i>Gene sRNA</i>	42	65.71 ± 5.90	87.53 ± 2.81	39	100.00 ± 0.00
<i>Gene tRNA</i>	1	85.55 ± 0.00	86.98 ± 0.00	0	94.94 ± 0.00
<i>Intron</i>	2	40.95 ± 9.05	86.70 ± 9.98	2	NaN

[†], *3SVM* model was trained on 200 human *pre-miRs* and 400 pseudo hairpins randomly selected using the latest libSVM 2.82 (the "-b 1" option was enabled) and the optimal hyperparameter pair (C, γ). %SP (Specificity). Values are expressed as mean ± standard error.

Table S5. F1 and F2 scores for features of *miPred* and *3SVM*, sorted by descending F1 scores.

<i>miPred</i>					<i>3SVM</i> [†]				
Rank	Features	F1 score	F2 score	$\Delta F = F1 - F2$	Features	F1 score	F2 score	$\Delta F = F1 - F2$	
01	<i>MFEI</i> ₁	1.28	1.52	-2.42E ⁻⁰¹	A(((8.20E ⁻⁰¹	6.97E ⁻⁰¹	1.22E ⁻⁰¹	
02	<i>zG</i>	1.27	1.48	-2.15E ⁻⁰¹	U(((7.58E ⁻⁰¹	6.12E ⁻⁰¹	1.46E ⁻⁰¹	
03	<i>dP</i>	1.03	1.18	-1.49E ⁻⁰¹	G...	4.57E ⁻⁰¹	2.05E ⁻⁰¹	2.52E ⁻⁰¹	
04	<i>zP</i>	9.67E ⁻⁰¹	1.03	-6.33E ⁻⁰²	A...	4.42E ⁻⁰¹	1.94E ⁻⁰¹	2.47E ⁻⁰¹	
05	<i>zQ</i>	8.33E ⁻⁰¹	7.29E ⁻⁰¹	1.04E ⁻⁰¹	C...	4.31E ⁻⁰¹	1.84E ⁻⁰¹	2.47E ⁻⁰¹	
06	<i>dG</i>	8.23E ⁻⁰¹	7.50E ⁻⁰¹	7.31E ⁻⁰²	G.(3.81E ⁻⁰¹	1.62E ⁻⁰¹	2.20E ⁻⁰¹	
07	<i>dQ</i>	7.99E ⁻⁰¹	6.67E ⁻⁰¹	1.32E ⁻⁰¹	A(..	3.50E ⁻⁰¹	1.31E ⁻⁰¹	2.19E ⁻⁰¹	
08	<i>zD</i>	7.92E ⁻⁰¹	6.70E ⁻⁰¹	1.23E ⁻⁰¹	A.(3.28E ⁻⁰¹	1.17E ⁻⁰¹	2.11E ⁻⁰¹	
09	<i>dD</i>	7.46E ⁻⁰¹	5.91E ⁻⁰¹	1.55E ⁻⁰¹	C(.	3.19E ⁻⁰¹	1.12E ⁻⁰¹	2.07E ⁻⁰¹	
10	<i>MFEI</i> ₂	4.41E ⁻⁰¹	1.53E ⁻⁰¹	2.88E ⁻⁰¹	G(..	3.07E ⁻⁰¹	9.75E ⁻⁰²	2.10E ⁻⁰¹	
11	%UA	3.87E ⁻⁰¹	1.56E ⁻⁰¹	2.31E ⁻⁰¹	U...	3.05E ⁻⁰¹	9.74E ⁻⁰²	2.08E ⁻⁰¹	
12	%G+C	3.06E ⁻⁰¹	1.04E ⁻⁰¹	2.02E ⁻⁰¹	C.(2.97E ⁻⁰¹	9.54E ⁻⁰²	2.02E ⁻⁰¹	
13	<i>zF</i>	2.88E ⁻⁰¹	7.13E ⁻⁰²	2.16E ⁻⁰¹	G(((2.84E ⁻⁰¹	8.95E ⁻⁰²	1.94E ⁻⁰¹	
14	%UU	2.83E ⁻⁰¹	8.91E ⁻⁰²	1.94E ⁻⁰¹	C..	2.70E ⁻⁰¹	7.93E ⁻⁰²	1.91E ⁻⁰¹	
15	%GU	2.64E ⁻⁰¹	7.71E ⁻⁰²	1.87E ⁻⁰¹	G(.	2.63E ⁻⁰¹	7.62E ⁻⁰²	1.87E ⁻⁰¹	
16	%GC	2.44E ⁻⁰¹	6.57E ⁻⁰²	1.79E ⁻⁰¹	G..	2.48E ⁻⁰¹	6.69E ⁻⁰²	1.81E ⁻⁰¹	
17	<i>dF</i>	2.42E ⁻⁰¹	5.16E ⁻⁰²	1.90E ⁻⁰¹	U.(2.19E ⁻⁰¹	5.20E ⁻⁰²	1.67E ⁻⁰¹	
18	%CC	2.04E ⁻⁰¹	4.59E ⁻⁰²	1.58E ⁻⁰¹	C.(1.89E ⁻⁰¹	3.92E ⁻⁰²	1.50E ⁻⁰¹	
19	%AA	1.83E ⁻⁰¹	3.73E ⁻⁰²	1.46E ⁻⁰¹	C(((1.87E ⁻⁰¹	3.88E ⁻⁰²	1.48E ⁻⁰¹	
20	%GG	1.82E ⁻⁰¹	3.68E ⁻⁰²	1.45E ⁻⁰¹	G.(1.82E ⁻⁰¹	3.52E ⁻⁰²	1.47E ⁻⁰¹	
21	%CA	1.77E ⁻⁰¹	3.48E ⁻⁰²	1.42E ⁻⁰¹	U.(1.71E ⁻⁰¹	2.88E ⁻⁰²	1.42E ⁻⁰¹	
22	%CG	1.73E ⁻⁰¹	3.30E ⁻⁰²	1.40E ⁻⁰¹	U(..	1.56E ⁻⁰¹	2.69E ⁻⁰²	1.30E ⁻⁰¹	
23	%GA	1.41E ⁻⁰¹	2.13E ⁻⁰²	1.19E ⁻⁰¹	U.(1.37E ⁻⁰¹	2.08E ⁻⁰²	1.16E ⁻⁰¹	
24	%AU	1.25E ⁻⁰¹	1.69E ⁻⁰²	1.08E ⁻⁰¹	A.(1.22E ⁻⁰¹	1.52E ⁻⁰²	1.07E ⁻⁰¹	
25	%AG	1.08E ⁻⁰¹	1.28E ⁻⁰²	9.54E ⁻⁰²	C.(1.10E ⁻⁰¹	1.32E ⁻⁰²	9.68E ⁻⁰²	
26	%UG	6.31E ⁻⁰²	4.42E ⁻⁰³	5.87E ⁻⁰²	G(.	1.02E ⁻⁰¹	1.13E ⁻⁰²	9.05E ⁻⁰²	
27	%AC	3.71E ⁻⁰²	1.53E ⁻⁰³	3.55E ⁻⁰²	C(.	6.68E ⁻⁰²	4.95E ⁻⁰³	6.19E ⁻⁰²	
28	%CU	3.21E ⁻⁰²	1.13E ⁻⁰³	3.09E ⁻⁰²	A(.	6.06E ⁻⁰²	4.06E ⁻⁰³	5.65E ⁻⁰²	
29	%UC	2.18E ⁻⁰²	5.21E ⁻⁰⁴	2.13E ⁻⁰²	A.(5.90E ⁻⁰²	3.87E ⁻⁰³	5.52E ⁻⁰²	
30	-	-	-	-	A(.	3.21E ⁻⁰²	1.14E ⁻⁰³	3.10E ⁻⁰²	
31	-	-	-	-	U.(3.28E ⁻⁰³	1.20E ⁻⁰⁵	3.26E ⁻⁰³	
32	-	-	-	-	U((.	6.80E ⁻⁰⁵	0.00E ⁻⁰⁰	6.80E ⁻⁰⁵	
		0.429 ± 0.0711	0.332 ± 0.0872	-			0.252 ± 0.0336	0.103 ± 0.0277	-

[†], *3SVM* model was trained on 200 human *pre-miRs* and 400 pseudo hairpins randomly selected using the latest libSVM 2.82 (the "-b 1" option was enabled) and the optimal hyperparameter pair (C, γ).

Table S6. Effects of feature selection on *miPred*'s accuracy.

Classifiers	Human pre-miRs (TR-H and TE-H)	Non-human pre-miRs (IE-NH)	ncRNAs (IE-NC)	mRNAs (IE-M)
<i>miPred</i>	93.60	95.64	68.68	87.10
<i>miPred</i> ₃	94.12	95.69	68.31	87.10
<i>miPred</i> _{3/5}	92.67	95.36	71.20	100.00
<i>miPred</i> _{3/10}	93.40	95.64	69.82	83.87
<i>miPred</i> _{3/15}	93.40	95.79	60.93	80.65
<i>miPred</i> _{3/20}	92.67	94.68	72.18	100.00
<i>miPred</i> _{3/21}	92.67	95.29	72.01	100.00
<i>miPred</i> _{3/22}	92.57	95.15	71.26	100.00
<i>miPred</i> _{3/23}	92.67	95.22	70.15	100.00
<i>miPred</i> _{3/24}	92.98	95.39	64.56	100.00
<i>miPred</i> _{3/25}	91.64	93.52	63.16	96.77
<i>miPred</i> _I	77.30	76.35	67.53	90.32
<i>miPred</i> _{II}	93.81	95.83	61.38	54.84
<i>miPred</i> _{III}	93.60	95.69	66.13	70.97

*miPred*₃ contains a subset of 26 features from *miPred* that excludes *dQ*, *dD*, and *zD*. Derived from *miPred*₃, the remaining nine variants denoted as *miPred*_{3/5}, *miPred*_{3/10}, ..., *miPred*_{3/24}, and *miPred*_{3/25} only include the top ranking 21, 16, 11, 6, 5, 4, 3, 2, and 1 feature(s), respectively. *miPred*_I (17 features: 16 dinucleotides frequencies and %G+C), *miPred*_{II} (12 features: *MFEI*₁, *MFEI*₂, *dP*, *dG*, *dQ*, *dD*, *dF*, *zP*, *zG*, *zQ*, *zD*, and *zF*), and *miPred*_{III} (9 features; a subset of *miPred*_{II} that excludes *dQ*, *dD*, and *zD*).

-	119625	94	AUCCUCGGCGACGGCGUGCAGUCGGGGCGUUUAGACACGCGGCC <u>gccuuaa</u> GGCCGAGUCCACCGUCGCGCCGAAGAGGACACCGACGAGGAU (<i>hcmv-mir-p45</i>)
-	36838	94	UCCUCUGCCUGGGCACGCGCUGCGCCGCGUCGCAAACGCUUGG <u>Guac</u> CCGAGGUCUUUUGCACGCGGACUUGGCCACCUUGCGUGCGGA (<i>hcmv-mir-p46</i>)
-	197467	90	GUGGGUGCCACGGACUUGGACCAUCACUCUGCAUUUGGUGC <u>cg</u> GCACCAAUUGCAAACCAUGUGGUGCCAGCCUCGGUACCAUUAU (<i>hcmv-mir-p47</i>)
+	119625	94	AUCCUCGUCGGUGCCUCUUCGGGGCGCAGCGGUGGACUCGGCCU <u>Uaag</u> GCGGCCGCGUGUCAUAACGCCCGACGUGCAGCCGUCGCCGAGGAU (<i>hcmv-mir-p48</i>)
+	147719	93	GACGGCGACGGUGAAAACAACGUCGUGGAAGUCAGCAGCAGCACC <u>ggc</u> GGGUGCGCACCCGCCGAGCGACGACGCCACUUUCACCGUGCAGUU (<i>hcmv-mir-p49</i>)
-	194965	93	UGACGUGACUCUUGACGUUUUAAACCGCAUGGGAAAGUACGGU <u>Gucgc</u> CACCGUUGACGUGGGCGGCGAUGAGAACGUCAGCGGUGGCCGAAA (<i>hcmv-mir-p50</i>)
+	128612	95	ACUGGGUCGUCUUACUGGGACCCGUGCCGUACCCUGUUUUU <u>Gcga</u> CGGUGAAGUGGAGGGCCACGGUGAACAUUCUGGUACCUACGACGCAGU (<i>hcmv-mir-p51</i>)

S (+/- strand), *SP* (start position), and *L* (length of the putative *pre-miRs*). 25 true-positives and 1 false-negative match 25 published *pre-miR* sequences (red regions) and their mature miRNAs (underlined regions) as obtained from miRBase 8.2 (Griffiths-Jones *et al.*, 2006); predicted terminal loop ≥ 3 -nts (bold lowercase nucleotides). [†]*kshv-mir-K12-9* are the accepted and incorrect positives of *kshv-mir-K12-9*. [‡]*ebv-mir-BHRF1-1* (0.437 *miPred* score) and [§]*mghv-mir-M1-8* (0.658).