2nd International Summit on Integrative Biology August 4-5, 2014 Chicago, USA

Pseudo DNA Sequence Generation of Non-coding Distributions using Stream Cipher Mechanism

Jeffrey Zheng School of Software, Yunnan University August 4, 2014

Content

- Frontier of Non-Coding DNAs/RNAs
- General Comparison Model for Pseudo DNAs & Real DNAs
- Sample Cases
- Conclusion



Frontier of Non–Coding DNAs/RNAs

Ratios on Non-Coding DNAs Tools for Analysis Current Situation Assumption & Question

Typical Ratios of Non-Coding DNAs/RNAs



3% U. Gibba



30% Arabidopsis



90% Takifugu



98% Human

 ENCODE: over 80% of DNA in the human genome "serves some purpose, biochemically speaking".
However, this conclusion is strongly criticized ...

Tools to Analyze Non-Coding DNAs/RNAs

- Frequency Distribution
- GC densities
- Repeat sub-sequences
- ...
- Machine Learning
- Bayesian Inference and Induction
- Neural Network
- Hidden Markov Model



A case of Non-Coding DNA: Hairpin



Refined Distributions on different parameters

Current Situation

- Total DNA varies widely between organisms
- Ratios of coding DNAs and Non-coding DNAs in genomes are different significantly
- > 98% human genomes are Non-coding DNAs
- Non-coding RNAs/DNAs may be drivers of complexity, they are a larger heterogeneous group

Due to various criteria, no a general classification can be used to subclassify this group

Assumption & Question

Assumption:

A general classification of Non-Coding DNA interactions could be relevant to higher levels of pair structures between a distance on a DNA sequence.

Both 0-1 outputs & DNA segments are random sequences

Question:

Can interaction models of Stream Cipher mechanism simulate a general classification for Non-Coding DNAs?

General Comparison Model for Pseudo DNAs & DNAs

Variant Logic DNAs & Pseudo DNAs General Model Main Procedure

Variant Logic

An unified 0-1 logic framework base on input/output and logic functions using four Meta symbols: {⊥, +, -, ⊤}

•
$$0-0$$
 : \perp , $0-1$: + ,

$$\circ$$
 1−0 : − , 1−1 : \top .

Multiple Maps of Variant Phase Spaces can be visualized



Variant Logic & DNA Sequencing



DNA Sequences	Variant Logic
G	0−0:⊥
A	0-1: +
Т	1–0: –
С	1–1: ⊤

Results of automated chaintermination DNA sequencing.

Four Meta States

A Comparison Model to simulate Non-Coding DNAs in Visual Maps

- Two input sources:
 - Pseudo DNAs Artificial Sequences using Stream Cipher on Interactions – HC256
 - Real DNAs Human DNAs
- Variant Construction to measure & quantity input sequences on 4 meta bases {ACGT}
- Using Visual Maps to identify higher levels of global symmetries between A&T and C&G maps for both artificial & real DNAs

General Comparison Model



Main Procedure

Input: Pseudo DNA/Real DNA Vector

X^t: GGTACTTGCAT...

Projected as Four 0-1 vectors

M_G: 11000001000 ...

M_A: 00010000010 ...

Calculated as four Probability Vectors

 $\left\{ \rho_l^V \right\}_{0 \leq l < m_t}$

Determine four pairs of map position





 $\sqrt[k]{\sum_{i=1}^{n} pi^k}$

 $\left(\sum_{i=1}^n \sqrt[k]{pi}\right)^k$

Collected all DNA Vectors $\forall t, X^{t} \in D^{N_{t}}$

Four Maps constructed

{Map_v}_{veD}



Sample Cases

2700 DNA Sequences Human DNAs vs. HC256 Pseudo DNAs Sets of Maps

Non-Coding DNA Sequence Information

Two Sets of T=2700 sequences

Non-Coding DNAs for Human Genomes

• SRR027956.xxxxxxx , N= 500bp

For a sample point, a sequence could be

Human DNAs vs. Pseudo DNAs



(a1-a5) Map_A for the file *Right*; (b1-b5) Map_A for the file hc256 mode = 1, r = 1.



Pseudo DNAs on various conditions



Figure 5. Four groups of sixteen 2D maps in the range of n = 15, $k = \{2,3,4,7\}$, $N \cong 500$, T = 2700; (a) group (a1 - a4) four Map_A maps; (b) group (b1-b4) four Map_T maps; (c) (c1 - c4) four Map_G maps; (d) (d1 - d4) four Map_C maps for the file *right*.

Pseudo DNA sequences on different parameters



Figure 6. Four groups of sixteen 2D maps in the range of n = 12, $k = \{2,3,4,7\}$, $N \cong 500$, T = 2700 for the file hc256, r = 1, mode = 1; (a) group (a1 - a4) four Map_A maps; (b) group (b1-b4) four Map_T maps; (c) (c1 - c4) four Map_G maps; (d) (d1 - d4) four Map_C maps.

Two Groups of Human DNAs



Figure 7. Two groups of eight 2D maps in the range of n = 15, k = 7, $N \cong 200 \sim 600$, T = 2700; (a) group (a1 - a4) four Map_V maps for the file *left*; (b) group (b1-b4) four Map_V maps for the file *right*.

Pseudo DNAs under Various Interactions



Human DNAs vs. Pseudo DNAs



RTP21

right; (b-c) on

Conclusion

Conclusion

- Using Variant Logic, Four DNA Meta States correspond Four Variant Meta States
- Pseudo DNAs can be generated under Various conditions to form Visual Maps
- Both Real & Artificial DNAs have stronger similarity
- Visual Maps may provide a General Classification for Genomic analysis on DNA Interactions
- Further Explorations are required...

References

- B. Banfai, H. Jia, J. Khatun et al. (2012) Long noncoding RNAs are rarely translated in two human cell lines, *Genome Research*, Cold Spring Harbor Laboratory Press, 22:1646-1657 Doi:10.1101/gr.134767.111
- 2. J.M. Engreitz, A. Pandya-Jones, P. McDonel et al. (2013) Large Noncoding RNAs can Localize to Regulatory DNA Targets by Exploriting the 3D Architecture of the Genome, *Proceedings of The Biology of Genomes*, Cold Spring Harbor Laboratory Press, 122
- 3. J. Zheng, C. Zheng and T. Kunii (2011) A Framework of Variant Logic Construction for Cellular Automata, in *Cellular Automata – Innovative Modelling for Science and Engineering*, Edited by A. Salcido, InTech Press, 325-352, 2011. <u>http://www.intechopen.com/chapters/20706</u>
- 4. J. Zheng, W. Zhang, J. Luo, W. Zhou and R. Shen (2013) Variant Map System to Simulate Complex Properties of DNA Interactions Using Binary Sequences, *Advances in Pure Mathematics*, 3 (7A) 5-24. doi: <u>10.4236/apm.2013.37A002</u>
- 5. J. Zheng, J. Luo and W. Zhou (2014) Pseudo DNA Sequence Generation of Non-Coding Distributions Using Variant Maps on Cellular Automata," *Applied Mathematics*, 5(1) 153-174. doi: <u>10.4236/am.2014.51018</u>
- 6. J. Zheng, W. Zhang, J. Luo, W. Zhou, V. Liesaputra (2014) Variant Map Construction to Detect Symmetric Properties of Genomes on 2D Distributions. *J Data Mining Genomics Proteomics* 5:150. doi: 10.4172/2153-0602.1000150

