

Additional file 1

1 Additional Figures

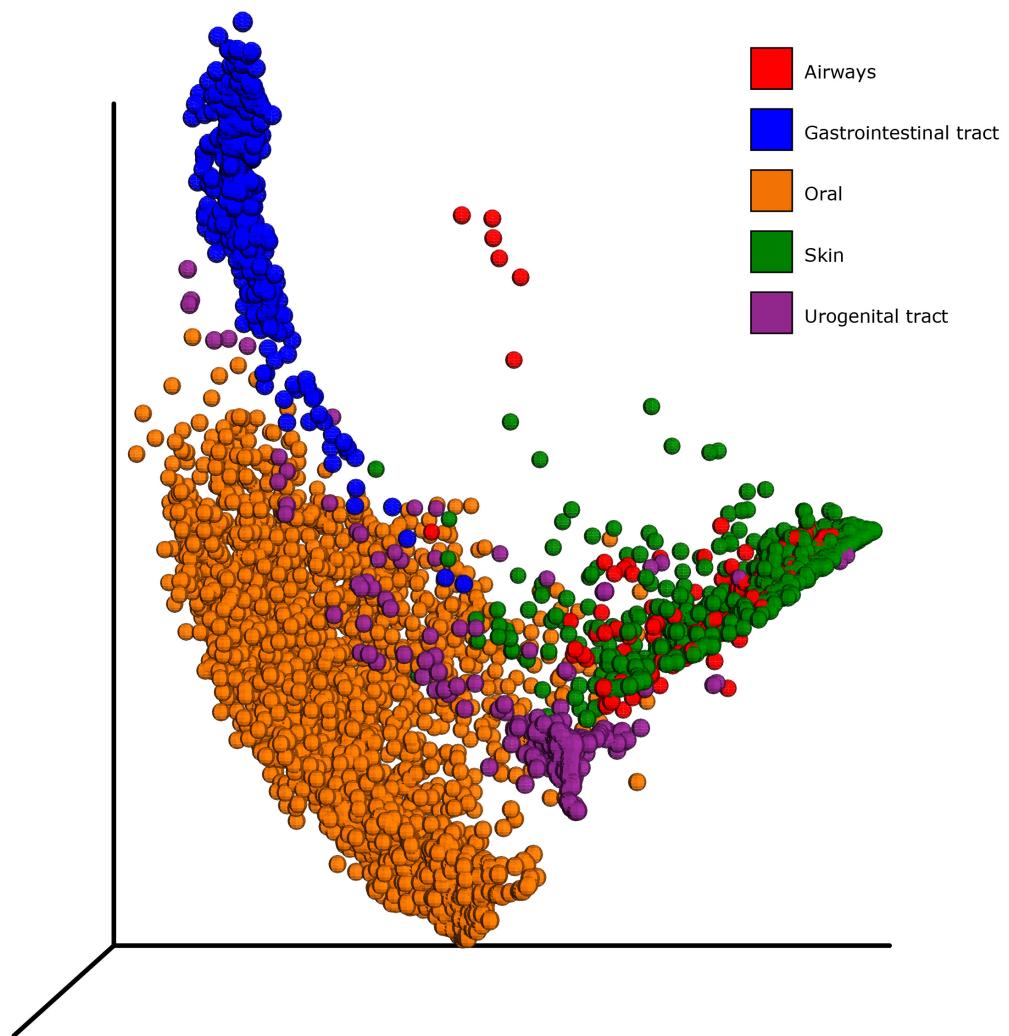


Figure S1: The PCoA plot of The Human Microbiome Project Consortium (2012) dataset, which is generated via the *beta_diversity_through_plots.py* script available by QIIME.

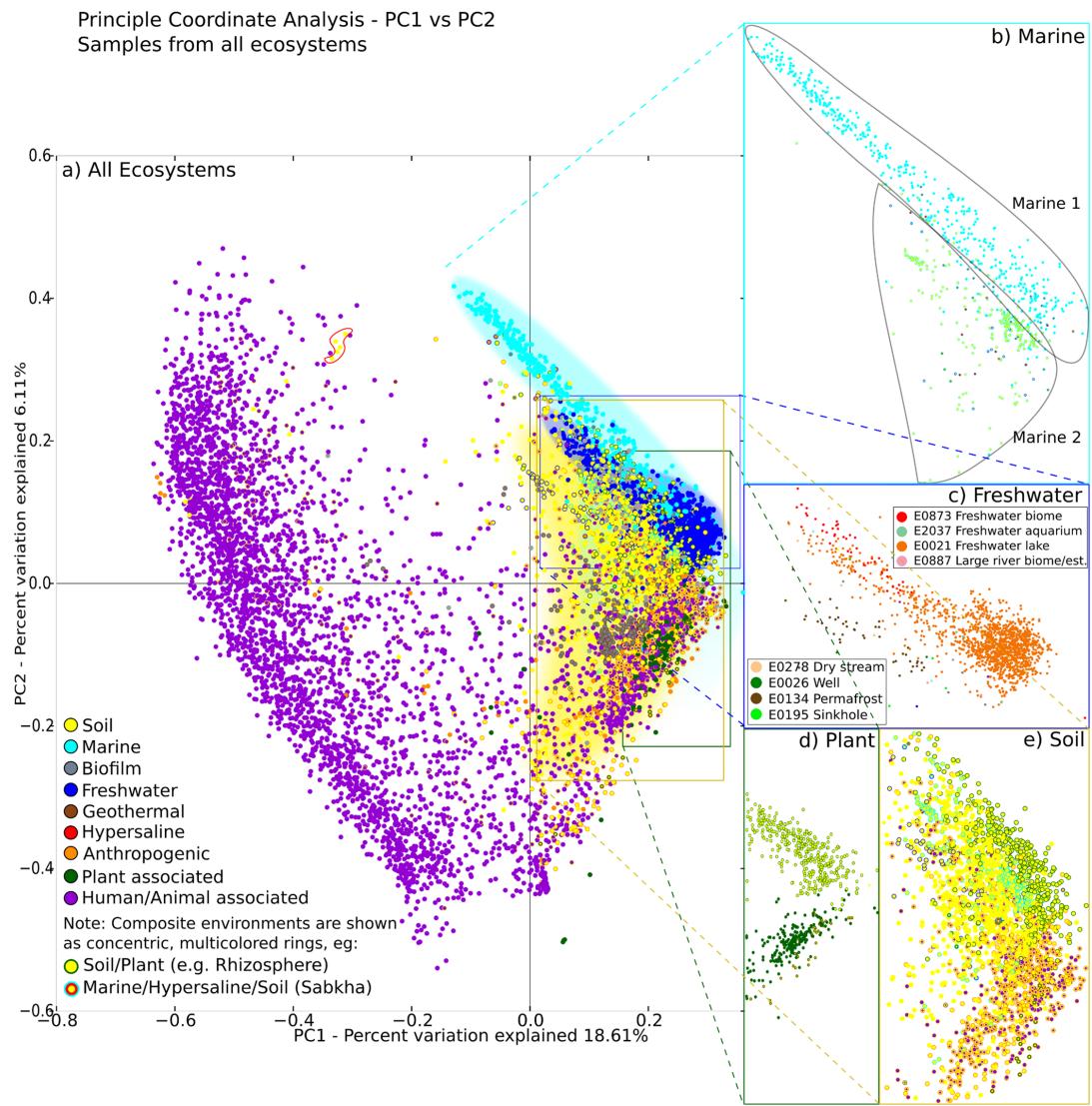


Figure S2: The PCoA plot provided in the Meta-analysis of environmental microbiomes conducted by Henschel *et al.* (2015).

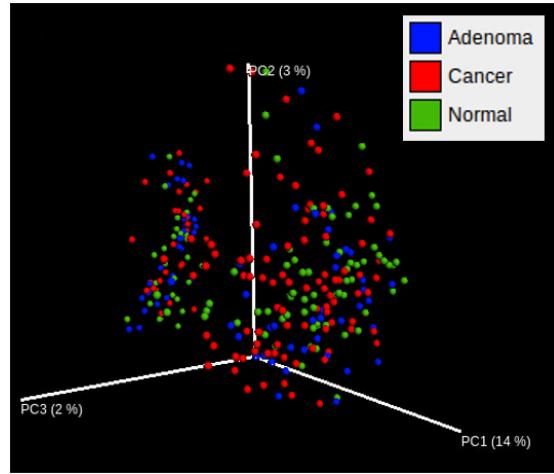


Figure S3: The PCoA plot of the combined CRC dataset.

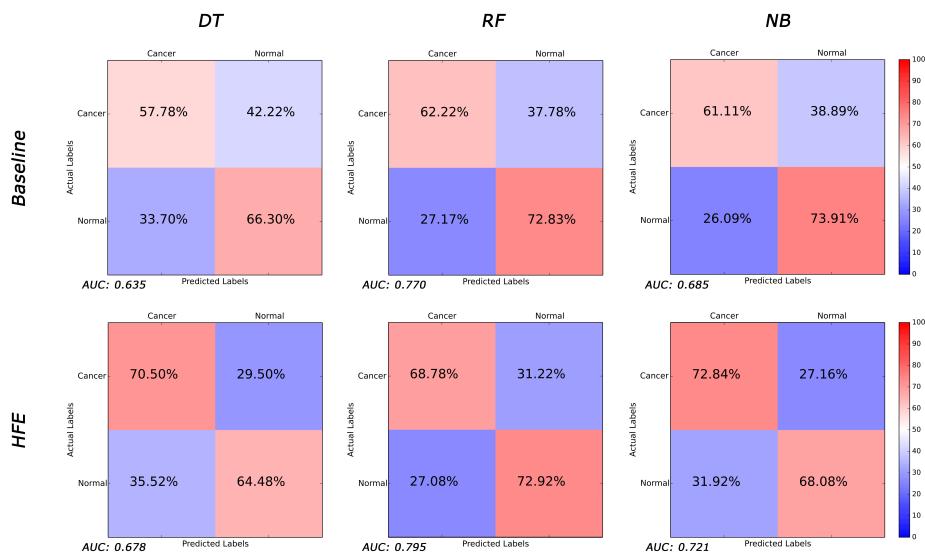


Figure S4: Comparison between the baseline and HFE confusion matrices when applied on CRC1 dataset (Zeller *et al.*, 2014) for Cancer vs. Normal classification.

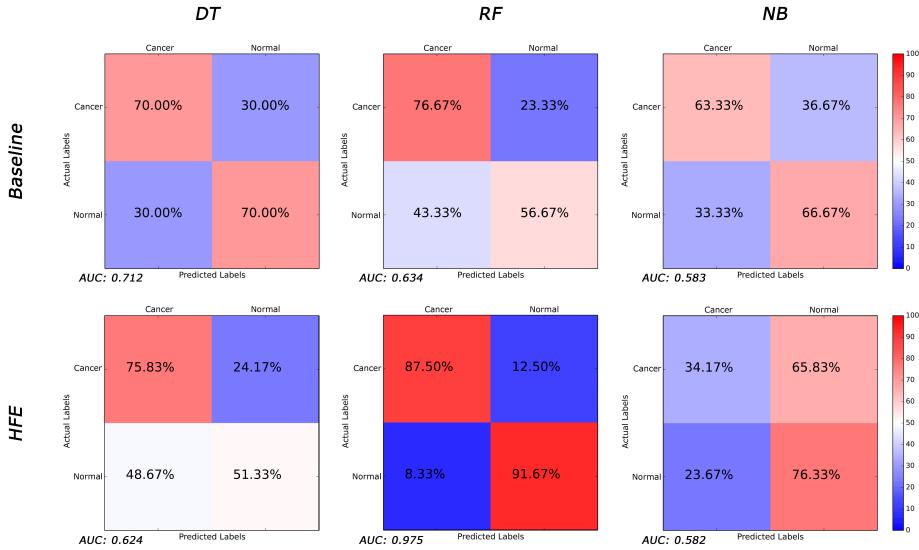


Figure S5: Comparison between the baseline and HFE confusion matrices when applied on CRC2 dataset (Zackular *et al.*, 2014) for Cancer vs. Normal classification.

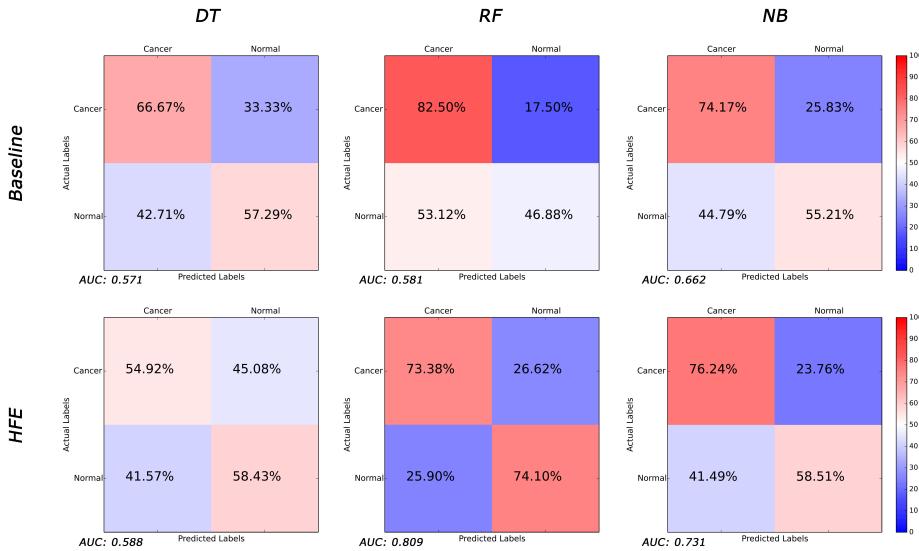


Figure S6: Comparison between the baseline and HFE confusion matrices when applied on CRC1+2 dataset for Cancer vs. Normal classification.

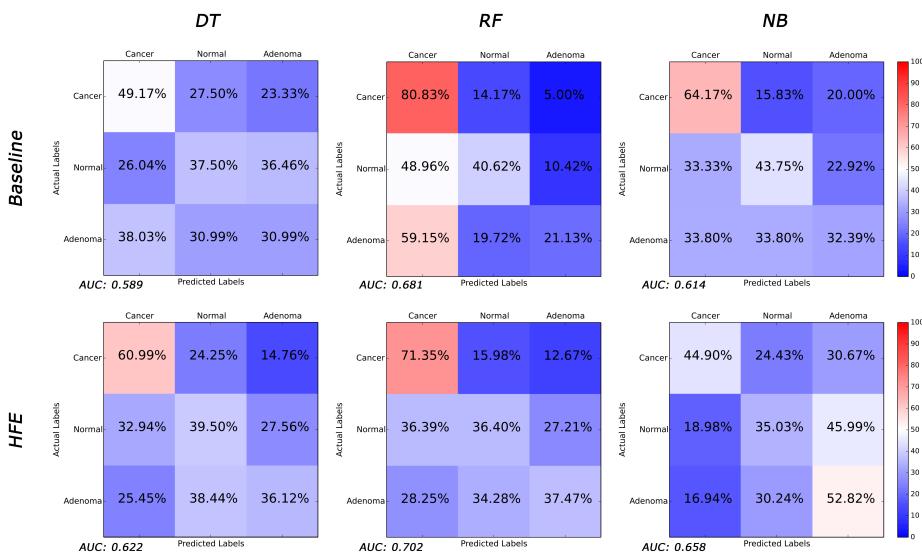


Figure S7: Comparison between the baseline and HFE confusion matrices when applied on CRC1+2 dataset for Cancer vs. Normal vs. Adenoma classification.

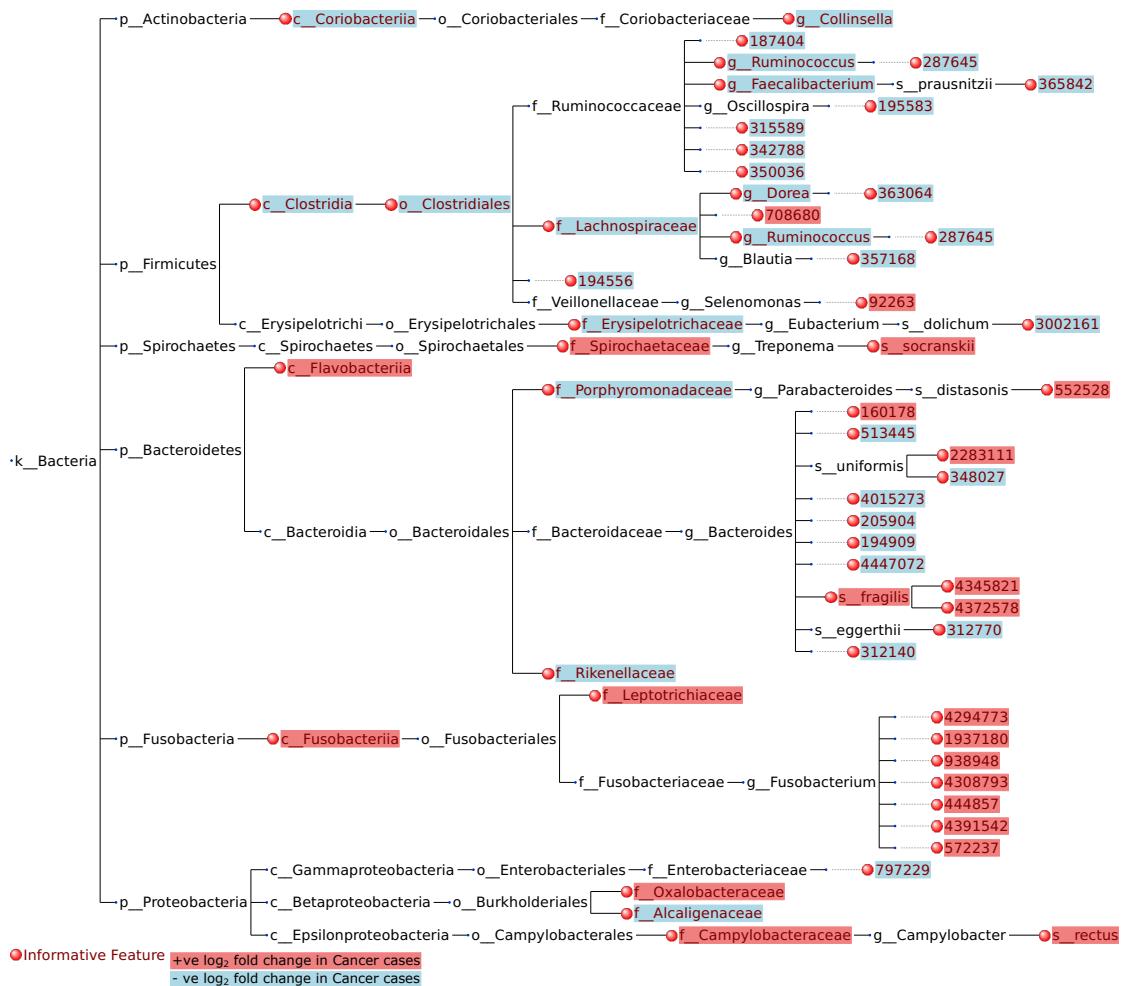


Figure S8: The taxonomic tree of all the informative features extracted by the HFE method for Cancer vs. Normal classification with respect to the dataset provided by Kostic *et al.* (2012).

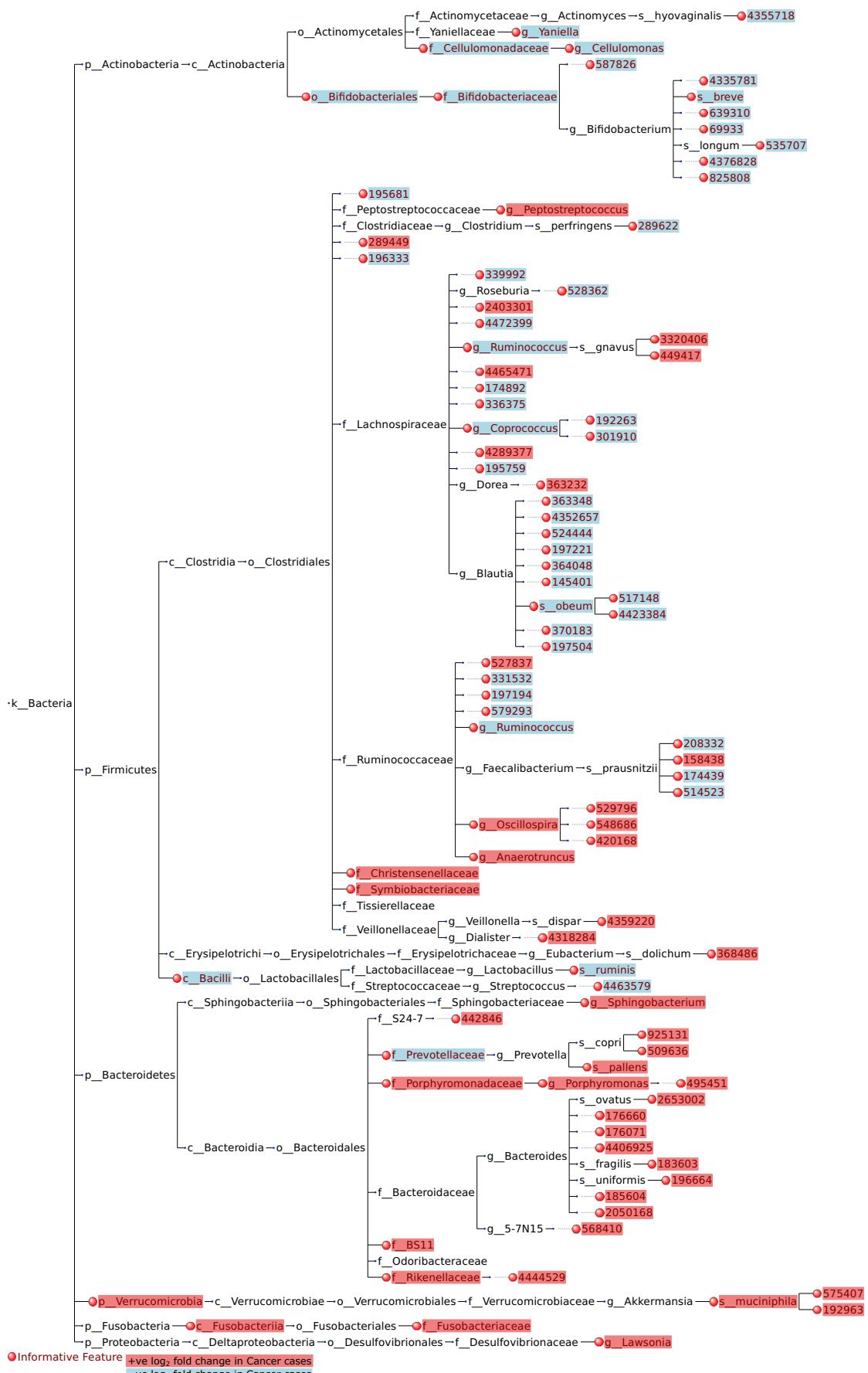


Figure S9: The taxonomic tree of all the informative features extracted by the HFE method for Cancer vs. Normal classification with respect to CRC1 dataset (Zeller *et al.*, 2014).

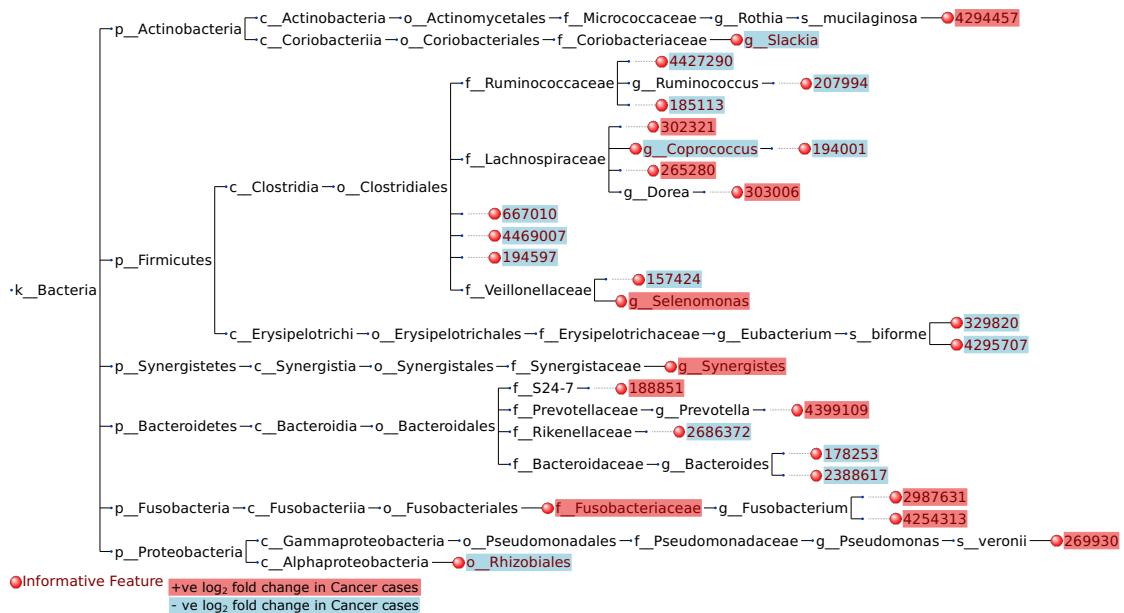


Figure S10: The taxonomic tree of all the informative features extracted by the HFE method for Cancer vs. Normal classification with respect to CRC2 dataset (Zackular *et al.*, 2014).

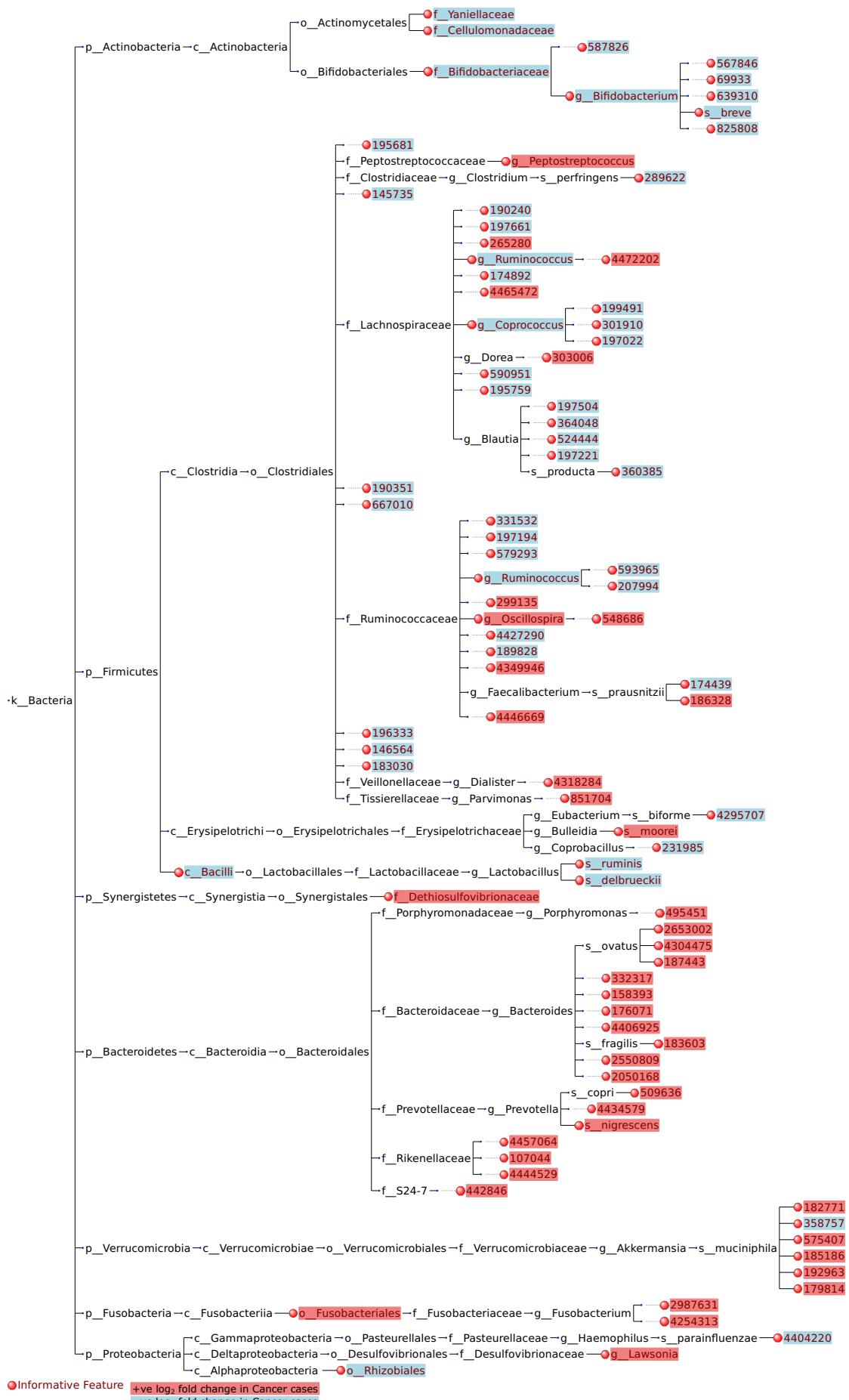


Figure S11: The taxonomic tree of all the informative features extracted by the HFE method for Cancer vs. Normal classification with respect to CRC1+2 dataset.

2 Supplementary Tables

Table **S1**: The cross-validation results of the proposed pipeline when applied for human body site prediction and environment prediction, in terms of AUC.

ML	Human Body Site Prediction		Environment Prediction	
	BL	HFE	BL	HFE
DT	0.992	0.985	0.973	0.960
RF	0.999	0.999	0.999	0.999
NB	0.994	0.995	0.903	0.949
#Features	5,430	84	30,860	267

BL and HFE refer to the baseline and HFE feature sets, respectively.

References

- Henschel, A., Anwar, M., and Manohar, V. (2015). Comprehensive meta-analysis of ontology annotated 16s rrna profiles identifies beta diversity clusters of environmental bacterial communities. *PLoS Computational Biology*, **11**, 1–24.
- Kostic, A. D., Gevers, D., Pedamallu, C. S., Michaud, M., Duke, F., Earl, A. M., Ojesina, A. I., Jung, J., Bass, A. J., Tabernero, J., et al. (2012). Genomic analysis identifies association of fusobacterium with colorectal carcinoma. *Genome research*, **22**(2), 292–298.
- The Human Microbiome Project Consortium (2012). Structure, function and diversity of the healthy human microbiome. *Nature*, **486**, 207–214.
- Zackular, J., Rogers, M., Ruffin, M., and Schloss, P. (2014). The human gut microbiome as a screening tool for colorectal cancer. *Cancer Prevention Research*, **7**, 1112–1121.
- Zeller, G., Tap, J., Voigt, A., Sunagawa, S., Kultima, J., Costea, P., Amiot, A., Bohm, J., Brunetti, F., Habermann, N., Hercog, R., Koch, M., Luciani, A., et al. (2014). Potential of fecal microbiota for early-stage detection of colorectal cancer. *Molecular Systems Biology*, **10**, 1–18.