| 1 | - Correlation coefficients are widely used to    | 1 | + This paper presents the Clustermatch         |
|---|--|---|--|
|   | identify patterns in data that may be of         |   | Correlation Coefficient (CCC), an efficient    |
|   | particular interest.                             |   | and not-only-linear correlation coefficient    |
|   |  |   | based on machine learning models, to identify  |
|   |  |   | linear and nonlinear patterns in               |
|   |  |   | transcriptomics data.                          |
| 2 | - In transcriptomics, genes with correlated      | 2 | + We aim to determine if CCC can detect        |
|   | expression often share functions or are part     |   | meaningful linear and nonlinear relationships  |
|   | of disease-relevant biological processes.        |   | in gene expression data, including those       |
|   |  |   | missed by linear-only correlation              |
|   |  |   | coefficients, and if highly-ranked gene pairs  |
|   |  |   | by CCC are enriched for interactions in        |
|   |  |   | integrated networks.                           |
| 3 | - Here we introduce the Clustermatch Correlation | 3 | + When applied to human gene expression data,  |
|   | Coefficient (CCC), an efficient, easy-to-use     |   | CCC identifies robust linear relationships and |
|   | and not-only-linear coefficient based on         |   | nonlinear patterns associated with sex         |
|   | machine learning models.                         |   | differences.                                   |
| 4 | - CCC reveals biologically meaningful linear and | 4 | + Our results suggest that CCC can detect      |
|   | nonlinear patterns missed by standard, linear-   |   | functional relationships not captured by       |
|   | only correlation coefficients.                   |   | linear-only methods.                           |
| 5 | - CCC captures general patterns in data by       | 5 | + CCC is a highly-efficient, next-generation   |
|   | comparing clustering solutions while being       |   | not-only-linear correlation coefficient that   |
|   | much faster than state-of-the-art coefficients   |   | can be applied to genome-scale data and other  |
|   | such as the Maximal Information Coefficient.     |   | domains across different data types.           |
| 6 | - When applied to human gene expression data,    |   |  |
|   | CCC identifies robust linear relationships       |   |  |
|   | while detecting nonlinear patterns associated,   |   |  |
|   | for example, with sex differences that are not   |   |  |
|   | captured by linear-only coefficients.            |   |  |
| 7 | - Gene pairs highly ranked by CCC were enriched  |   |  |
|   | for interactions in integrated networks built    |   |  |
|   | from protein-protein interaction,                |   |  |
|   | transcription factor regulation, and chemical    |   |  |
|   | and genetic perturbations, suggesting that CCC   |   |  |
|   | could detect functional relationships that       |   |  |
|   | linear-only methods missed.                      |   |  |
| 8 | - CCC is a highly-efficient, next-generation     |   |  |
|   | not-only-linear correlation coefficient that     |   |  |
|   | can readily be applied to genome-scale data      |   |  |
|   | and other domains across different data types.   |   |  |