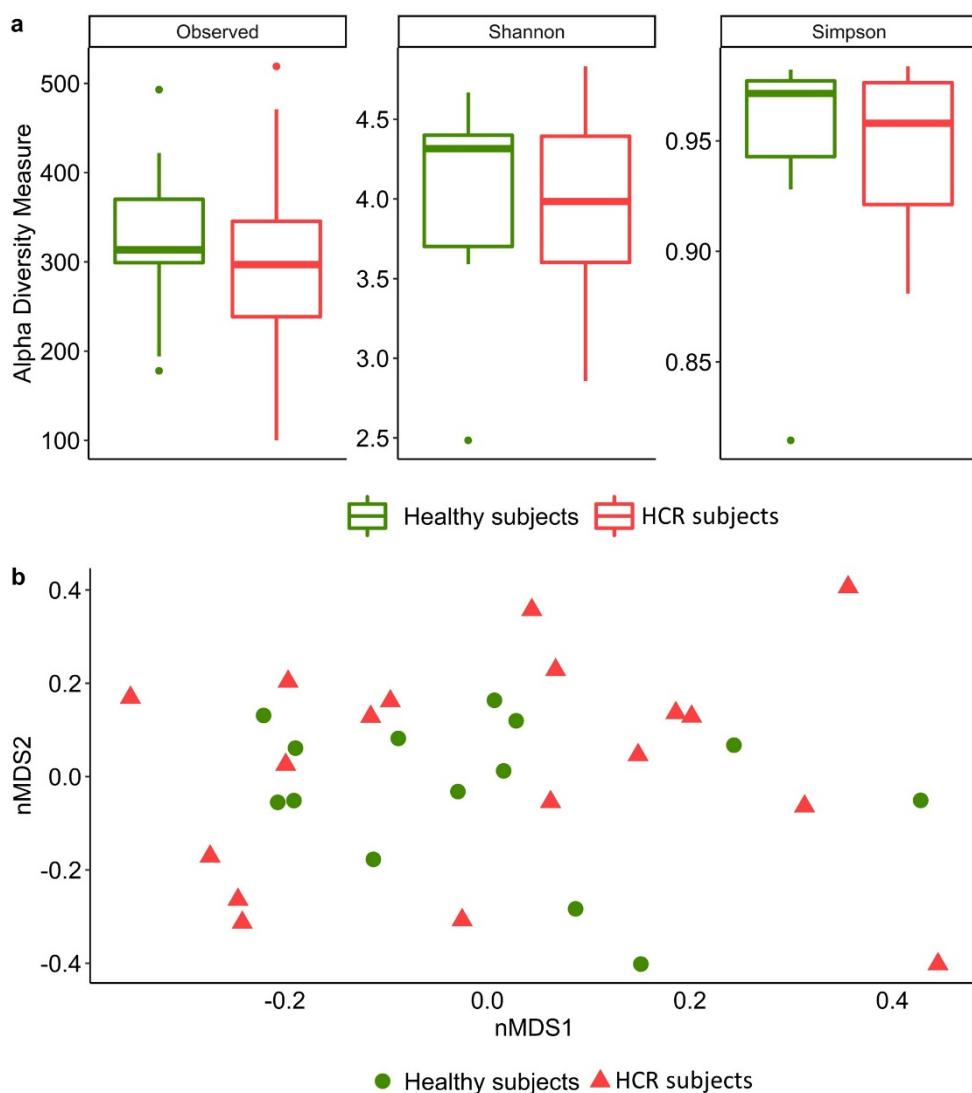
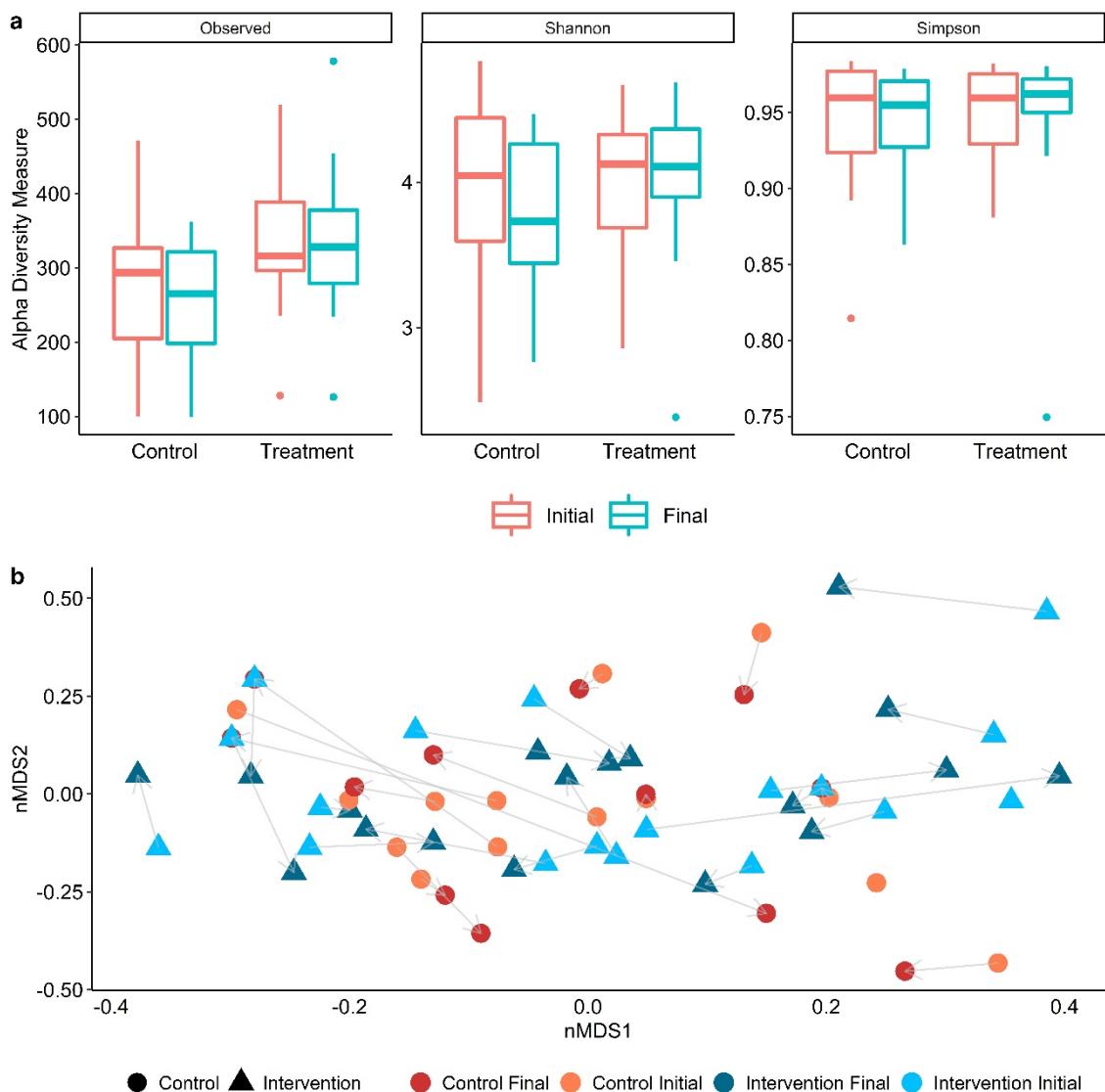


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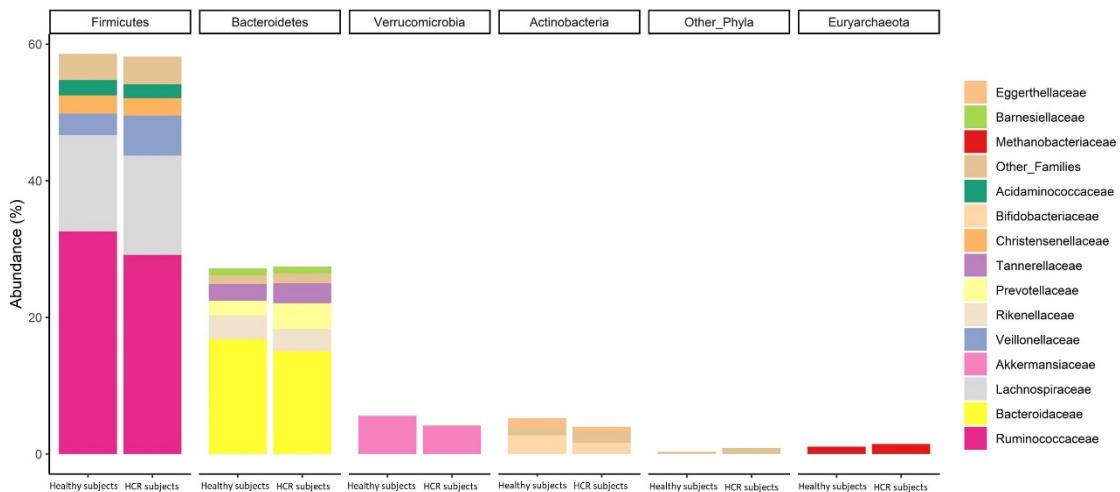
Supporting Information



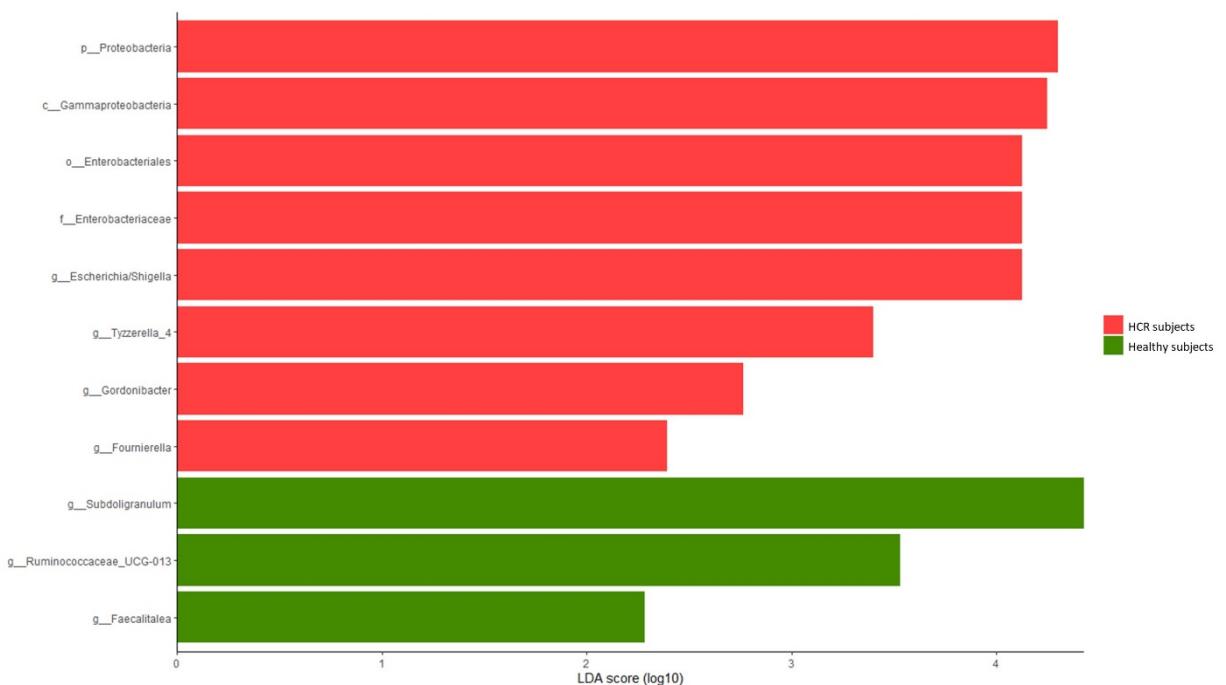
**Figure S1.** Bacterial diversity analysis of faecal samples before the intervention period. a) Shannon ( $H'$ ) diversity index as a measurement of alpha-diversity. Pairwise group comparisons confirmed no significant differences (Wilcoxon rank sum test) between healthy subjects and HCR subjects. b) Beta-diversity (nMDS) of healthy subjects and HCR subjects.



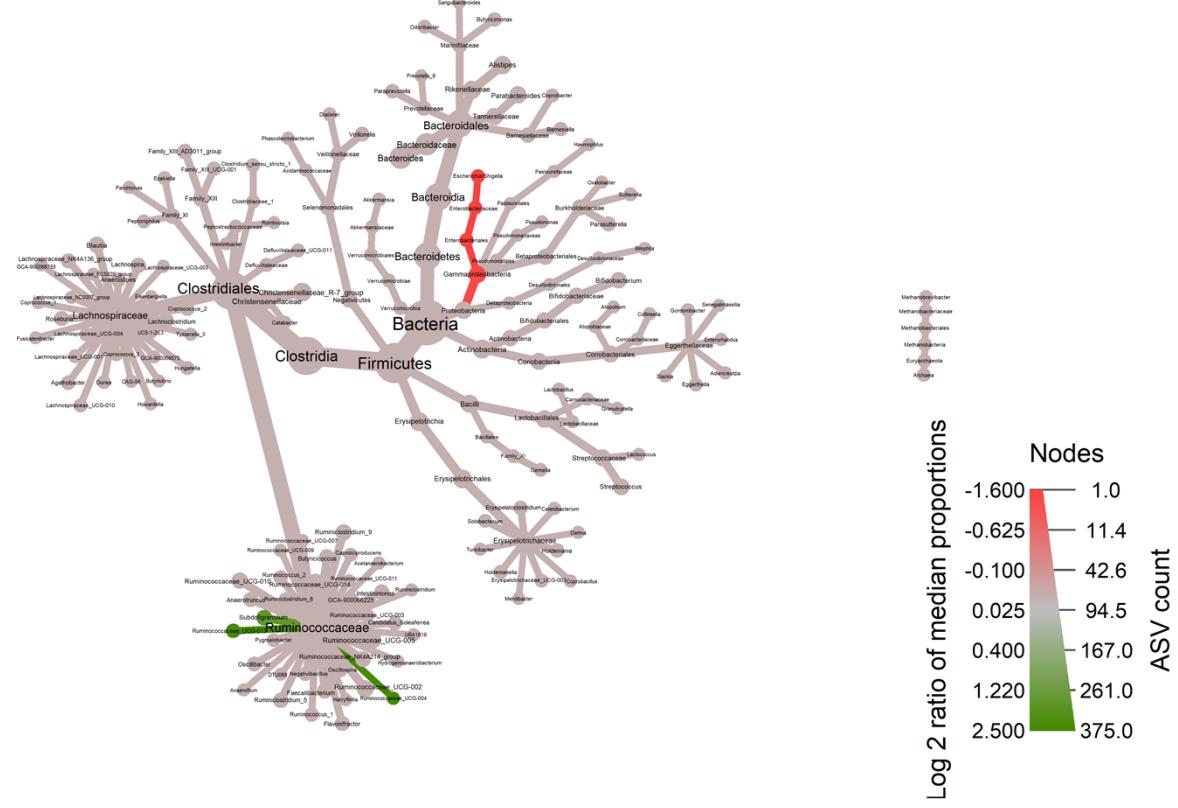
**Figure S2.** Bacterial diversity analysis of faecal samples before and after the intervention period. a) Shannon ( $H'$ ) diversity index as a measurement of alpha-diversity. Pairwise group comparisons confirmed no significant differences (Wilcoxon signed-rank test) between groups and between sampling times (before and after wine consumption) at an intra-group level. b) Beta-diversity (nMDS) of the control and intervention groups before and after the intervention. Lines join data from the same individual.



**Figure S3.** Relative abundance of the main bacterial phyla and families present in faecal samples and the comparison between healthy subjects and HCR subjects.



**Figure S4.** LDA scores (log10) for the significant biomarkers obtained from the LEfSe analysis between the groups of healthy subjects and HCR subjects.



**Figure S5.** Heat tree showing differential abundance between healthy subjects (blue) and HCR subjects (red) according to a Wilcoxon rank sum test carried out with faecal samples at baseline and only with that ASVs present in at least 10 individuals.

**Table S1.** Amplicon sequence variants (ASV) correlated to clinical parameters and/or microbial-derived metabolites in the four communities obtained after bacterial co-occurrence networking.

| ASV                             | correlated to                    | Community                        | Keystoness |
|---------------------------------|----------------------------------|----------------------------------|------------|
| G_Bacteroides                   | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Bacteroides                   | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Bacteroides                   | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Erysipelatoclostridium        | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| F_Lachnospiraceae               | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Marvinbryantia                | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Ruminococcaceae UCG-010       | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Subdoligranulum               | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.86245124 |
| G_Bacteroides                   | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| G_Christensenellaceae R-7 group | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| S_Porphyromonas uenonis         | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| G_Ruminiclostridium             | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| F_Ruminococcaceae               | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| F_Ruminococcaceae               | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.83489266 |
| S_Bacteroides fragilis          | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.80761591 |
| G_Coprobacter                   | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.80238715 |
| G_Rikenella                     | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.80132353 |
| G_Ruminococcaceae NK4A214 group | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.80104992 |
| F_Christensenellaceae           | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.80049188 |
| G_Ruminococcaceae NK4A214 group | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.79809087 |
| O_Clostridiales                 | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.79412456 |
| G_Prevotella                    | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.78183566 |
| G_Ruminiclostridium 5           | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.76771115 |
| F_Ruminococcaceae               | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.7631378  |

|                                 |                                  |                                  |            |
|---------------------------------|----------------------------------|----------------------------------|------------|
| G_Eubacterium                   | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.76062861 |
| G_Ruminococcaceae UCG-005       | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.74795148 |
| G_Methanobrevibacter            | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.72353679 |
| G_Intestinimonas                | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.71722376 |
| O_DTU014                        | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.71509562 |
| G_Coprobacter                   | (+)BP(-)Butyric acid(-)Phenolics | (+)BP(-)Butyric acid(-)Phenolics | 0.71443475 |
| S_Actinomyces odontolyticus     | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.70202522 |
| G_Alistipes                     | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.70202522 |
| G_DTU089                        | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.70202522 |
| F_Saccharimonadaceae            | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.70202522 |
| F_Saccharimonadaceae            | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.70202522 |
| S_Gordonibacter pamelaeae       | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.70093747 |
| G_Enterococcus                  | (+)BP(-)Phenolics                | (+)BP(-)Butyric acid(-)Phenolics | 0.69949413 |
| G_Methanobrevibacter            | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.68582049 |
| G_Olsenella                     | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.6786717  |
| G_Ruminococcaceae NK4A214 group | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.6776808  |
| G_Christensenellaceae R-7 group | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.6710752  |
| F_Christensenellaceae           | (-)Butyric acid(-)Phenolics      | (+)BP(-)Butyric acid(-)Phenolics | 0.67083745 |
| S_Fusobacterium nucleatum       | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.65717787 |
| F_Eggerthellaceae               | (-)Butyric acid                  | (+)BP(-)Butyric acid(-)Phenolics | 0.64592985 |
| F_Eggerthellaceae               | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| F_Erysipelotrichaceae           | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| G_Faecalitalea                  | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| G_Butyricimonas                 | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| F_Muribaculaceae                | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| F_Muribaculaceae                | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| F_Muribaculaceae                | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |
| F_Muribaculaceae                | (-)Phenolics                     | (-)Phenolics                     | 0.75806908 |

|                                |                          |                          |            |
|--------------------------------|--------------------------|--------------------------|------------|
| G_Ruminococcaceae UCG-014      | (-)Phenolics             | (-)Phenolics             | 0.75806908 |
| F_Ruminococcaceae              | (-)Phenolics             | (-)Phenolics             | 0.75806908 |
| G_Ruminococcaceae UCG-005      | (-)Phenolics             | (-)Phenolics             | 0.75806908 |
| G_Subdoligranulum              | (-)Phenolics             | (-)Phenolics             | 0.75806908 |
| G_Dialister                    | (-)Phenolics             | (-)Phenolics             | 0.75806908 |
| G_Bacteroides                  | (-)Phenolics             | (-)Phenolics             | 0.71085076 |
| F_Ruminococcaceae              | (-)Phenolics             | (-)Phenolics             | 0.7041538  |
| G_Angelakisella                | (-)Phenolics             | (-)Phenolics             | 0.65861735 |
| G_Faecalibacterium             | (-)Phenolics             | (-)Phenolics             | 0.65420968 |
| G_Bacteroides                  | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| S_Barnesiella intestinihominis | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| S_Parasutterella secunda       | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Holdemanella                 | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Holdemanella                 | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Holdemanella                 | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| F_Prevotellaceae               | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Prevotella 9                 | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Prevotella 9                 | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| F_Prevotellaceae               | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Ruminococcaceae UCG-004      | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Ruminococcaceae UCG-005      | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Negativibacillus             | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Subdoligranulum              | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Subdoligranulum              | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Parabacteroides              | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Allisonella                  | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Dialister                    | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.86818654 |
| G_Parabacteroides              | (+)Cholesterol(+)Glucose | (+)Cholesterol(+)Glucose | 0.82935041 |

|                                 |                                     |                                     |            |
|---------------------------------|-------------------------------------|-------------------------------------|------------|
| G_Parabacteroides               | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.81883202 |
| F_Ruminococcaceae               | (+)Cholesterol(+)Glucose            | (+)Cholesterol(+)Glucose            | 0.78778786 |
| G_Lachnospiraceae UCG-010       | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.77699938 |
| G_Negativibacillus              | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.77699938 |
| G_Christensenellaceae R-7 group | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.76614238 |
| G_Parasutterella                | (+)Cholesterol(+)Glucose            | (+)Cholesterol(+)Glucose            | 0.76096304 |
| G_Anaerofilum                   | (+)Cholesterol(+)Glucose            | (+)Cholesterol(+)Glucose            | 0.76096304 |
| F_Ruminococcaceae               | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.74454675 |
| G_Catenibacterium               | (+)Glucose                          | (+)Cholesterol(+)Glucose            | 0.72582345 |
| F_Ruminococcaceae               | (+)Cholesterol(+)Glucose            | (+)Cholesterol(+)Glucose            | 0.7036379  |
| G_Ruminococcaceae UCG-014       | (+)Cholesterol                      | (+)Cholesterol(+)Glucose            | 0.70334566 |
| S_Faecalicoccus pleomorphus     | (+)Cholesterol(+)Glucose            | (+)Cholesterol(+)Glucose            | 0.67999734 |
| F_Lachnospiraceae               | (+)Cholesterol                      | (+)Cholesterol(+)Glucose            | 0.67318677 |
| G_Family XIII AD3011 group      | (+)Cholesterol                      | (+)Cholesterol(+)Glucose            | 0.66847504 |
| S_Flavonifractor plautii        | (+)Cholesterol                      | (+)Cholesterol(+)Glucose            | 0.66538279 |
| G_Paraprevotella                | (+)Cholesterol                      | (+)Cholesterol(+)Glucose            | 0.60656758 |
| G_Phascalarctobacterium         | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Phascalarctobacterium         | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Phascalarctobacterium         | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Bacteroides                   | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Bacteroides                   | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Bacteroides                   | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Butyricimonas                 | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Butyricimonas                 | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Paraprevotella                | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Paraprevotella                | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Alistipes                     | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Alistipes                     | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |

|                                 |                                     |                                     |            |
|---------------------------------|-------------------------------------|-------------------------------------|------------|
| G_Ruminococcaceae UCG-014       | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Ruminococcaceae UCG-014       | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| F_Ruminococcaceae               | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Ruminococcaceae NK4A214 group | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Ruminococcaceae UCG-005       | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Ruminococcaceae_UCG-010       | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| F_Ruminococcaceae               | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Parabacteroides               | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Parabacteroides               | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.76998151 |
| G_Ruminococcaceae UCG-005       | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.7317685  |
| G_Alistipes                     | (-)Weight                           | (-)Creatinine(-)Leukocytes(-)Weight | 0.72581931 |
| G_Ruminococcaceae UCG-014       | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.68315734 |
| S_Phascolarctobacterium         | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.68261174 |
| succinatutens                   | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.68261174 |
| G_Bacteroides                   | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.67483768 |
| G_Ruminococcus 1                | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.67188655 |
| G_Parabacteroides               | (-)Creatinine                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.67073368 |
| G_Barnesiella                   | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.66497361 |
| G_Butyricimonas                 | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.66209737 |
| G_Bacteroides                   | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.65917934 |
| G_Pygmaiovibrio                 | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.65763952 |
| S_Prevotella timonensis         | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.65760176 |
| G_Paraprevotella                | (-)Creatinine(-)Leukocytes(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.65591882 |
| G_Butyricimonas                 | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight | 0.64979569 |
| G_Barnesiella                   | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.64812211 |
| G_Porphyromonas                 | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.64794966 |
| G_Defluviitaleaceae UCG-011     | (-)Creatinine(-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.64757782 |
| S_Prevotella 6 corporis         | (-)Leukocytes                       | (-)Creatinine(-)Leukocytes(-)Weight |            |

|                   |                        |                                     |            |
|-------------------|------------------------|-------------------------------------|------------|
| G_Porphyrimonas   | (-)Weight              | (-)Creatinine(-)Leukocytes(-)Weight | 0.64205069 |
| C_Bacteroidia     | (-)Leukocytes          | (-)Creatinine(-)Leukocytes(-)Weight | 0.6234686  |
| F_Ruminococcaceae | (-)Creatinine(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.62257628 |
| F_Ruminococcaceae | (-)Creatinine(-)Weight | (-)Creatinine(-)Leukocytes(-)Weight | 0.56587745 |