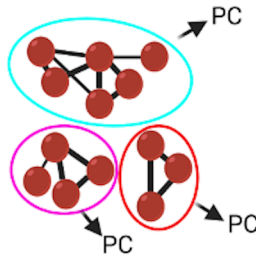


Analysis pipeline



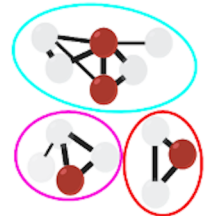
1019 metabolites
following quality
control

1) Group metabolites into modules of highly coexpressed metabolites using weighted gene coexpression network analysis (WGCNA). Derive the first principal component (PC) of each module to be used for association analyses



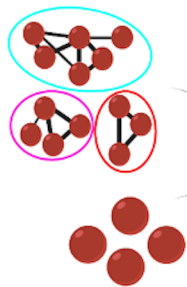
WGCNA modules

2) Identify metabolites highly correlated to the module that are likely to be driving module expression



Hub metabolites

Are modules and hub metabolites associated with imaging measures? $N=437$



Amyloid status
Mean hippocampal volume
Whole-brain volume

Are key modules and hub metabolites additionally associated with polygenic risk scores (PRS) for Alzheimer's disease? $N=1638$

APOE⁺ PRS



APOE⁻ PRS

Alzheimer's disease PRS

