

Data: Correct GO predictions, T , for the selected set of genes
Data: Key list representing parent classes for every GO class
parameter: Vector of L Noise Levels, NL
default: $NL = [0, 0.1, 0.2...1]$
parameter: Number of repetitions, K , within each NL
default: $K = 1000$
Result: K by L Matrix, scores with selected Evaluation Metric
begin
 Define L as the length of NL ;
 Define $output$ as K by L matrix ;
 Set $N_{neg} = 4$ (GO classes per gene in neg. set) ;
 foreach l in $[1, 2, \dots, L]$ **do**
 $p = NL[l]$, current noise level ;

 #1. Create positive and negative data
 $P_{pos} = GeneratePosData(T, p)$;
 $P_{neg} = GenerateNegData(T, N_{neg})$;

 #2. Create artificial classifier scores to sets
 foreach row in P_{pos} **do**
 Select predictor score, s , from $Normal(\mu = 1, \sigma = 0.5)$;
 Add s to the current row;
 end
 foreach row in P_{neg} **do**
 Select predictor score, s , from $Normal(\mu = -1, \sigma = 0.5)$;
 Add s to the current row;
 end

 # 3. Combine positive and negative datasets
 $P = P_{pos} \cup P_{neg}$;

 # 4. Run Evaluation Metric, EvM
 $Output[k, l] = EvM(P, T)$;
 end
end

Algorithm 1: Artificial Dilution Series (ADS) pipeline. Code uses sub-functions *GeneratePosData* and *GenerateNegData*, explained later.

Data: Correct GO predictions, T , for the selected set of genes

Data: Key list representing parent classes for every GO class

parameter: Noise proportion, p

output : Modified GO predictions, P

begin

Define N_T as the size of T ;

$P = T$;

$th_{noise} = round(N_T * p)$, the size threshold for the Noise Set ;

Shifting step

Define N_{shift} , a random integer between 0 and N_T ;

Select P_{shift} , a random subset of P of size N_{shift} ;

foreach *row* in P_{shift} **do**

 | Replace GO class with one of its nearest parents;

end

Permutation step

Define $NoiseSet = []$;

Define $N_{err} = 0$;

while $th_{noise} > N_{err}$ **do**

 | Select two random rows A and B from $P \setminus NoiseSet$;

 | # Genes of these rows will be $gene_A$ and $gene_B$;

 | # GO classes of these rows will be GO_A and GO_B ;

 | **if** $IsNoiseClass(gene_A, GO_B)$ and $IsNoiseClass(gene_B, GO_A)$ **then**

 | Swap GO classes between rows A and B in P ;

 | Add A and B to $NoiseSet$;

 | $N_{err} = N_{err} + 2$;

 | **end**

end

return P ;

end

Algorithm 2: GeneratePosData: Generation of positive data with noise proportion p in ADS pipeline. Code uses IsNoiseClass function, explained later.

Data: Correct GO predictions, T
parameter: N_{neg} , number of reported negative GO classes per gene
output : Negative GO predictions, P_{neg}
begin
 Define $genes(T)$, the set of unique gene names in T ;
 Define $P_{neg} = []$;
 foreach $gene$ *in* $genes(T)$ **do**
 $GO_{count} = 0$;
 while $GO_{count} < N_{neg}$ **do**
 pick random GO class, GO_{rand} , from GO structure ;
 if $IsNoiseClass(GO_{rand}, gene)$ **then**
 Add ($gene, GO_{rand}$) as item to P_{neg} ;
 $GO_{count} = GO_{count} + 1$;
 end
 end
 end
 Return P_{neg} ;
end

Algorithm 3: GenerateNegData: Generation of negative data in ADS pipeline. Code uses IsNoiseClass function, explained later.

Data: Correct GO predictions, T
Data: Key list representing all ancestor classes for every GO class
Data: Evaluated GO class, GO_{eval}
Data: Evaluated gene, $gene_{eval}$
parameter: Noise threshold, th
 default: $th = 0.1$
output : TRUE or FALSE
begin
 | Get the set of ancestor classes, A_{eval} , for GO_{eval} ;
 | Define $J_{max} = 0$;
 | **foreach** row in T **do**
 | | Let $gene_{test}$ be the gene of the current row ;
 | | **if** $gene_{test}$ is $gene_{eval}$ **then**
 | | | Let GO_{test} be the GO class of the current row;
 | | | Get the set of ancestor classes, A_{test} , for GO_{test} ;
 | | | Calculate J_{test} , Jaccard correlation between A_{test} and A_{eval} ;
 | | | **if** $J_{test} > J_{max}$ **then**
 | | | | $J_{max} = J_{test}$
 | | | **end**
 | | **end**
 | **end**
 | **if** $J_{max} < th$ **then**
 | | Return TRUE ;
 | **else**
 | | Return FALSE ;
 | **end**
end

Algorithm 4: IsNoiseClass. This algorithm tests if GO class qualifies as noise. This is tested against the annotations of the selected gene