

netAna: A tool for network analysis

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Abstract

In present study, a software tool for network analysis is presented. The tool is mainly used to calculate topological properties of the network, detect type of the network, and calculate node centralities of the network. Various network properties include total number of nodes, total number of links, clustering coefficient of network, mean link density of network, connectance of network, skewness of degree distribution, aggregation index of network, variation coefficient of network, entropy of network, etc. Binomial distribution, Poisson distribution, exponential distribution, and particularly power law distribution are used for detection of network type. Node centralities used include degree centrality, closeness centrality, and betweenness centrality.

Keywords network analysis; software; network topology; power law; node centrality; crucial nodes.

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1 Introduction

Network analysis is one of the research focuses in network biology (Zhang, 2012, 2016g, 2018). In the past, various algorithms and tools on network analysis have been proposed and used in network biology (Jiang and Zhang, 2015; Jiang et al., 2015; Chen and Zhang, 2020; Xing and Zhang, 2020, 2021; Yang and Zhang, 2022; Zhang and Zhang, 2019; Zhang and Feng, 2017; Zhang and Qi, 2023; Zhang, 2012, 2016a-e, 2017a-b, 2018a-b, 2021, 2023). In present study, I will use Delphi developing environment (Zhang, 2023) to develop a standalone executable software tool for network analysis. It is expected to be powerful and easily used.

2 Algorithms and Software

2.1 Algorithms

All algorithms for detection of network type and calculation of network properties and node centralities, are based on Zhang and Zhan (2011), and Zhang (2016f), etc.

Various network properties include total number of nodes, total number of links, clustering coefficient of network, mean link density of network, connectance of network, skewness of degree distribution, aggregation index of network, variation coefficient of network, entropy of network (Zhang and Zhang, 2011; Zhang, 2012, 2016g, 2018). Binomial distribution, Poisson distribution, exponential distribution ($F(x)=1-e^{-\lambda x}$, $x \geq 0$),

particularly power law distribution ($p(x)=x^{-\alpha}$, $x \geq x_{\min}$) are used for detection of network type.

Node centralities used in present study include degree centrality, closeness centrality, and betweenness centrality (Zhang, 2012, 2016f, 2018).

2.2 Software codes

The full Object Pascal codes of netAna (version 1.0), developed in Delphi environment, are as follows:

```

unit Unit1;

interface

uses
  Windows, Messages, SysUtils, Variants, Classes, Graphics, Controls, Forms,
  Dialogs, StdCtrls, Math;

type
  TForm1 = class(TForm)
    OpenDialog1: TOpenDialog;
    SaveDialog1: TSaveDialog;
    Memo1: TMemo;
    Button1: TButton;
    Button2: TButton;
    procedure Rankings();
    procedure Dijkstra();
    procedure Button2Click(Sender: TObject);
    procedure Button1Click(Sender: TObject);
    procedure FormCreate(Sender: TObject);
  private
    { Private declarations }
  public
    { Public declarations }
  end;

var
  Form1: TForm1;
  nname: array[1..50000,1..3] of String;
  nodes: array[1..10000] of String;
  d: array of array of integer;
  distances: array of array of Double;
  centr: array of Extended;
  pat, rank: array of Integer;
  sss: Extended;
  v: integer;

implementation

{$R *.dfm}

procedure TForm1.Rankings();
var

```

```

s, i, j, k: Integer;
u: Double;
begin
setLength(rank,v+1);
for i:=1 to v do
rank[i]:=i;
for i:=1 to v-1 do
begin
k:=i;
for j:=i to v-1 do
if(centr[j+1]>centr[k]) then
k:=j+1;
s:=rank[i];
rank[i]:=rank[k];
rank[k]:=s;
u:=centr[i];
centr[i]:=centr[k];
centr[k]:=u;
end;
end;

procedure TForm1.Dijkstra();
var
a, b, p, w: array of Integer;
n, h, su, c, sd, ma, iv, i, j, k: integer;
begin
//d: weighted adjacency matrix; ss: total number of paths;
//pat: number of paths passing through each node;
//distances: matrix of distances between different nodes;
//paths: string of paths and distances between any of two nodes.
setLength(distances,v+1,v+1);
setLength(pat,v+1);
setLength(a,v+1);
setLength(b,v+1);
setLength(p,v+1);
setLength(w,v+1);
for i:=1 to v do
for j:=1 to v do
if ((d[i,j]=0) and (i<>j)) then
d[i,j]:=12345;
su:=0;
for j:=1 to v-1 do
begin
for k:=j+1 to v do
begin
for i:=1 to v do
begin
p[i]:=0;
w[i]:=0;
a[i]:=12345;

```

```

end;
a[j]:=0;
w[j]:=1;
n:=j;
h:=0;
while(v>0) do
begin
ma:=12345;
for i:=1 to v do
begin
if(w[i]=1) then
continue;
iv:=d[n,i]+a[n];
if(iv<a[i]) then
begin
a[i]:=iv;
b[i]:=n;
end;
if(a[i]>ma) then
continue;
ma:=a[i];
h:=i;
end;
w[h]:=1;
if(h=k) then
break;
n:=h;
end;
sd:=a[k];
p[1]:=k;
c:=k;
for i:=2 to v do
begin
if(c=j) then
break;
p[i]:=b[c];
c:=b[c];
end;
for i:=v downto 1 do
for h:=1 to v do
if(p[i]=h) then
begin
pat[h]:=pat[h]+1;
break;
end;
if(sd<>12345) then
begin
distances[j,k]:=sd;
distances[k,j]:=sd;
end;

```

```

if(sd=12345) then
su:=su+1;
end;
end;
for i:=1 to v do
for j:=1 to v do
if ((d[i,j]=12345) and (i<>j)) then
d[i,j]:=0;
sss:=v*(v-1)/2-su;
end;

procedure TForm1.Button2Click(Sender: TObject);
begin
Application.terminate;
end;

procedure TForm1.Button1Click(Sender: TObject);
begin
if SaveDialog1.Execute then
Memo1.Lines.SaveToFile(SaveDialog1.FileName+'.txt');
Application.terminate;
end;

procedure TForm1.FormCreate(Sender: TObject);
var
F: TextFile;
filename, str: String;
n, m, r, rr, rexp, i, j, k, tem, xm, xmin: Integer;
it, pp, qq, ss, meann, varr, skew, h, pr0, hk, chi, k1, k2: Extended;
a, alpha, alpha1, L, dd: Extended;
deg, frq, x, xminss, xmins, z: array of Integer;
p, pr, fr, dat, cx, cf: array of Single;
begin
Form1.visible:=false;
try
if OpenDialog1.Execute then
begin
filename:=OpenDialog1.Filename;
AssignFile(F,filename);
Reset(F);
n:=0;
while not Eof(F) do
begin
n:=n+1;
Readln(F,str);
i:=pos(',', str);
nname[n,1]:=copy(str,1,i-1);
str:=copy(str,i+1,length(str)-i);
i:=pos(',', str);
nname[n,2]:=copy(str,1,i-1);

```

```

nname[n,3]:=copy(str,i+1,length(str)-i);
nname[n,1]:=trim(nname[n,1]);
nname[n,2]:=trim(nname[n,2]);
nname[n,3]:=trim(nname[n,3]);
end;
CloseFile(F);
end;
v:=1;
nodes[1]:=nname[1,1];
for i:=1 to n do
for k:=1 to 2 do
begin
tem:=0;
for j:=1 to v do
if(sametext(nodes[j],nname[i,k])) then
begin
tem:=1;
break;
end;
if(tem=0) then
begin
v:=v+1;
nodes[v]:=nname[i,k];
end;
end;
memo1.Lines.Append("");
memo1.Lines.Append('I. Global properties of network');
memo1.Lines.Append("");
memo1.Lines.append('Total number of nodes='+inttostr(v));
memo1.Lines.append('Total number of links='+inttostr(n));
memo1.Lines.append("");
setLength(d,v+1,v+1);
for tem:=1 to n do
begin
for k:=1 to v do
if(sametext(nodes[k],nname[tem,1])) then
begin
i:=k;
break;
end;
for k:=1 to v do
if(sametext(nodes[k],nname[tem,2])) then
begin
j:=k;
break;
end;
d[i,j]:=1;
d[j,i]:=1;
end;
//Network type detection

```

```

rr:=10;
setLength(deg,v+1);
setLength(centr,v+1);
setLength(frq,rr+1);
a:=0;
for i:=1 to v do
begin
deg[i]:=0;
for j:=1 to v do
begin
if (abs(d[i,j])=1) then
deg[i]:=deg[i]+1;
end;
a:=a+deg[i];
centr[i]:=deg[i];
end;
memo1.Lines.append('Clustering coefficient of network: '+floattostr(a/v));
memo1.Lines.append('Mean link density of network: '+floattostr(n/v));
memo1.Lines.append('Connectance of network: '+floattostr(2*n/(v*v-v)));
memo1.Lines.Append(");
for i:=v downto 1 do
for j:=1 to v-1 do
if(deg[j]<deg[j+1]) then
begin
tem:=deg[j];
deg[j]:=deg[j+1];
deg[j+1]:=tem;
end;

pp:=0;

qq:=0;
for i:=1 to v do
begin
pp:=pp+deg[i];
qq:=qq+deg[i]*(deg[i]-1);
end;
it:=(deg[1]-deg[v])/rr;
for i:=1 to rr do
begin
frq[i]:=0;
for j:=1 to v do
begin
if((deg[j]>=(deg[v]+(i-1)*it)) and (deg[j]<(deg[v]+i*it))) then
frq[i]:=frq[i]+1;
end;
end;
rexp:=rr;
rr:=0;
for i:=1 to rexp do

```

```

if(frq[i]<>0) then
rr:=rr+1;
j:=0;
for i:=1 to rr do
begin
if(frq[i]<>0) then
j:=j+1;
end;
setLength(fr,j+1);
j:=0;
for i:=1 to rr do
begin
if(frq[i]<>0) then
begin
j:=j+1;
fr[j]:=frq[i];
end;
end;
memo1.Lines.Append('II. Network properties and network type detection');
memo1.Lines.Append('');
meann:=0;
for i:=1 to v do
meann:=meann+deg[i];
meann:=meann/v;
varr:=0;
for i:=1 to v do
varr:=varr+power(deg[i]-meann,2);
varr:=varr/(v-1);
skew:=v/((v-2)*sqrt(varr));
memo1.Lines.Append('Skewness of degree distribution: '+floattostr(skew)+');
memo1.Lines.Append('');
h:=v*qq/(pp*(pp-1));
memo1.Lines.Append('Aggregation index of network: '+floattostr(h)+');
if(h<=1) then
memo1.Lines.Append('It is likely a random network.')
else memo1.Lines.Append('It is likely a complex network.');
memo1.Lines.Append('');
h:=varr/meann;
memo1.Lines.Append('Variation coefficient H of network: '+floattostr(h)+');
memo1.Lines.Append('Entropy E of network: '+floattostr(varr-meann)+');
if(h<=1) then
memo1.Lines.Append('It is likely a random network.')
else memo1.Lines.Append('It is likely a complex network.');
memo1.Lines.Append('');
setLength(p,rr);
setLength(pr,rr);
//Binomial distri.: pr:= Crn pr qn-r, r:=0,1,2,..., n;
ss:=0;
for i:=1 to rr do
ss:=ss+fr[i]*(i-1);

```

```

pp:=ss/(v*(rr-1));
qq:=1-pp;
pr0:=power(qq,rr-1);
for i:=1 to rr-1 do
begin
if(i=1) then
pr[i]:=(rr-i)*pp*pr0/(i*qq)
else pr[i]:=(rr-i)*pp*pr[i-1]/(i*qq);
end;
chi:=0;
for i:=0 to rr-1 do
begin
if(i=0) then
hk:=pr0*v
else hk:=pr[i]*v;
if((i=0) and (pr0=0)) then
hk:=fr[i+1];
if((i>0) and (p[i]=0)) then
hk:=fr[i+1];
if((i=0) and (1/hk>5)) then
chi:=chi+power(pr0*v-fr[i+1],2)/hk
else chi:=chi+power(pr[i]*v-fr[i+1],2)/hk;
end;
memo1.Lines.Append('Binomial distribution Chi-square:='+floattostr(chi)+");
memo1.Lines.Append('Binomial p:='+floattostr(pp)+");
k1:=20.09;
if (chi<=k1) then
begin
memo1.Lines.Append('Node degrees are binomially distributed. ');
memo1.Lines.Append('It is likely a random network. ');
end
else
memo1.Lines.Append('It is likely not a random network.');
memo1.Lines.Append("");
//Poisson distri., pr := e^(-λ)λr/r!, r:=0,1,2,...
pr0:=exp(-meann);
for r:=1 to rr-1 do
begin
if(r=1) then
pr[r]:=meann/r*pr0
else pr[r]:=meann/r*pr[r-1];
end;
chi:=0;
for i:=0 to rr-1 do
begin
if(i=0) then
hk:=pr0*v
else hk:=pr[i]*v;
if((i=0) and (pr0=0)) then
hk:=fr[i+1];

```

```

if((i>0) and (p[i]=0)) then
hk:=fr[i+1];
if((i=0) and (1/hk>5)) then
chi:=chi+power(pr0*v-fr[i+1],2)/hk
else chi:=chi+power(pr[i]*v-fr[i+1],2)/hk;
end;
memo1.Lines.Append('Poisson distribution Chi-square:='+floattostr(chi)+");
memo1.Lines.Append('Poisson distribution λ:='+floattostr(meann)+");
k1:=20.09;
if(chi<=k1) then
begin
memo1.Lines.Append('Node degrees are Poisson distributed.');
memo1.Lines.Append('It is likely a random network.');
end
else memo1.Lines.Append('It is likely not a random network.');
memo1.Lines.Append(""));
//Exponential distri., F(x) =1-e^(-λx), x≥0
chi:=0;
for i:=1 to rexp do
begin
if(frq[i]<>0) then
begin
k1:=deg[v]+it/2+(i-1)*it;
k2:=deg[v]+it/2+i*it;
pp:=v*(exp(-k1/meann)-exp(-k2/meann));
chi:=chi+power(frq[i]-pp,2)/pp;
end;
end;
memo1.Lines.Append('Exponential distribution Chi-square:='+floattostr(chi)+");
memo1.Lines.Append('Exponential distribution λ (Exponential distri.: F(x)=1-e^(-λx), x≥0; x: node
degree):='+floattostr(1/meann)+");
k1:=20.09;
if(chi<=k1) then
memo1.Lines.Append('Node degrees are exponentially distributed.')
else
memo1.Lines.Append('Node degrees are not exponentially distributed.');
memo1.Lines.Append(""));
//Power law distri., p(x):=x^(-α), x≥xmin
setLength(x,v+1);
for i:=1 to v do
x[i]:=deg[i];
setLength(xminss,v+1);
m:=1;
xminss[1]:=x[1];
for i:=2 to v do
begin
tem:=0;
for j:=1 to m do
if(xminss[j]=x[i]) then
begin

```

```

tem:=1;
break;
end;
if(tem=0) then
begin
m:=m+1;
xminss[m]:=x[i];
end;
end;
setLength(xmins,m);
for i:=1 to m do
xminss[i]:=xminss[m-i+1];
setLength(dat,m);
setLength(z,v+1);
for i:=1 to v do
z[i]:=deg[v-i+1];
n:=v;
for xm:=1 to m-1 do
begin
xmin:=xmins[xm];
j:=0;
for i:=1 to n do
if(z[i]>=xmin) then
begin
j:=j+1;
z[j]:=z[i];
end;
n:=j;
a:=0;
for j:=1 to n do
a:=a+ln(z[j]/xmin);
a:=n/a;
setLength(cx,n+1);
setLength(cf,n+1);
for j:=1 to n do
cx[j]:=(j-1)/n;
for j:=1 to n do
cf[j]:=1-power(xmin/z[j],a);
dat[xm]:=-1e+20;
for j:=1 to n do
if(abs(cf[j]-cx[j])>dat[xm]) then
dat[xm]:=abs(cf[j]-cx[j]);
end;
dd:=1e+20;
for j:=1 to m-1 do
if(dat[j]<dd) then
dd:=dat[j];
for i:=1 to m-1 do
if(dat[i]<=dd) then
begin

```

```

xmin:=xmins[i];
break;
end;
j:=0;
for i:=1 to v do
if(x[i]>=xmin) then
j:=j+1;
setLength(z,j+1);
j:=0;
for i:=1 to v do
if(x[i]>=xmin) then
begin
j:=j+1;
z[j]:=x[i];
end;
n:=j;
alpha1:=0;
for i:=1 to n do
alpha1:=alpha1+ln(z[i]/xmin);
alpha:=1+n/alpha1;
L:=n*ln((alpha-1)/xmin)-alpha*alpha1;
//L is the log-likelihood of the data x>=xmin under the fitted power law.
memo1.Lines.Append('Power law distribution Kolmogorov-Smirnov goodness-of-fit statistic D:='+floattostr(dd)+");
if(dd<(1.63/sqrt(n))) then
memo1.Lines.Append('Node degrees are power-law distributed (Power law distri.: p(x)=x^(-α), x≥xmin; x: node degree), thus it
is a scale-free complex network.');
memo1.Lines.Append('Power law α:='+floattostr(alpha)+");
memo1.Lines.Append('Power law xmin:='+floattostr(xmin)+");
memo1.Lines.Append(");
memo1.Lines.Append('III. Node centralities');
memo1.Lines.Append(");
//Node centrality calculation
memo1.Lines.Append('Degree centrality');
memo1.Lines.Append(");
memo1.Lines.Append('Rank'+' '+'Node'+' '+'Degree centrality');
Ranks();
for i:=1 to v do
memo1.Lines.Append(inttostr(i)+' '+'nodes[rank[i]]+' +'floattostr(centr[i]));
memo1.Lines.Append(");
memo1.Lines.Append('In terms of degree centrality, '+nodes[rank[1]]+' and '+nodes[rank[2]]+', etc., are the most crucial nodes in
the network.');
memo1.Lines.Append('And '+nodes[rank[v]]+' and '+nodes[rank[v-1]]+', etc., are the most trivial nodes in the network.');
memo1.Lines.Append(");
memo1.Lines.Append('Closeness centrality');
memo1.Lines.Append(");
memo1.Lines.Append('Rank'+' '+'Node'+' '+'Closeness centrality');
Dijkstra();
for i:=1 to v do
begin
ss:=0;

```

```

for j:=1 to v do
ss:=ss+distances[i,j];
centr[i]:=1/ss;
end;
Rankings();
for i:=1 to v do
memo1.Lines.Append(inttostr(i)+" "+nodes[rank[i]]+" "+floattostr(centr[i]));
memo1.Lines.Append(");
memo1.Lines.Append('In terms of closeness centrality, '+nodes[rank[1]]+" and "+nodes[rank[2]]+", etc., are the most crucial
nodes in the network.');
memo1.Lines.Append('And '+nodes[rank[v]]+" and "+nodes[rank[v-1]]+", etc., are the most trivial nodes in the network.");
memo1.Lines.Append(");
memo1.Lines.Append('Betweenness centrality');
memo1.Lines.Append(");
memo1.Lines.Append('Rank'+" "+'Node'+" "+'Betweenness centrality');
for i:=1 to v do
centr[i]:=pat[i]/sss;
Rankings();
for i:=1 to v do
memo1.Lines.Append(inttostr(i)+" "+nodes[rank[i]]+" "+floattostr(centr[i]));
memo1.Lines.Append(");
memo1.Lines.Append('In terms of betweenness centrality, '+nodes[rank[1]]+" and "+nodes[rank[2]]+", etc., are the most crucial
nodes in the network.');
memo1.Lines.Append('And '+nodes[rank[v]]+" and "+nodes[rank[v-1]]+", etc., are the most trivial nodes in the network.");
memo1.Lines.Append(");
except
exit;
end;
end;

end.

```

2.3 Data

The data for the network is stored in a plain text file. Suppose there are totally m links in the network, then there are m rows in the data file. There are three items in each row: the first item (node) and the second item (node) represent a link between two nodes and the third item is an integer 1. The three items are separated by the comma “,” (so by default, it means that any comma is not allowed in the first and second items).

The software and demo data files are included in the package:

[http://www.iaees.org/publications/journals/nb/articles/2023-13\(4\)/e-suppl/Zhang-Supplementary-Material3.rar](http://www.iaees.org/publications/journals/nb/articles/2023-13(4)/e-suppl/Zhang-Supplementary-Material3.rar)

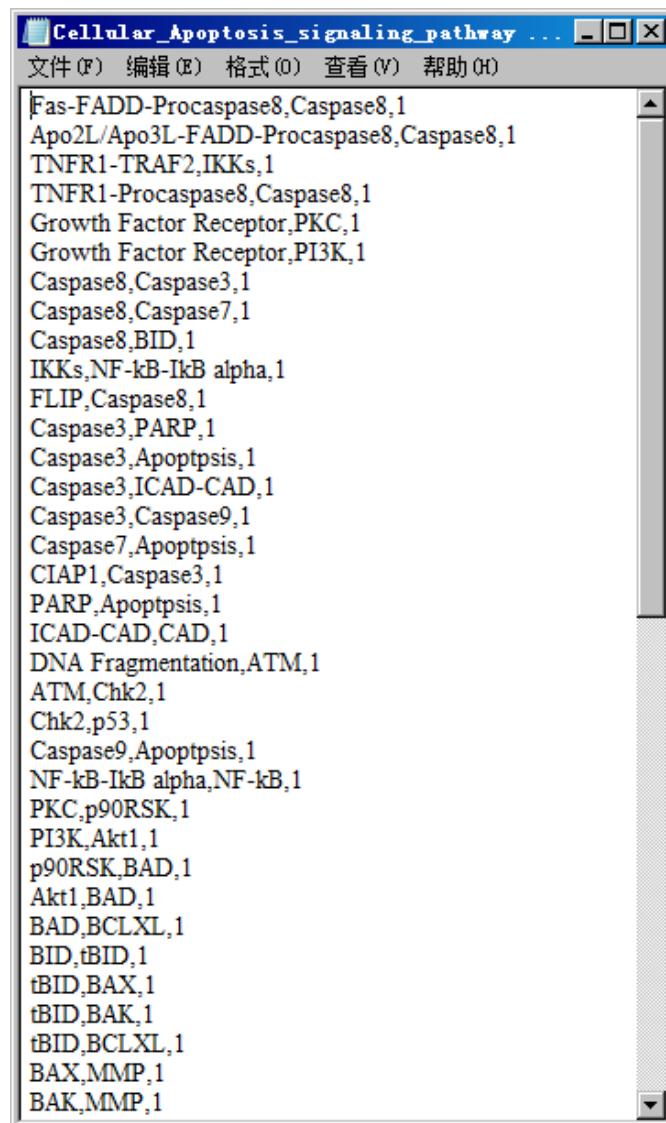


Fig. 2 A data file for netAna.

3 Demonstration

The network data for cellular apoptosis signaling pathway were obtained from Huang and Zhang (2012), Li and Zhang (2013), which was stored in a text file (Cellular_Apoptosis_signaling_pathway.txt), as shown in Fig. 2.

Running netAna and loading the data file, the results (Fig. 3) are listed bellow:

I. Global properties of network

Total number of nodes=49

Total number of links=67

Clustering coefficient of network: 2.73469387755102

Mean link density of network: 1.36734693877551

Connectance of network: 0.0569727891156463

II. Network properties and network type detection

Skewness of degree distribution: 0.53606751912101

Aggregation index of network: 1.13825608798115

It is likely a complex network.

Variation coefficient H of network: 1.38308457711443

Entropy E of network: 1.04761904761905

It is likely a complex network.

Binomial distribution Chi-square:=47.7728601257816

Binomial p:=0.255102040816327

It is likely not a random network.

Poisson distribution Chi-square:=105.214297758277

Poisson distribution λ :=2.73469387755102

It is likely not a random network.

Exponential distribution Chi-square:=65.7383685892767

Exponential distribution λ (Exponential distri.: $F(x)=1-e^{-\lambda x}$, $x \geq 0$; x: node degree):=0.365671641791045

Node degrees are not exponentially distributed.

Power law distribution Kolmogorov-Smirnov goodness-of-fit statistic D:=0.214274525642395

Node degrees are power-law distributed (Power law distri.: $p(x)=x^{-\alpha}$, $x \geq x_{\min}$; x: node degree), thus it is a scale-free complex network.

Power law α :=3.27938463678571

Power law x_{\min} :=3

III. Node centralities

Degree centrality

| Rank | Node | Degree centrality |
|------|-------------------|-------------------|
| 1 | MMP | 11 |
| 2 | Caspase8 | 7 |
| 3 | Caspase3 | 6 |
| 4 | Apoptpsis | 6 |
| 5 | BAK | 6 |
| 6 | BCL2 | 6 |
| 7 | p53 | 5 |
| 8 | CIAP1 | 4 |
| 9 | DNA Fragmentation | 4 |
| 10 | BCLXL | 4 |
| 11 | tBID | 4 |

| | | |
|----|------------------------------|---|
| 12 | Caspase9 | 3 |
| 13 | BAD | 3 |
| 14 | BAX | 3 |
| 15 | BIM | 3 |
| 16 | ICAD-CAD | 2 |
| 17 | BID | 2 |
| 18 | PKC | 2 |
| 19 | CAD | 2 |
| 20 | PI3K | 2 |
| 21 | ATM | 2 |
| 22 | Chk2 | 2 |
| 23 | Growth Factor Receptor | 2 |
| 24 | p90RSK | 2 |
| 25 | Akt1 | 2 |
| 26 | NF-kB-IkB alpha | 2 |
| 27 | Caspase7 | 2 |
| 28 | IKKs | 2 |
| 29 | VDAC | 2 |
| 30 | PARP | 2 |
| 31 | PUMA | 2 |
| 32 | Noxa | 2 |
| 33 | SMAC | 2 |
| 34 | Arts | 2 |
| 35 | HTRA2 | 2 |
| 36 | Cytoc | 2 |
| 37 | EndoG | 2 |
| 38 | AIF | 2 |
| 39 | APAF1 | 2 |
| 40 | MDM2 | 2 |
| 41 | TNFR1-Procaspase8 | 1 |
| 42 | NF-kB | 1 |
| 43 | FLIP | 1 |
| 44 | TNFR1-TRAF2 | 1 |
| 45 | Fas-FADD-Procaspase8 | 1 |
| 46 | p53AIP1 | 1 |
| 47 | p14(ARF) | 1 |
| 48 | Apo2L/Apo3L-FADD-Procaspase8 | 1 |
| 49 | POD | 1 |

In terms of degree centrality, MMP and Caspase8, etc., are the most crucial nodes in the network. And POD and Apo2L/Apo3L-FADD-Procaspase8, etc., are the most trivial nodes in the network.

Closeness centrality

| Rank | Node | Closeness centrality |
|------|------|----------------------|
|------|------|----------------------|

| | | |
|----|------------------------------|---------------------|
| 1 | IKKs | 0.25 |
| 2 | NF-kB-IkB alpha | 0.25 |
| 3 | TNFR1-TRAF2 | 0.1666666666666667 |
| 4 | NF-kB | 0.1666666666666667 |
| 5 | MMP | 0.0091743119266055 |
| 6 | BAK | 0.00806451612903226 |
| 7 | Apoptosis | 0.00793650793650794 |
| 8 | BCLXL | 0.0078125 |
| 9 | BCL2 | 0.00763358778625954 |
| 10 | Caspase3 | 0.00709219858156028 |
| 11 | BAX | 0.00709219858156028 |
| 12 | tBID | 0.00699300699300699 |
| 13 | SMAC | 0.00699300699300699 |
| 14 | Arts | 0.00699300699300699 |
| 15 | HTRA2 | 0.00699300699300699 |
| 16 | EndoG | 0.00699300699300699 |
| 17 | AIF | 0.00699300699300699 |
| 18 | Cytoc | 0.00671140939597315 |
| 19 | CIAP1 | 0.00641025641025641 |
| 20 | Caspase7 | 0.00641025641025641 |
| 21 | Caspase9 | 0.00641025641025641 |
| 22 | BIM | 0.00636942675159236 |
| 23 | BID | 0.00628930817610063 |
| 24 | Caspase8 | 0.00628930817610063 |
| 25 | PARP | 0.00628930817610063 |
| 26 | VDAC | 0.00628930817610063 |
| 27 | BAD | 0.0062111801242236 |
| 28 | PUMA | 0.00617283950617284 |
| 29 | Noxa | 0.00617283950617284 |
| 30 | DNA Fragmentation | 0.00617283950617284 |
| 31 | APAF1 | 0.00591715976331361 |
| 32 | POD | 0.00591715976331361 |
| 33 | ICAD-CAD | 0.00588235294117647 |
| 34 | CAD | 0.00558659217877095 |
| 35 | p53 | 0.00531914893617021 |
| 36 | ATM | 0.00523560209424084 |
| 37 | Akt1 | 0.005 |
| 38 | p90RSK | 0.005 |
| 39 | FLIP | 0.00495049504950495 |
| 40 | Apo2L/Apo3L-FADD-Procaspase8 | 0.00495049504950495 |
| 41 | Fas-FADD-Procaspase8 | 0.00495049504950495 |
| 42 | TNFR1-Procaspase8 | 0.00495049504950495 |
| 43 | Chk2 | 0.00469483568075117 |
| 44 | MDM2 | 0.00436681222707424 |
| 45 | p53AIP1 | 0.00432900432900433 |

| | | |
|----|------------------------|---------------------|
| 46 | PKC | 0.00418410041841004 |
| 47 | PI3K | 0.00418410041841004 |
| 48 | p14(ARF) | 0.00367647058823529 |
| 49 | Growth Factor Receptor | 0.00359712230215827 |

In terms of closeness centrality, IKKs and NF-kB-IkB alpha, etc., are the most crucial nodes in the network. And Growth Factor Receptor and p14(ARF), etc., are the most trivial nodes in the network.

Betweenness centrality

| Rank | Node | Betweenness centrality |
|------|------------------------------|------------------------|
| 1 | Caspase8 | 2.94477911646586 |
| 2 | Apo2L/Apo3L-FADD-Procaspase8 | 1.93373493975904 |
| 3 | BID | 0.938755020080321 |
| 4 | p53 | 0.76004016064257 |
| 5 | MMP | 0.613453815261044 |
| 6 | IKKs | 0.610441767068273 |
| 7 | Chk2 | 0.610441767068273 |
| 8 | BCL2 | 0.345381526104418 |
| 9 | TNFR1-TRAF2 | 0.336345381526104 |
| 10 | Apoptpsis | 0.295180722891566 |
| 11 | BCLXL | 0.295180722891566 |
| 12 | NF-kB-IkB alpha | 0.28714859437751 |
| 13 | BAD | 0.261044176706827 |
| 14 | Noxa | 0.234939759036145 |
| 15 | Caspase3 | 0.189759036144578 |
| 16 | tBID | 0.178714859437751 |
| 17 | BAK | 0.141566265060241 |
| 18 | DNA Fragmentation | 0.13855421686747 |
| 19 | Caspase7 | 0.134538152610442 |
| 20 | Akt1 | 0.133534136546185 |
| 21 | AIF | 0.132530120481928 |
| 22 | MDM2 | 0.0953815261044177 |
| 23 | p90RSK | 0.0913654618473896 |
| 24 | PI3K | 0.0893574297188755 |
| 25 | ICAD-CAD | 0.0883534136546185 |
| 26 | ATM | 0.0833333333333333 |
| 27 | CAD | 0.0803212851405622 |
| 28 | Cytoc | 0.0773092369477912 |
| 29 | HTRA2 | 0.0742971887550201 |
| 30 | CIAP1 | 0.0742971887550201 |
| 31 | Caspase9 | 0.073293172690763 |
| 32 | BIM | 0.0592369477911647 |
| 33 | APAF1 | 0.0582329317269076 |
| 34 | BAX | 0.0481927710843374 |

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35 VDAC      0.0481927710843374
36 PUMA      0.0481927710843373
37 SMAC      0.0481927710843373
38 Arts       0.0481927710843373
39 EndoG     0.0481927710843373
40 p53AIP1   0.0481927710843373
41 p14(ARF)  0.0481927710843373
42 POD       0.0481927710843373
43 PKC       0.0471887550200803
44 FLIP      0.0471887550200803
45 PARP      0.0471887550200803
46 Growth Factor Receptor 0.0471887550200803
47 TNFR1-Procaspase8    0.0461847389558233
48 Fas-FADD-Procaspase8 0.0441767068273092
49 NF-kB      0.0240963855421687

```

In terms of betweenness centrality, Caspase8 and Apo2L/Apo3L-FADD-Procaspase8, etc., are the most crucial nodes in the network.

And NF-kB and Fas-FADD-Procaspase8, etc., are the most trivial nodes in the network.

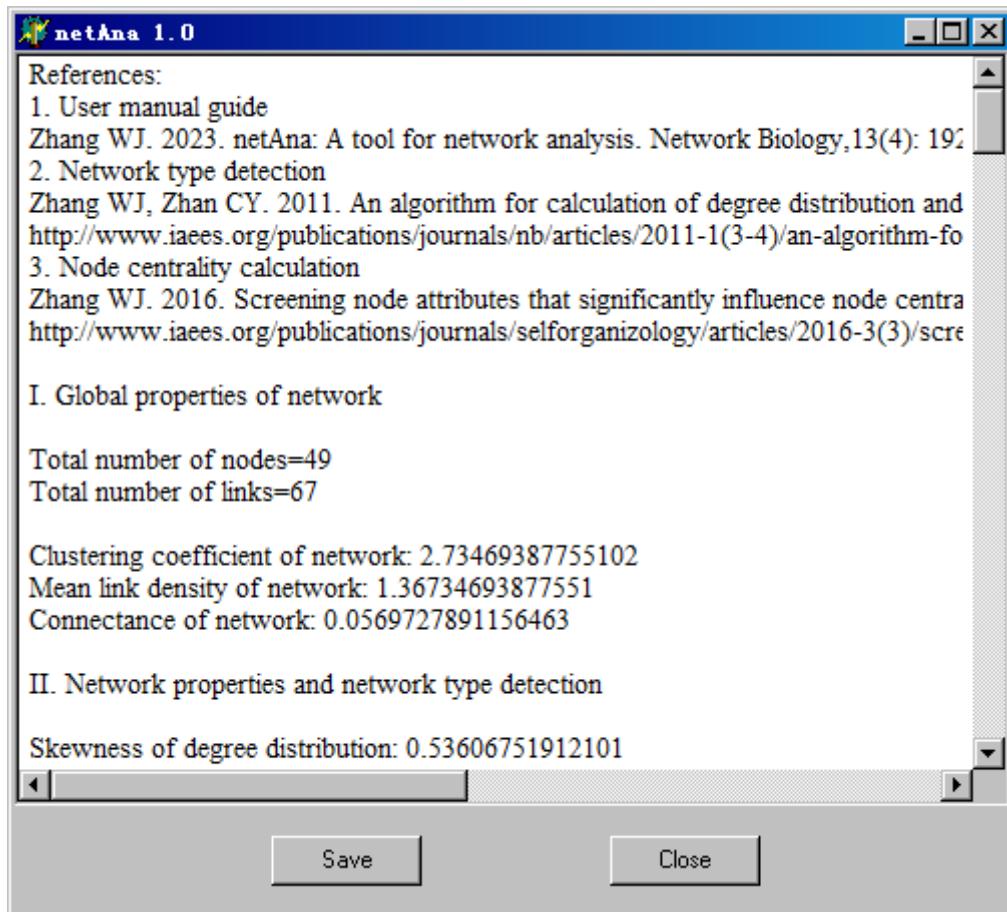


Fig. 3 Results of network analysis from netAna.

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