Draw loops from enumerated reaction networks December 2014

Initializations

```
> restart:
> interface(rtablesize = 400):
> with(ListTools):
> with(LinearAlgebra):
> with(VectorCalculus):
> with(GraphTheory):
> with(combinat):
> with(ArrayTools):
> with(FileTools):
> _Envsignum0 := 0:
```

Functions for constructing graph from stoichiometric matrix

```
| findZ := proc(A)
| local Z, n, m, i, j :
| n := Dimension(A)[1]:
| m := Dimension(A)[2]:
| Z := Matrix(n, m) :
| for i from 1 to n by 1 do
| for j from 1 to m by 1 do
| if A[i,j] < 0 then Z[i,j] := z[i,j]; end if; ### what is the z?
| end do:
| return(Z) :
| end proc:</pre>
```

Find the DSR graph from labels, A and Z

```
> ##Create signed DSR graph: entries are two matrices and the labels of the nodes createDSRgraphsigned := proc(mynodes, A, Z)
local G, n, m, Adj, varsZ, Zsign, varsA, Asign, X:
n := Dimension(A)[1]: m := Dimension(A)[2]:
X := Transpose(Z):
```

```
varsZ := indets(X) :
Zsign := subs(seq(varsZ[i] = 1, i = 1 ..numelems(varsZ)), X) :
Adj := Matrix(n + m, n + m) :
Adj[[n + 1 ..n + m], [1 ..n]] := Transpose(map(signum, A)) :
Adj[[1 ..n], [n + 1 ..n + m]] := Transpose(Zsign) :
G := GraphTheory[Graph](mynodes, Adj, weighted = true) :
return(G) :
end proc:
```

Find the DSR graph from labels and A and return the list of edges:

```
> findedgesDSR := proc(mynodes, A)
    local G, Z:
    Z := findZ(A):
    G := createDSRgraphsigned(mylabels, A, Z):
    return(Edges(G, weights)):
    end proc:

>
```

Functions for draw loops from graph

Auxiliary procedures

This procedure computes the polynomial $p_{A, Z}$ in the main text. The input are the matrices A and Z (in the function denoted N and X).

```
end if:
  bigdet := expand(Determinant(Mt)):
  return(bigdet):
  end proc:
```

This function returns the list of monomials that have the wrong sign. The input are the determinant and the wrong sign.

```
> ##Given a determinant and a wrong sign, return the list of wrong monomials
badterms := proc(deter, mysign)
    local vars, coeflist, monomlist, coeflistsign, wterms, i:
    vars := indets(deter):
    coeflist := [coeffs(deter, vars, 't')]:
    monomlist := [t]:
    coeflistsign := map(sign, coeflist):
    wterms := []:
    for i from 1 by 1 to numelems(coeflistsign) do
        if mysign = coeflistsign[i] then wterms := [op(wterms), monomlist[i]]: end if:
    end do:
    return(wterms):
```

Given a monomial on the entries of a matrix Amatrix, this function finds a matrix from Amatrix such that the variables in the monomial become 1 and the rest are zero.

```
> ##Find submatrix of a matrix corresponding to a monomial
findmatrix := proc(wmonom, Amatrix)
    local vars, wvarscomp, Anew, i, Anew1:
    vars := indets(wmonom):
    wvarscomp := indets(Amatrix) minus vars:
    Anew := subs(seq(wvarscomp[i] = 0, i = 1 ..numelems(wvarscomp)), Amatrix):
    Anew1 := subs(seq(vars[i] = 1, i = 1 ..numelems(vars)), Anew):
    return(Anew1):
    end proc:
```

Find the two submatrices of A and Z corresponding to the monomial, and make A symbolic by introducing a new variable x.

```
> ## Find the two matrices A,Z corresponding to a monomial
twomatrices := proc(wmonom, A, Z)
local vars, wvarscomp, Zembed, i, row, col, Aembed, Aembedx, nZ, X:
    X := Transpose(Z):
    vars := indets(wmonom):
    wvarscomp := indets(X) minus vars:
    Zembed := subs(seq(wvarscomp[i] = 0, i = 1 ..numelems(wvarscomp)), X):
    row, col := ArrayTools[SearchArray](Zembed):
```

```
Zembed := Zembed[convert(row, list), convert(col, list)]:
       nZ := numelems(vars):
      Aembed := A[convert(col, list), convert(row, list)]:
      Aembed := map(signum, Aembed):
      row, col := ArrayTools[SearchArray](Aembed):
      Aembedx := Matrix(nZ, nZ):
      for i from 1 by 1 to numelems (col) do
       Aembedx[row[i], col[i]] := Aembed[row[i], col[i]] \cdot x_i:
      end do:
      return (Aembedx, Zembed):
   end proc:
Extract the indices of the species and the reaction in the given monomial in the variables of Z
> ## Extract indices of species and reaction in the monomial from Z
   extractsr := \mathbf{proc}(wmonom, Z)
      local vars, wvarscomp, Zembed, i, row, col, X:
      X := Transpose(Z):
       vars := indets(wmonom):
      wvarscomp := indets(X)  minus vars:
                                              ## indeterminates not in the monomial
      Zembed := subs(seq(wvarscomp[i] = 0, i = 1 ..numelems(wvarscomp)), X):
       ##set the entries of the indeterminates not in the monomial to zero
      row, col := ArrayTools[SearchArray](Zembed):
       ##find the nonzero entries of the resulting matrix.
      return (row, col): ##return the species and reaction indices
    end proc:
   ##Create DSR graph: entries are two matrices and the labels of the nodes
   createDSRgraph := \mathbf{proc}(mynodes, A, Z)
      local G, n, m, Adj, varsZ, Zsign, varsA, Asign, X:
     n := Dimension(A)[1]: m := Dimension(A)[2]:
     X := Transpose(Z):
      varsZ := indets(X):
      Zsign := subs(seq(varsZ[i] = 1, i = 1 ..numelems(varsZ)), X):
      varsA := indets(A):
      Asign := subs(seq(varsA[i] = 1, i = 1 ..numelems(varsA)), A):
    Adj := Matrix(n + m, n + m):
    Adj[[n+1..n+m], [1..n]] := Transpose(map(signum, Asign)):
    Adj[[1..n], [n+1..n+m]] := Transpose(Zsign):
     G := GraphTheory[Graph](mynodes, Adj, weighted = true):
     return(G):
```

This function selects the subgraphs that give rise to the monomials with the wrong sign.

end proc:

```
## Select the subgraphs that correspond to the wrong terms of A and Z
graphlist := \mathbf{proc}(mydet, A, Z)
local srlist, row, col, Gsub, s, wsign, wrongterms, k, wcurrent, Aembedx, Zembed, detZ,
    detAx, wsignA, wrongtermsA, wcurrentA, j, Aembedx1, mynodes:
Gsub := []:
srlist := []:
s := Rank(A):
wsign := (-1)^{s+1}:
                                          ## find wrong sign
wrong terms := badterms(mydet, wsign): ## select the monomials with the wrong sign
for k from 1 by 1 to numelems (wrongterms) do
    ## for each such monomial, find the associated subgraph
   wcurrent := wrongterms[k]:
   row, col := extractsr(wcurrent, Z):
    ## find the indices of the species and the reactions in the monomial
   mynodes := [seq(S_{col[i]}, i = 1 ..numelems(col)), seq(R_{row[i]}, i = 1)]
    ..numelems(row))]:
   Aembedx, Zembed := two matrices(wcurrent, A, Z):
    ## the returned Zembed is giving half of the edges of the subgraphs
    detZ := subs(seq(indets(Zembed))[i] = 1, i = 1 ...numelems(indets(Zembed))),
    Determinant(Zembed)):
   detAx := expand(Determinant(Aembedx)):
   wsignA := wsign \cdot detZ:
   wrong terms A := badterms(det Ax, wsign A):
    ## select the monomials with the wrong sign of the subsystem
    for j from 1 by 1 to numelems (wrongtermsA) do
      wcurrentA := wrongtermsA[j]:
      Aembedx1 := findmatrix(wcurrentA, Aembedx):
    ## find the other half of the edges of the subgraphs
       Gsub := [op(Gsub), createDSRgraph(mynodes, Aembedx1, Zembed)]:
   end do:
 end do:
 return (Gsub): ##return the list of graphs
 end proc:
```

Given a list of edges that form a loop, the function returns the edges ordered such that connected they form the loop.

```
> ## Order the edges to have a loop
orderedge := proc(myedges)
local orderededges, endpoint, total, control, k:
orderededges := [myedges[1]]:
endpoint := myedges[1][1][2]:
total := numelems(myedges):
while numelems(orderededges) < total do
control := 0: k := 2:
while control = 0 do
if endpoint = myedges[k][1][1] then</pre>
```

```
orderededges := [op(orderededges), myedges[k]]:
    control := 1 :
    endpoint := myedges[k][1][2]:

end if:
    k := k + 1 :
    end do:
    end do:
    return(orderededges) :
    end proc:
```

Find the sequence of signs of the loop

```
> ##Extract the sequence of signs of a loop
extractsign := proc(orderededges)
    local graphsign, i:
    graphsign := []:
    for i from 1 by 1 to numelems(orderededges) do
        graphsign := [op(graphsign), orderededges[i][2]]:
    end do:
    return(graphsign):
    end proc:
```

Given a list of graphs, we find the positive feedback loops that they contain and return the sign pattern of each positive feedback loop as well (as those given in Table 1 in the main text).

```
> ##Find the positive feedback loops in the list of graphs
  positive feed := proc(Gsub)
     local selected, j, mygraph, Gsubcomp, k, mycomp, newgraph, wedges, myprod, i,
      signcycle:
     selected := []:
     signcycle := []:
     Gsubcomp := []:
     for j from 1 by 1 to numelems (Gsub) do
       mygraph := Gsub[j]:
      Gsubcomp := ConnectedComponents(mygraph):
      for k from 1 by 1 to numelems (Gsubcomp) do
        mycomp := Gsubcomp[k]:
       newgraph := InducedSubgraph(mygraph, mycomp):
       wedges := Edges(newgraph, weights):
       myprod := mul(wedges[i][2], i = 1 ..numelems(wedges)):
       if myprod = 1 then
                                   ##if the loop is positive, select it
          selected := [op(selected), [op(wedges)]]:
       end if
     end do:
    end do:
```

```
selected := ListTools[MakeUnique](selected):
for k from 1 by 1 to numelems(selected) do
    selected[k] := orderedge(selected[k]):
    signcycle := [op(signcycle), extractsign(selected[k])]:
    end do:
    return(selected, signcycle):
    end proc:
```

The main procedure to find the positive loop is the following:

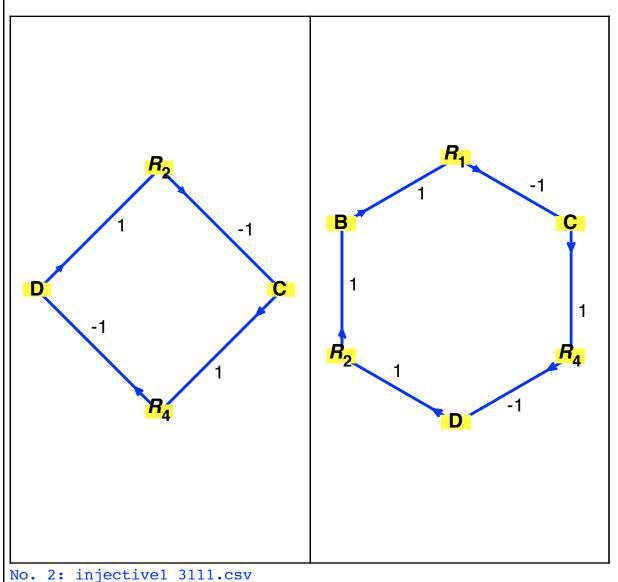
```
*##main program: find the positive loops
findloops := proc(A, Z)
    local Gsub, selected, signcycle, mydet:
        mydet := computdetS(A, Z): ## find the polynomial p<sub>A, Z</sub>
    Gsub := graphlist(mydet, A, Z):
        ## find the list of subgraphs corresponding to the wrong signs
        selected, signcycle := positivefeed(Gsub):
        ## find the positive feedback loops and their sign pattern
    return(selected):
    end proc:
```

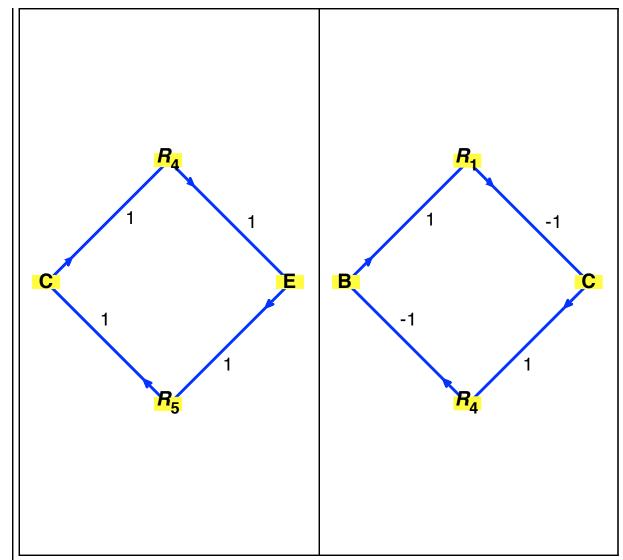
The second main procedure of the method is the function that draws the selected positive feedback loops. It requires a list with the names of the nodes (see the examples below)

```
| ## draw the positive feedback loops
| drawloops := proc(selected, speciesord)
| local loops, i, vertices, speciesdic, selected2 :
| loops := []:
| speciesdic := {seq(S<sub>i</sub> = speciesord[i], i = 1 ..numelems(speciesord))}:
| selected2 := subs(speciesdic, selected) :
| for i from 1 by 1 to numelems(selected2) do
| vertices := ListTools[MakeUnique]([seq(op(selected2[i][j][1]), j = 1 |
| ..numelems(selected2[i]))]):
| loops := [op(loops), Digraph(vertices, {op(selected2[i])})]:
| end do:
| DrawGraph(loops, style = circle);
| end proc:
```

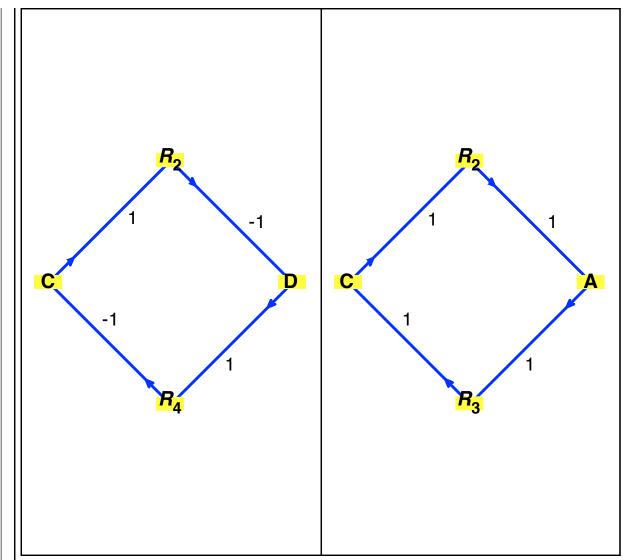
Non-multistationary: with competition loop, with intersecting loops

```
| uniquefolder := "5species/nonmultistationary/unique_competitionloop_intersectingloops" :
| uniquefiles := ListDirectory(uniquefolder) :
| m := nops(uniquefiles) :
| for i from 1 to m by 1 do uni := op(i, uniquefiles) : A := ImportMatrix(cat(uniquefolder, "/", uni)) : Z := findZ(A) : selected := findloops(A, Z) : printf("No. %d: %s \n", i, uni); print(drawloops(selected, speciesord)); end do:
| No. 1: injective1_2962.csv
```

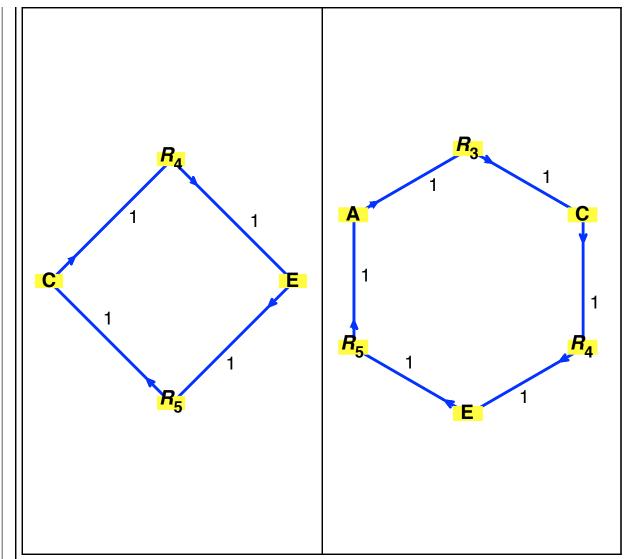




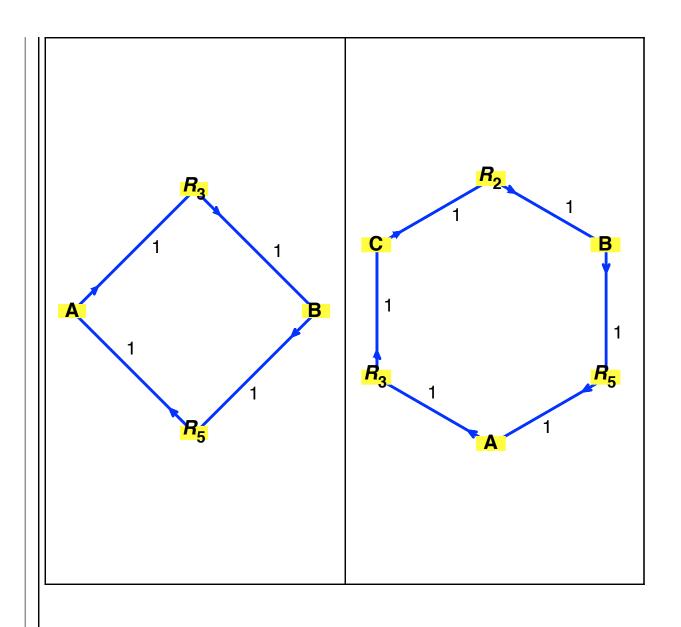
No. 3: injective1_3119.csv

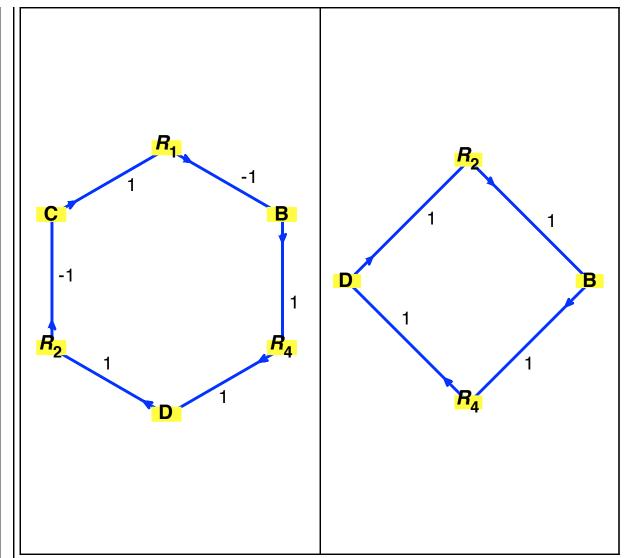


No. 4: injectiveEx1_1267.csv

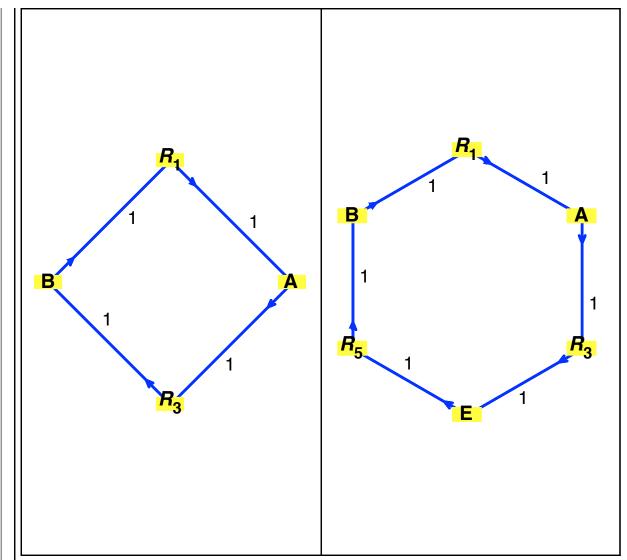


No. 5: injectiveEx1_2956.csv

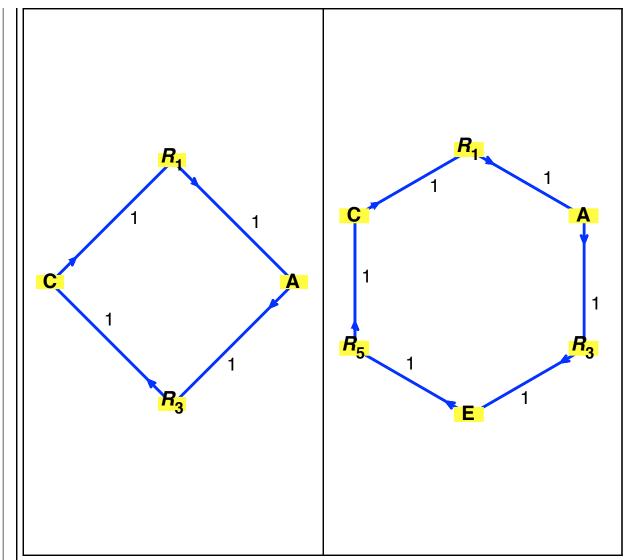




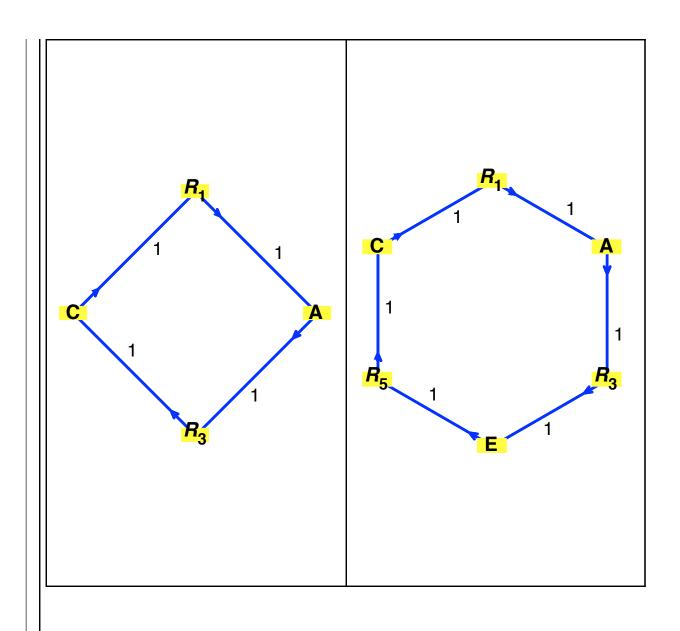
No. 6: injectiveEx1_2979.csv

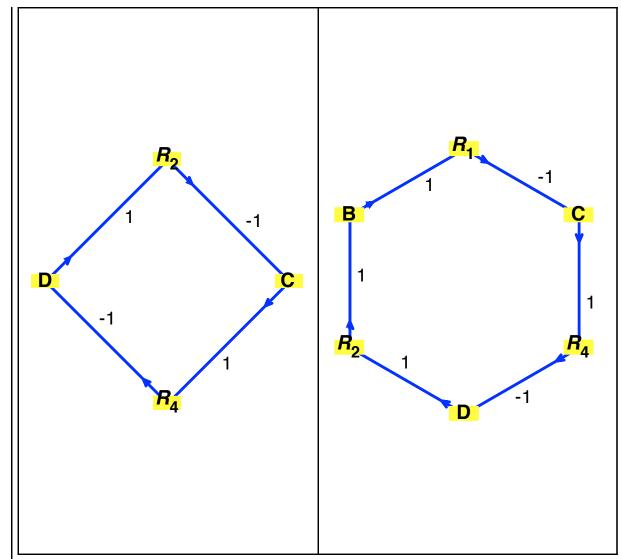


No. 7: injectiveEx1_2987.csv

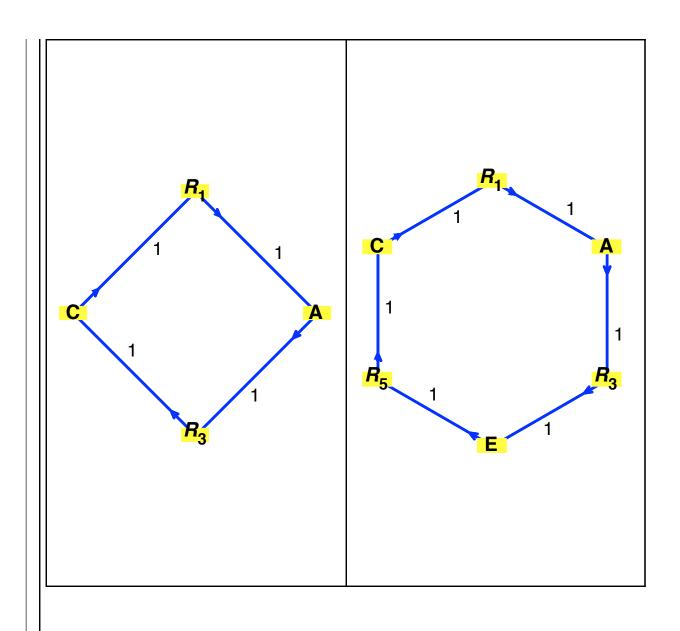


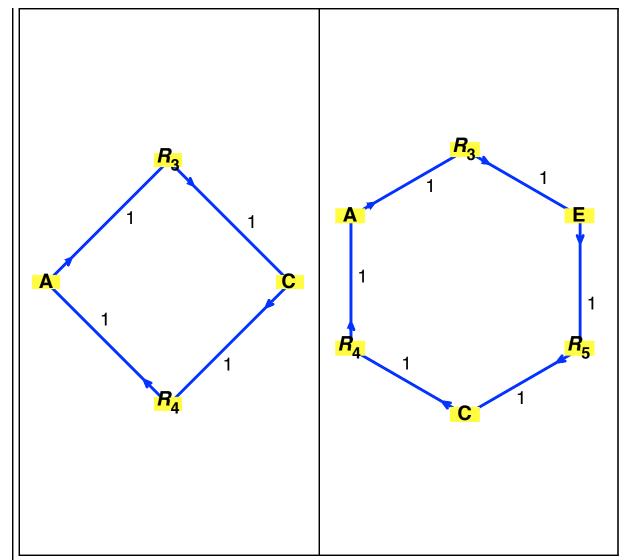
No. 8: injectiveEx1_2988.csv



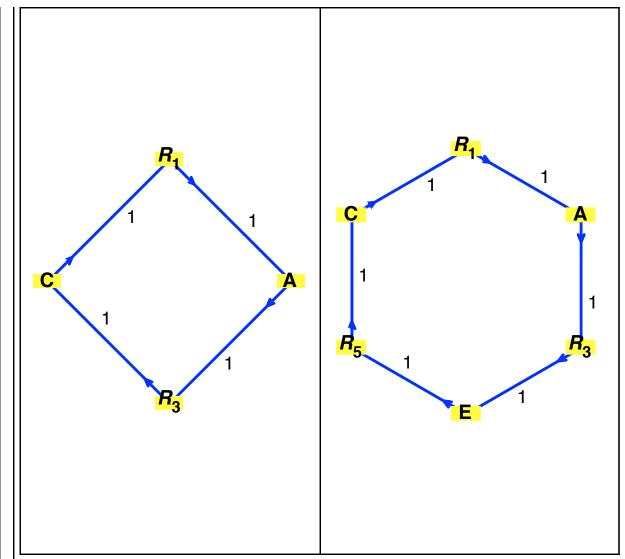


No. 9: injectiveEx1_2989.csv

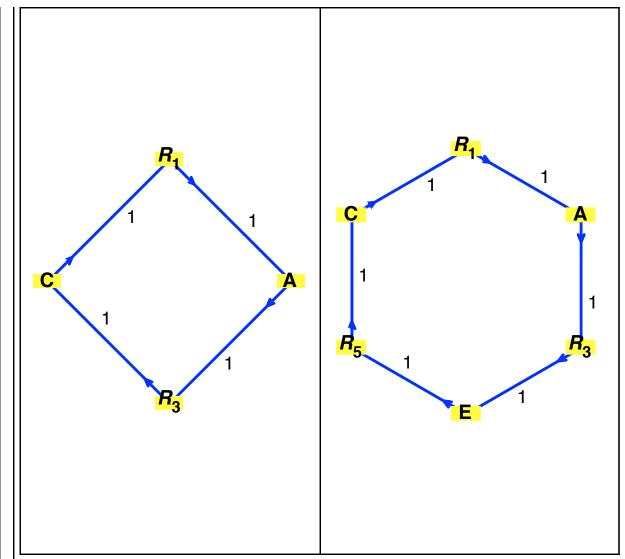




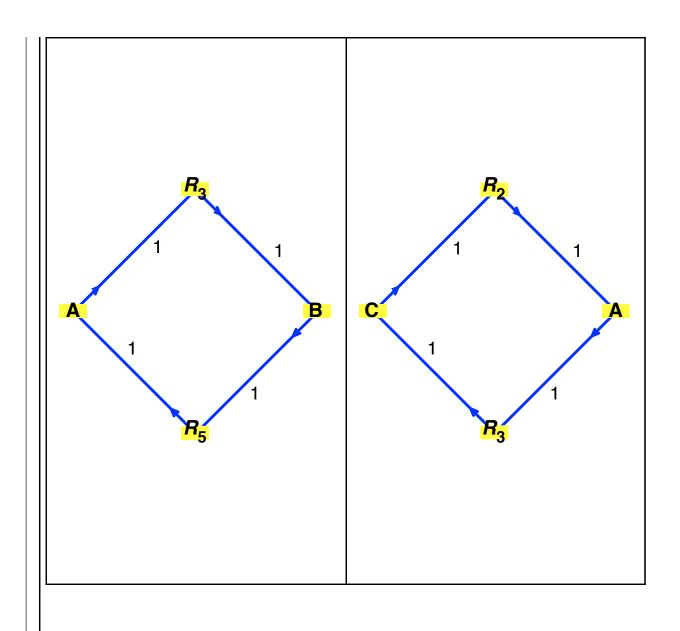
No. 10: injectiveEx1_2991.csv

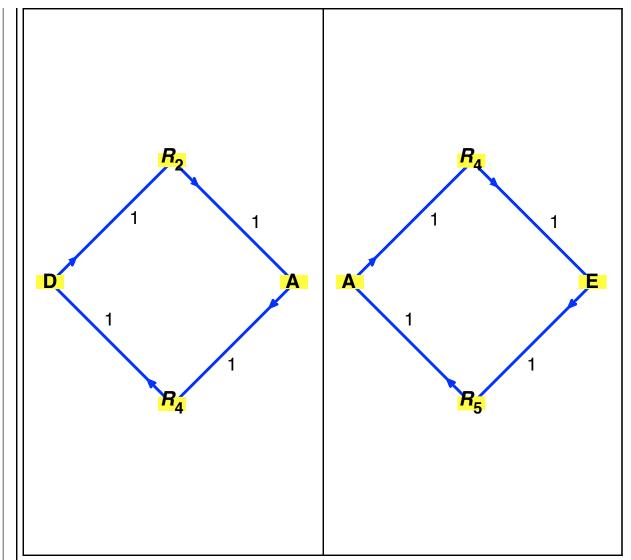


No. 11: injectiveEx1_2993.csv

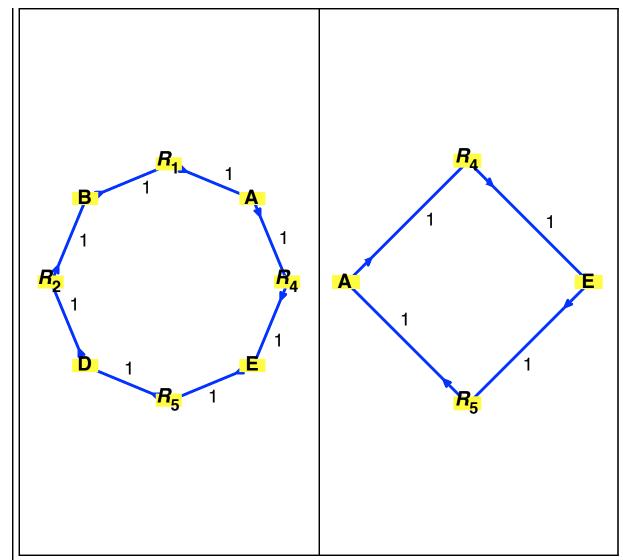


No. 12: injectiveEx1_3103.csv

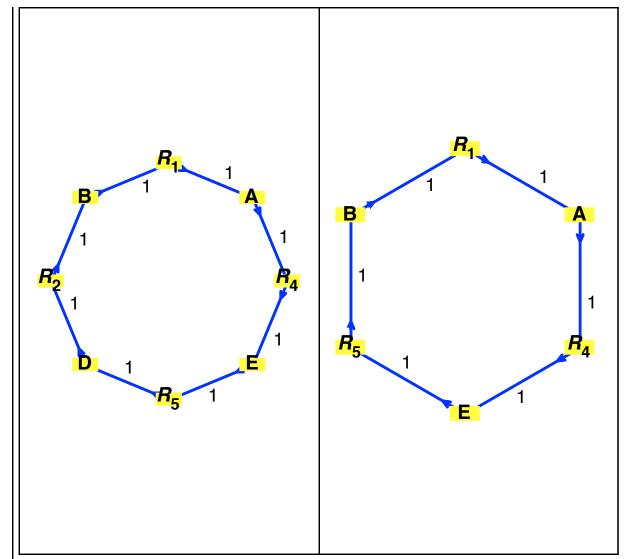




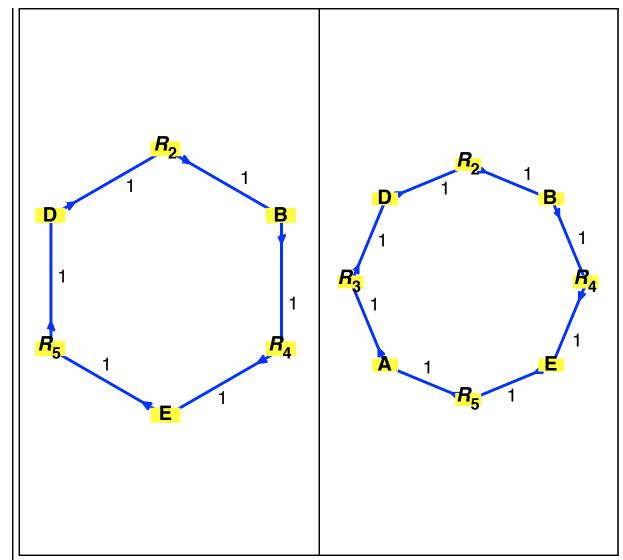
No. 13: injectiveEx3_1250.csv



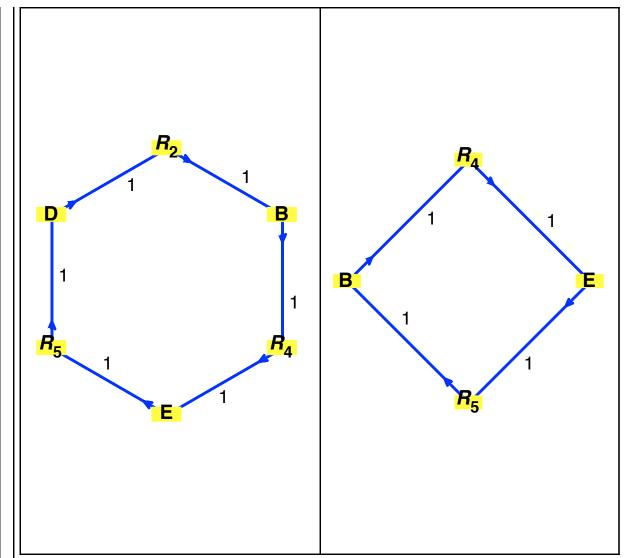
No. 14: injectiveEx3_1252.csv



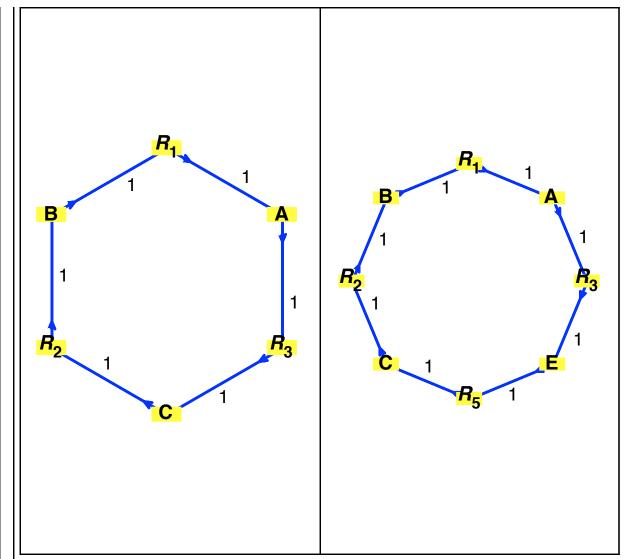
No. 15: injectiveEx3_1268.csv



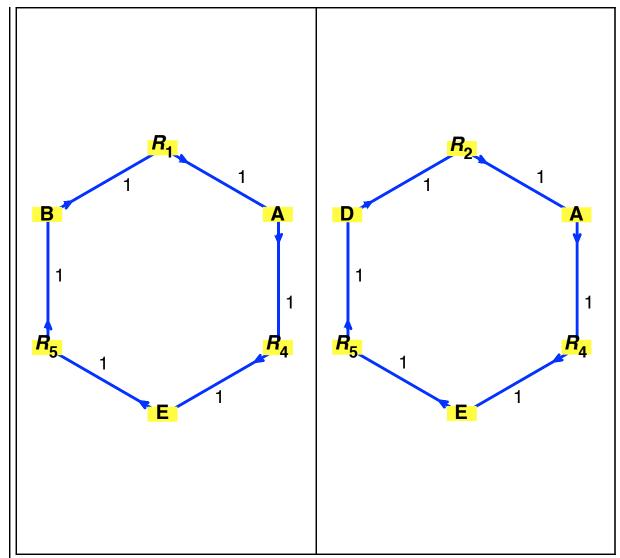
No. 16: injectiveEx3_1270.csv



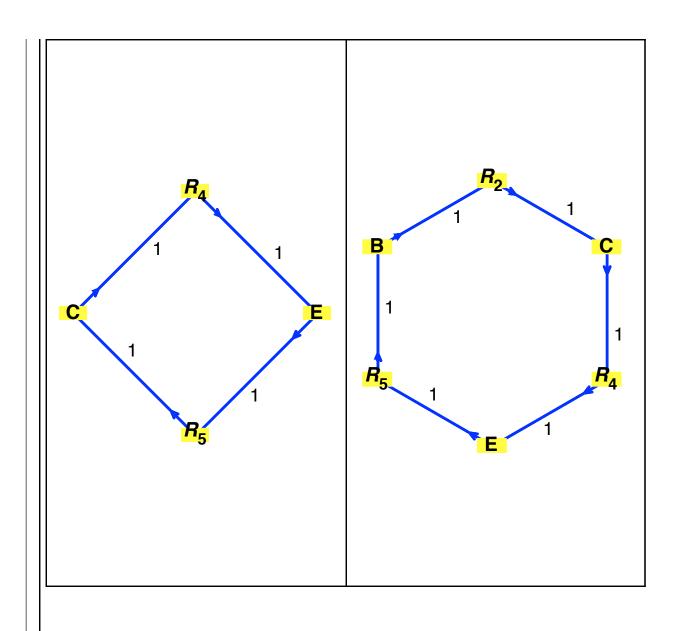
No. 17: injectiveEx3_1302.csv

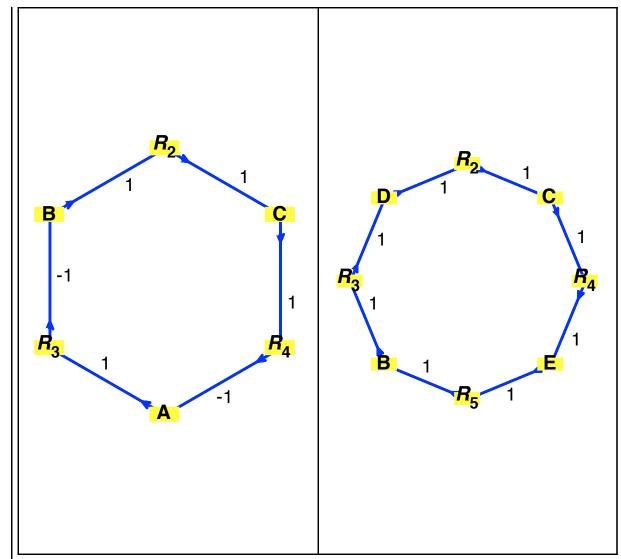


No. 18: injectiveEx3_1369.csv

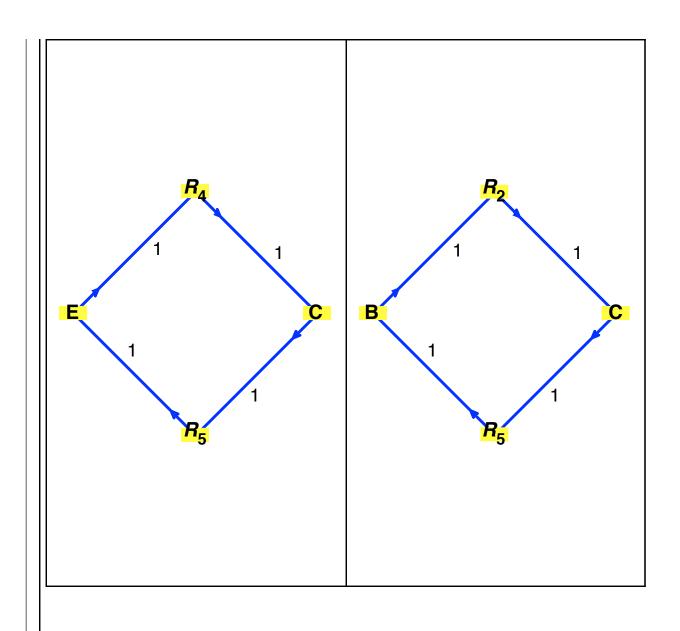


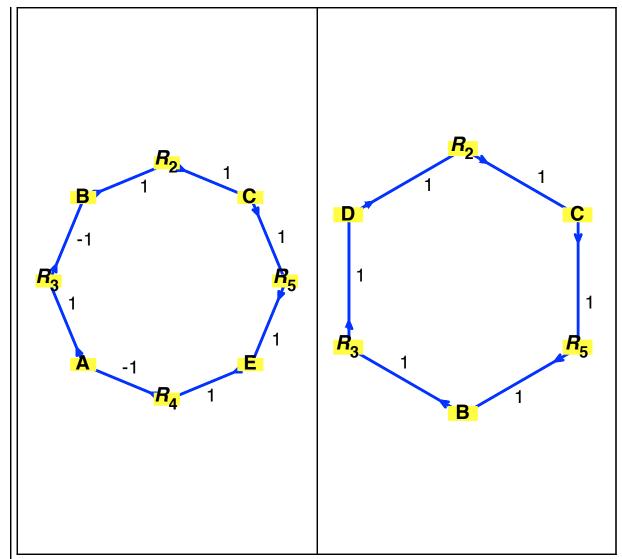
No. 19: injectiveEx3_1473.csv



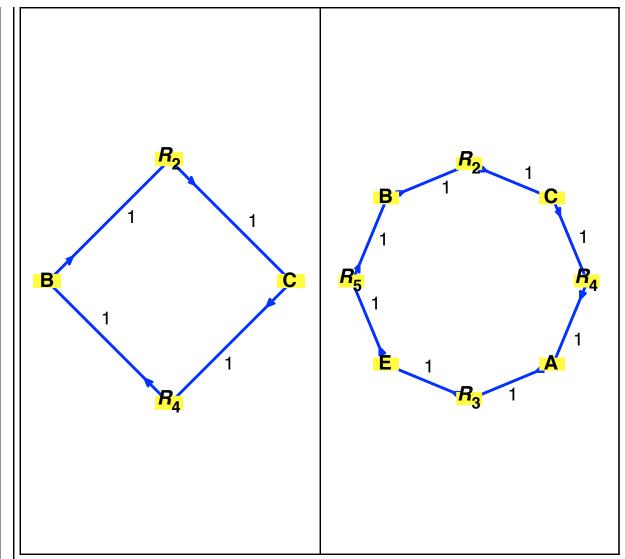


No. 20: injectiveEx3_1474.csv

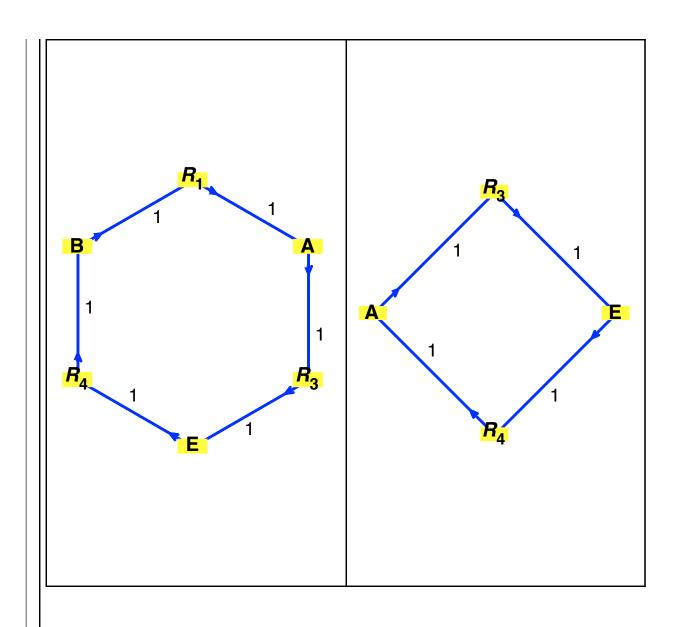


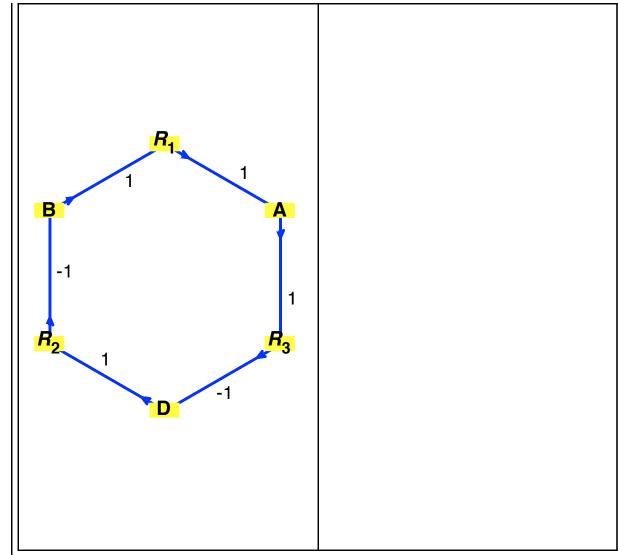


No. 21: injectiveEx3_1516.csv

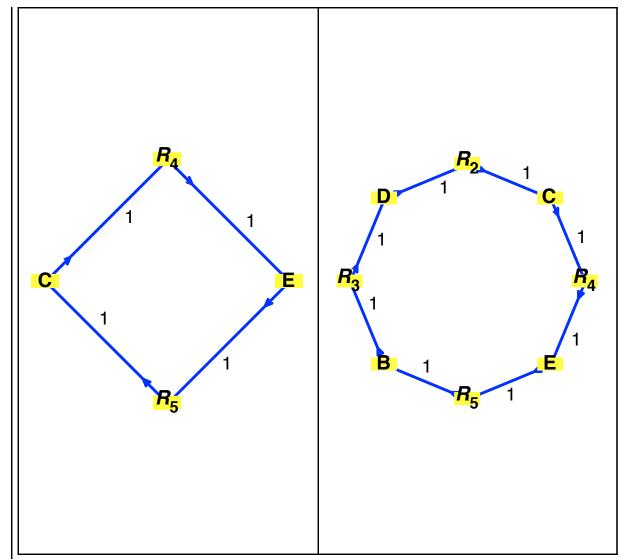


No. 22: injectiveEx3_1518.csv

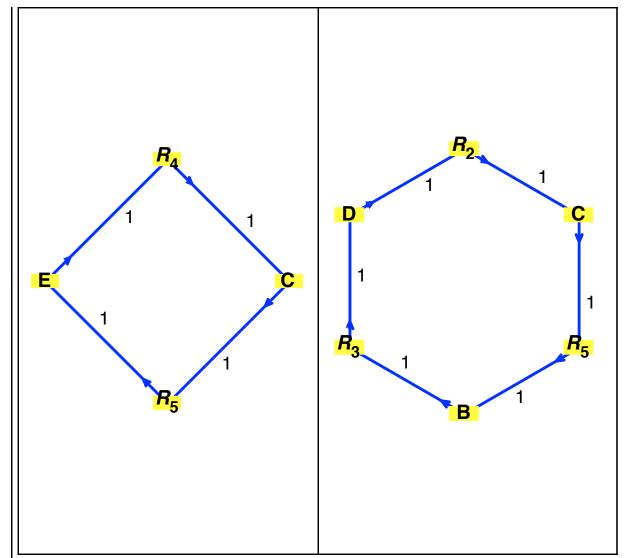




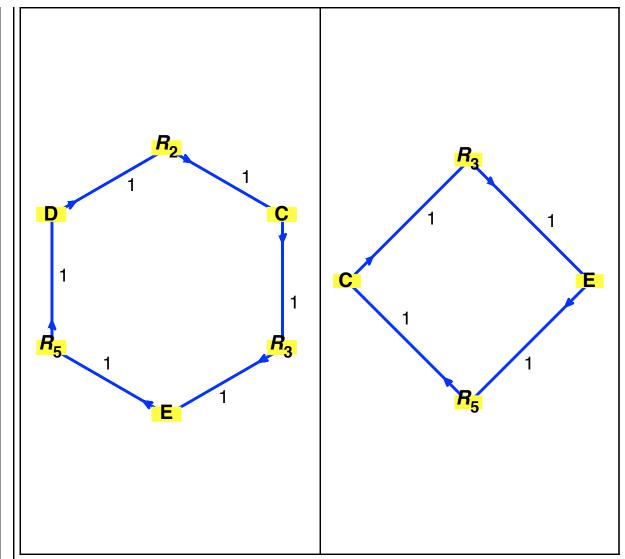
No. 23: injectiveEx3_1721.csv



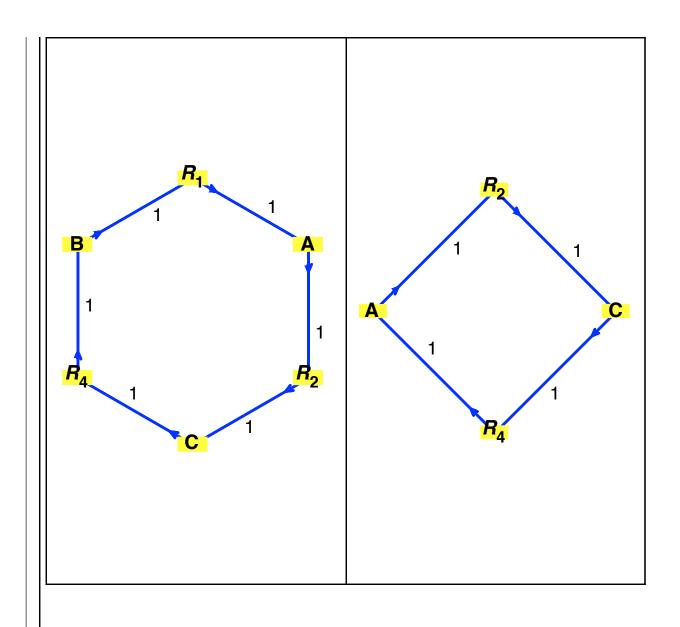
No. 24: injectiveEx3_1722.csv

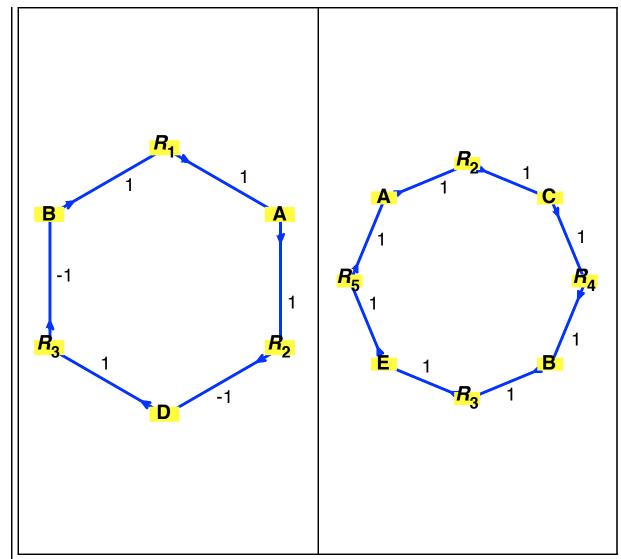


No. 25: injectiveEx3_1743.csv

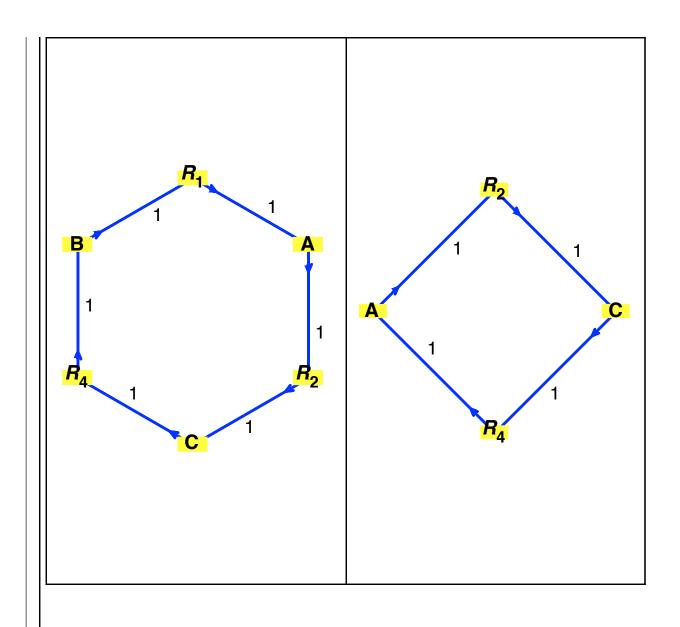


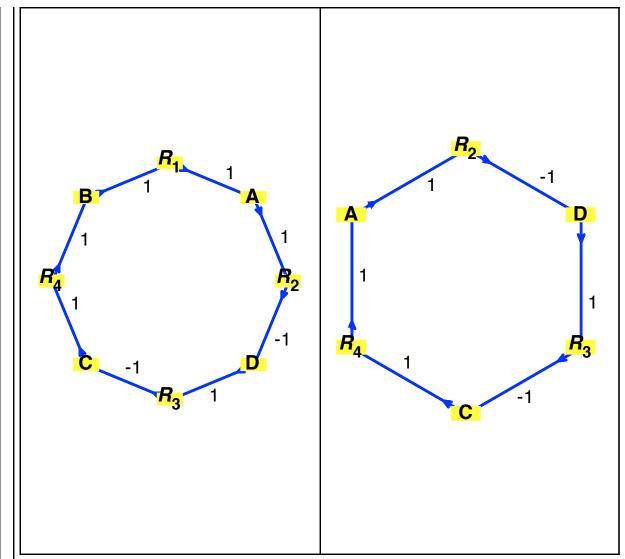
No. 26: injectiveEx3_1796.csv



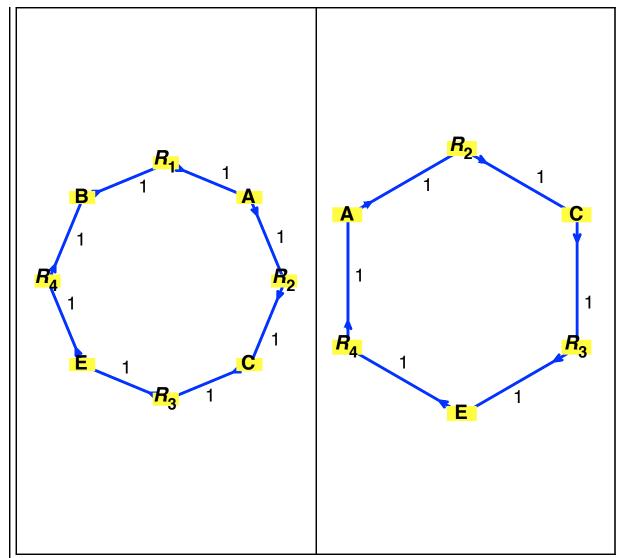


No. 27: injectiveEx3_1814.csv

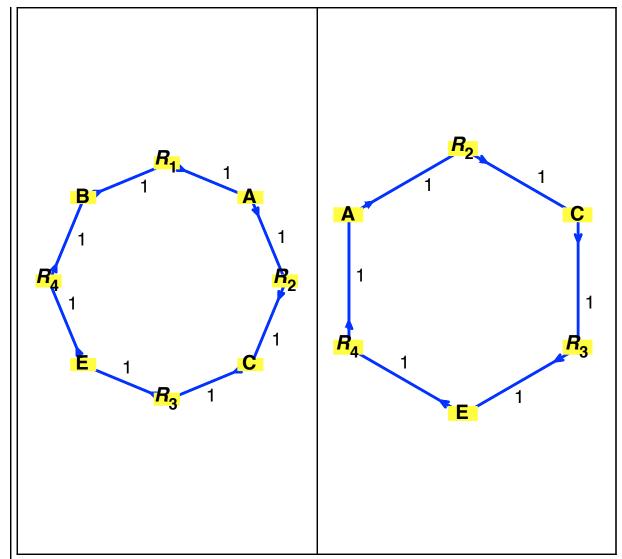




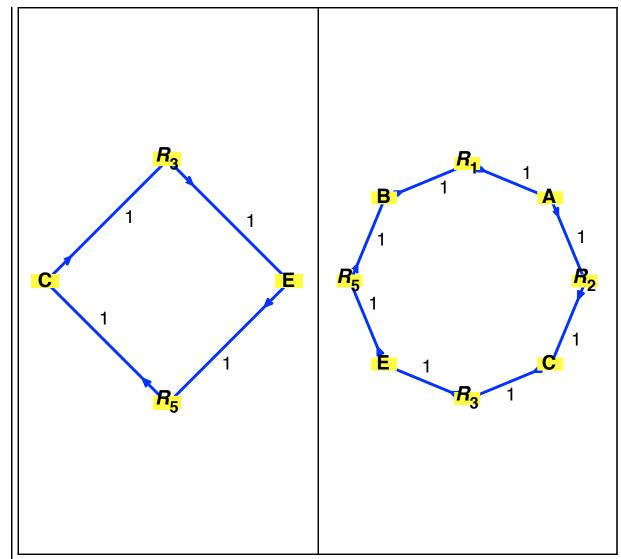
No. 28: injectiveEx3_1816.csv



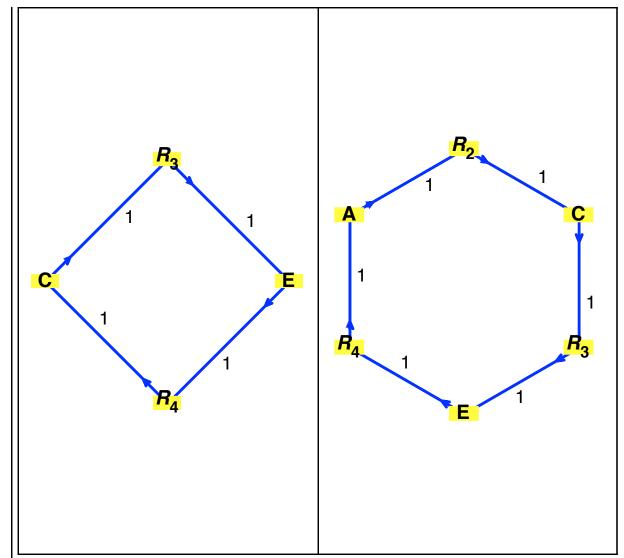
No. 29: injectiveEx3_1817.csv



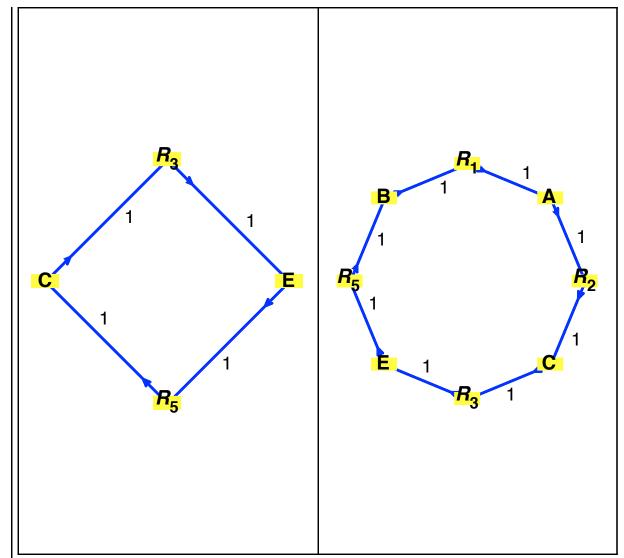
No. 30: injectiveEx3_1819.csv



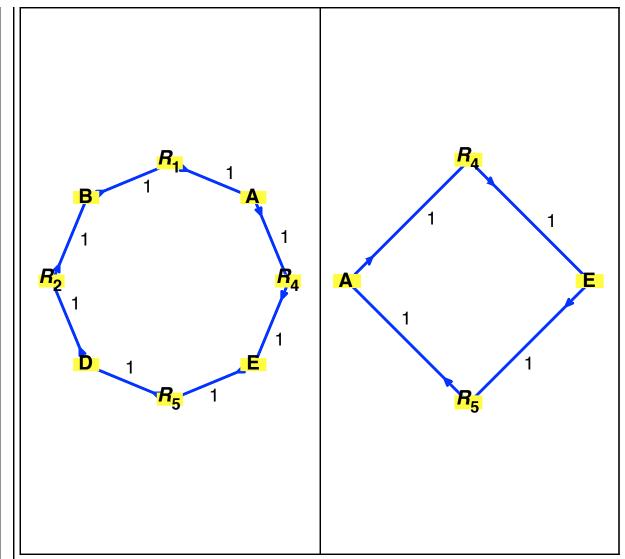
No. 31: injectiveEx3_1820.csv



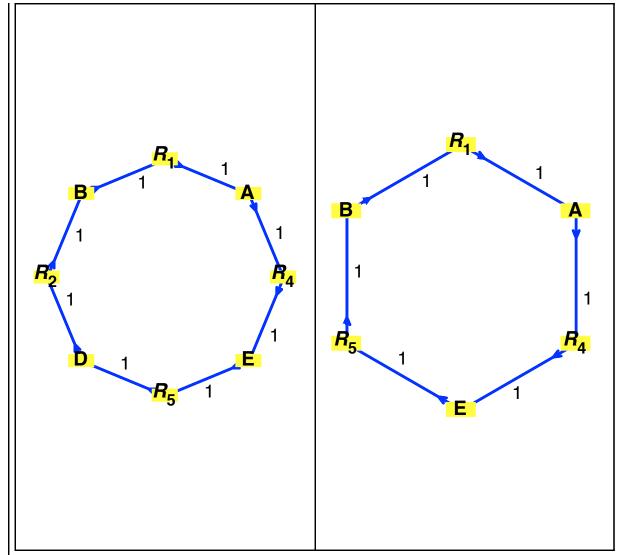
No. 32: injectiveEx3_1821.csv



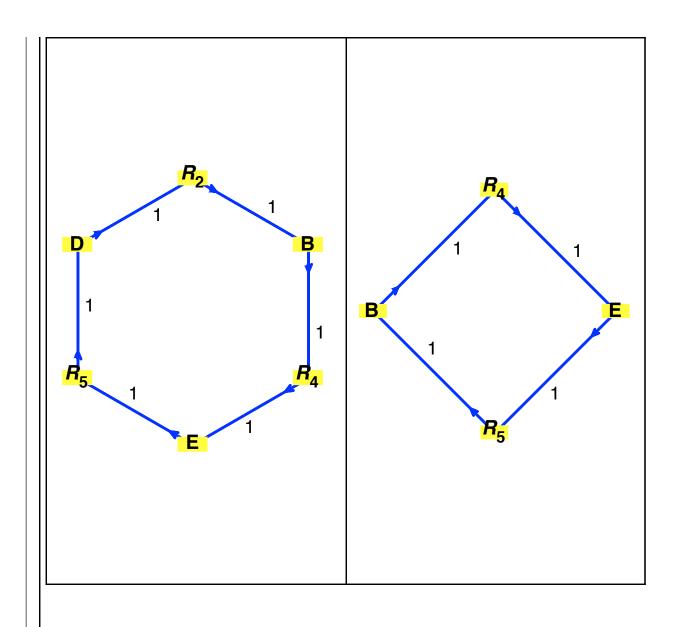
No. 33: injectiveEx3_2945.csv

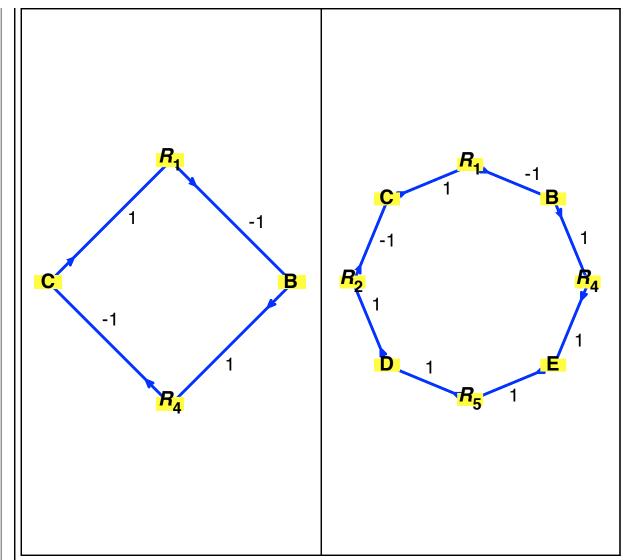


No. 34: injectiveEx3_2946.csv

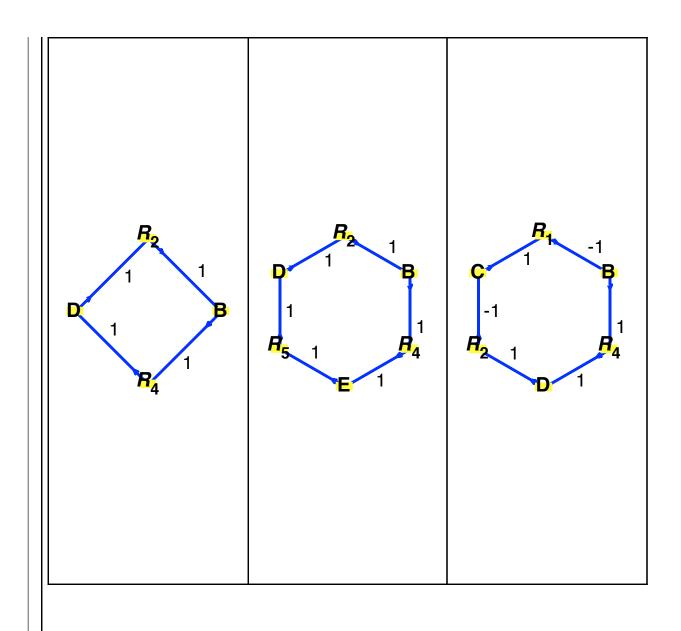


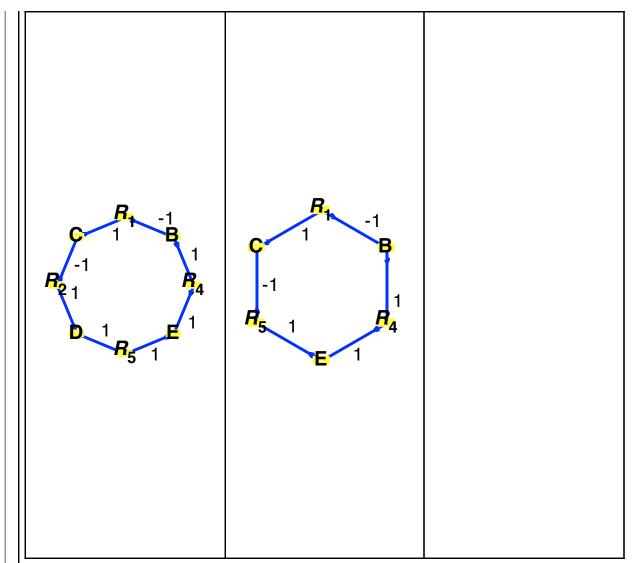
No. 35: injectiveEx3_2954.csv



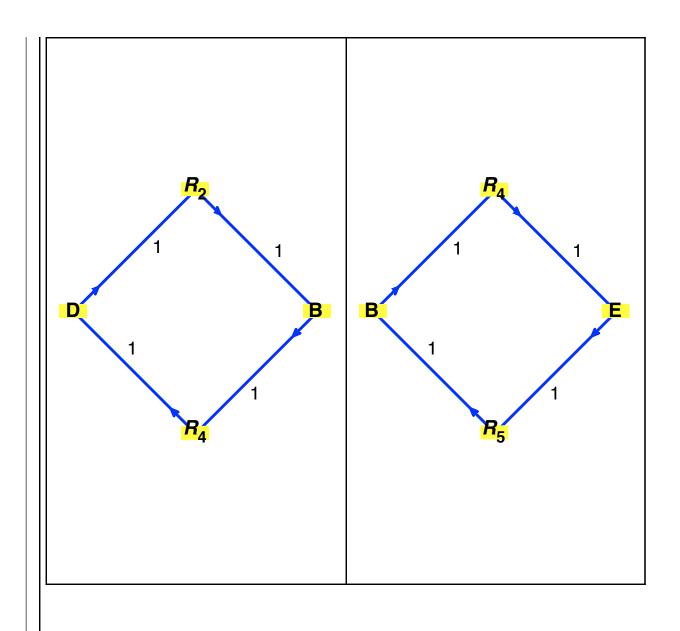


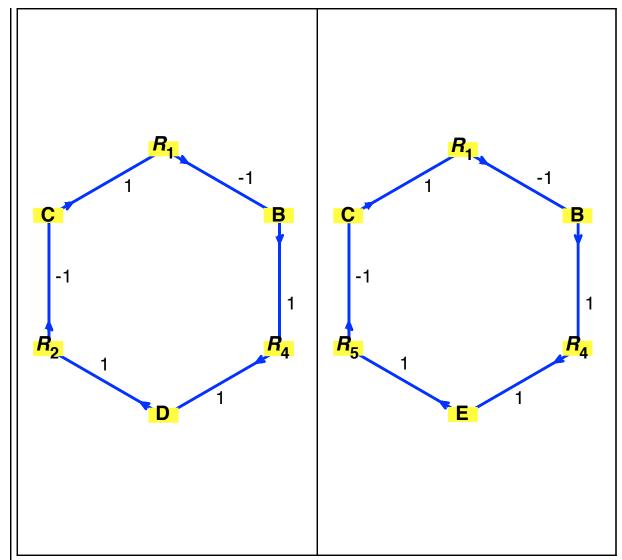
No. 36: injectiveEx3_2955.csv



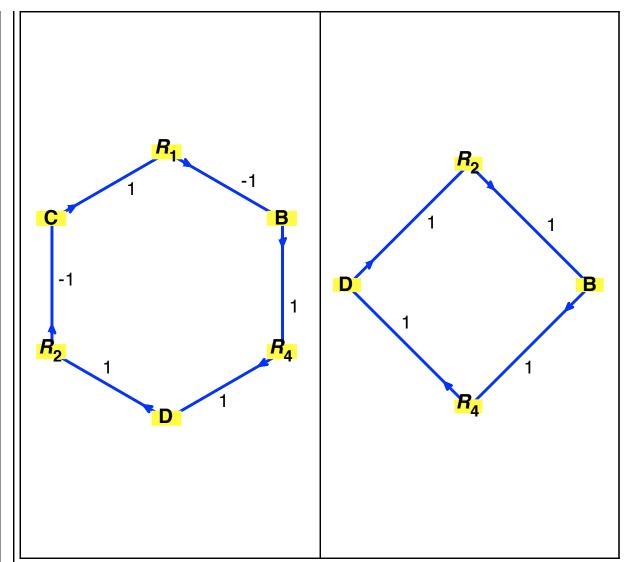


No. 37: injectiveEx3_2957.csv

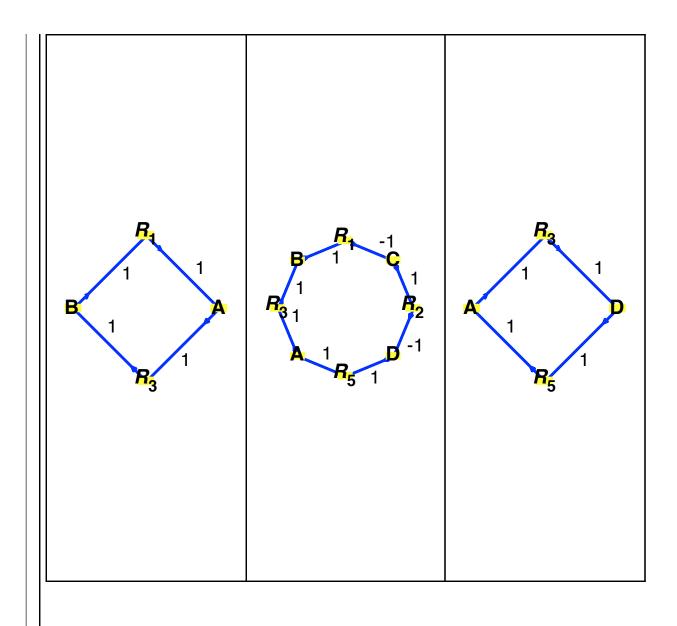


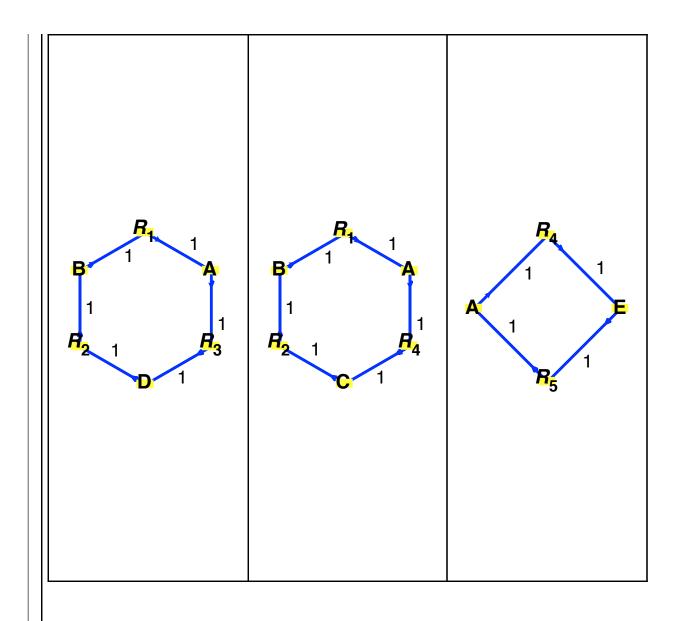


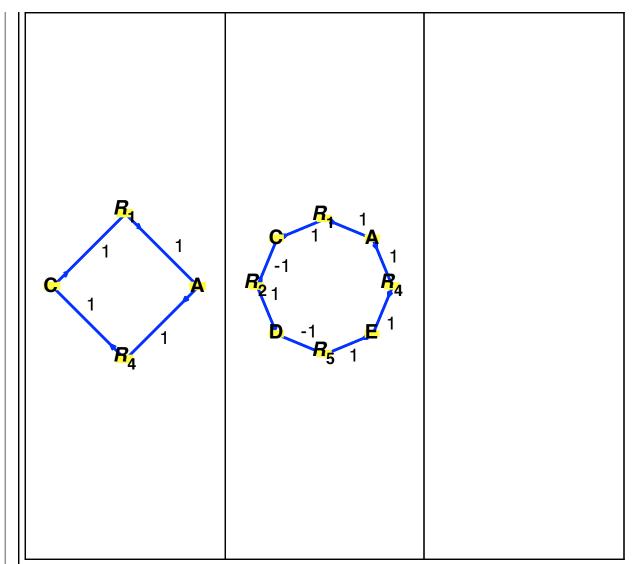
No. 38: injectiveEx3_2958.csv



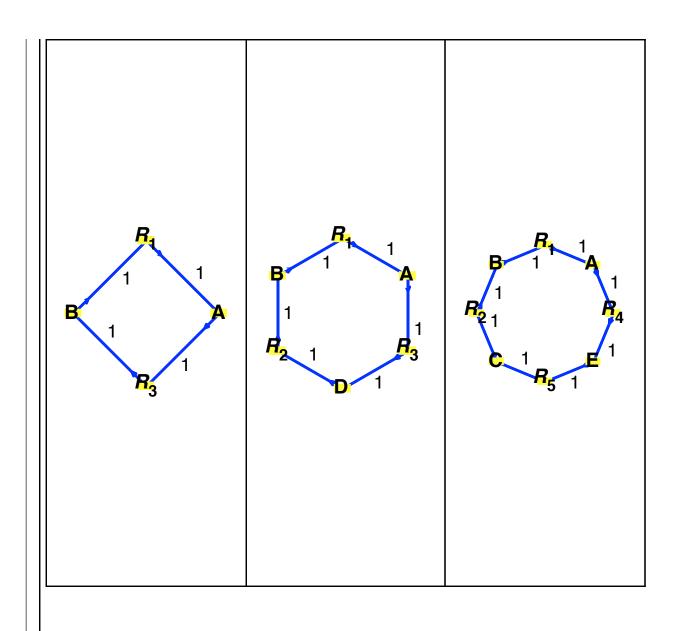
No. 39: injectiveEx3_2965.csv

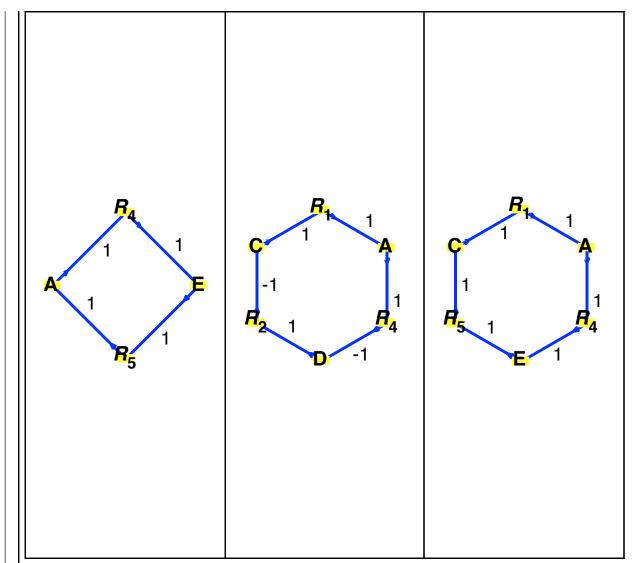




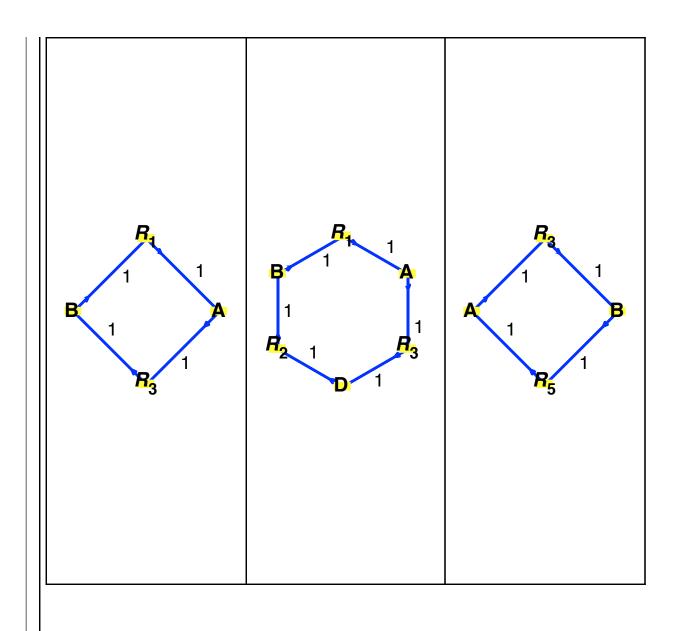


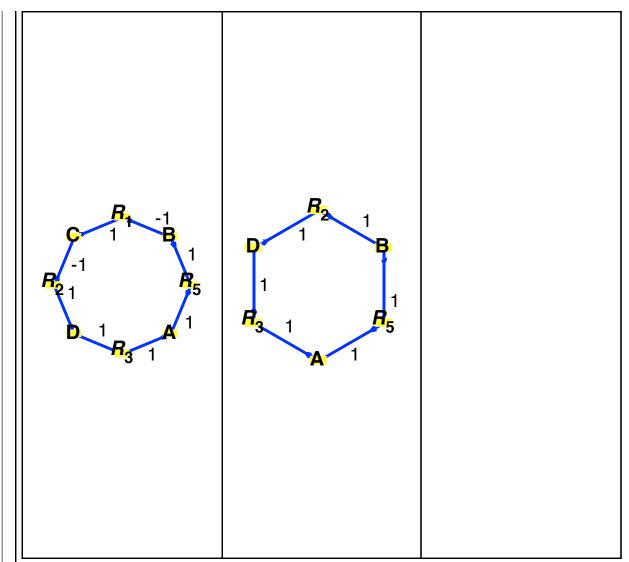
No. 40: injectiveEx3_2969.csv



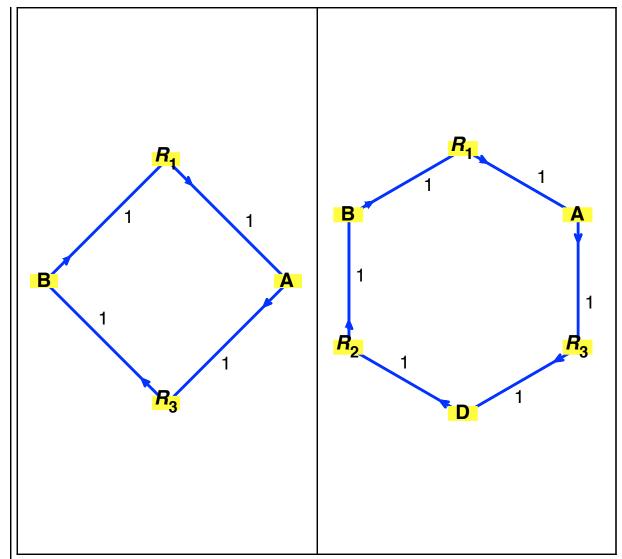


No. 41: injectiveEx3_2970.csv

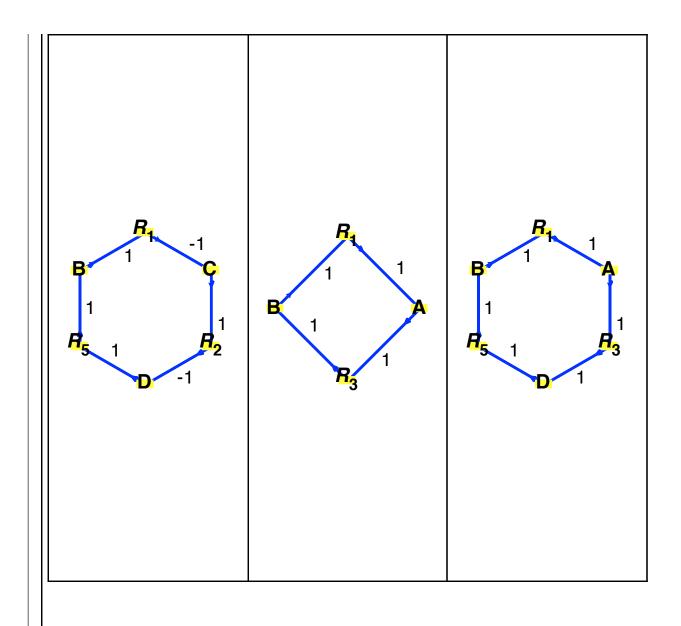


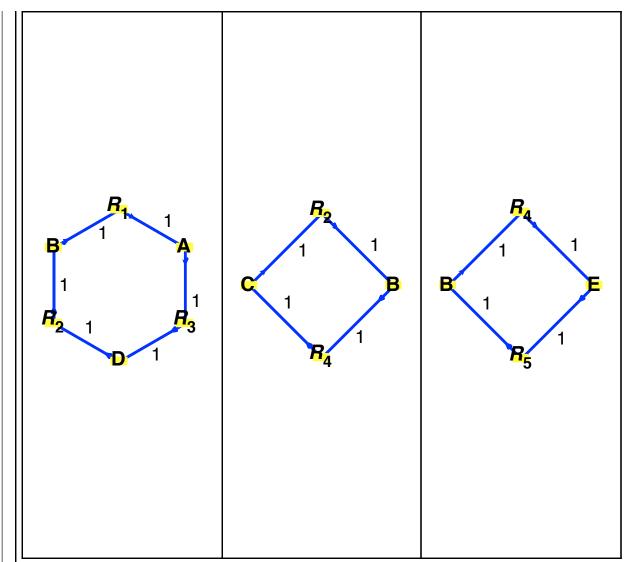


No. 42: injectiveEx3_2971.csv

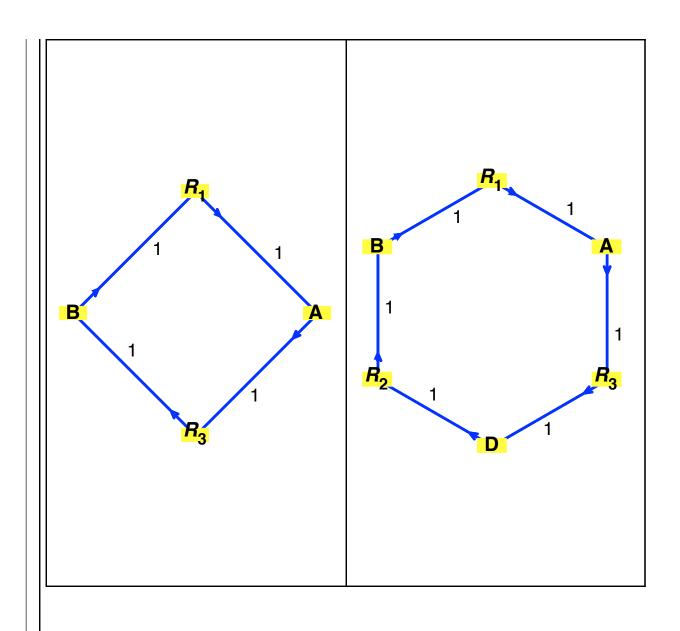


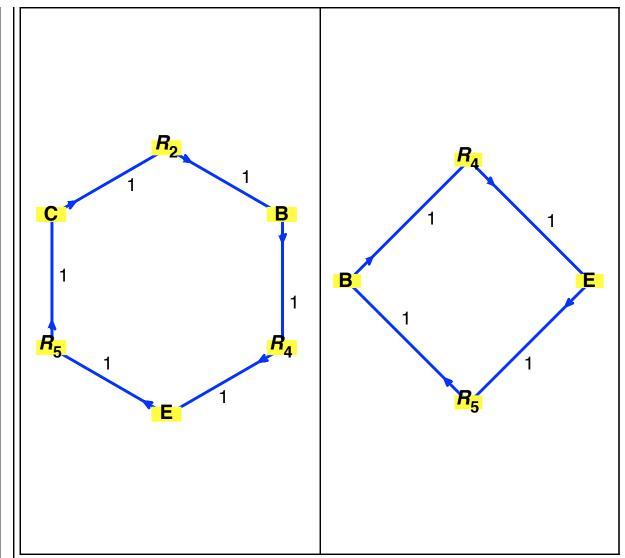
No. 43: injectiveEx3_2972.csv



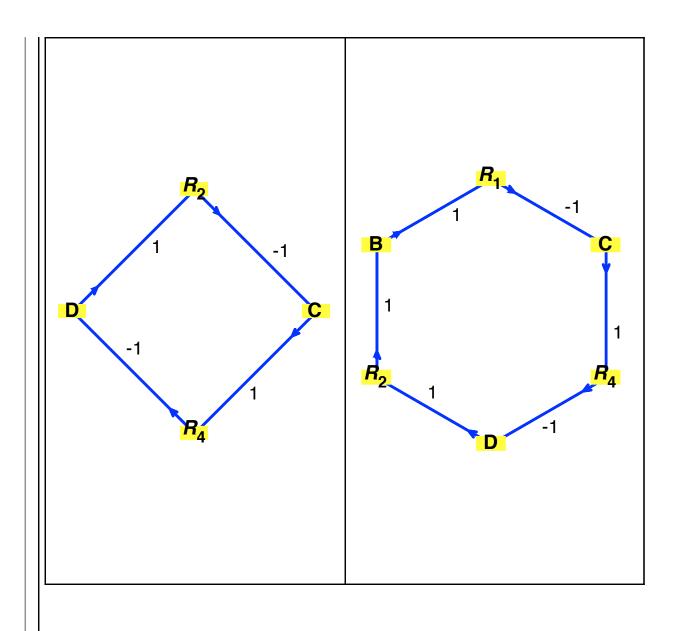


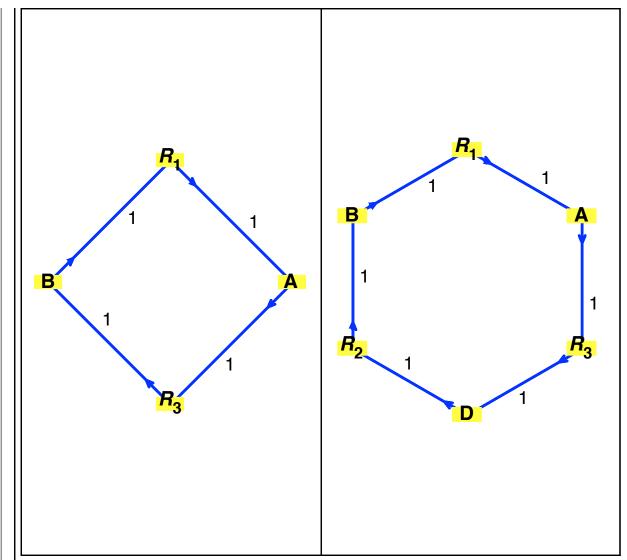
No. 44: injectiveEx3_2974.csv



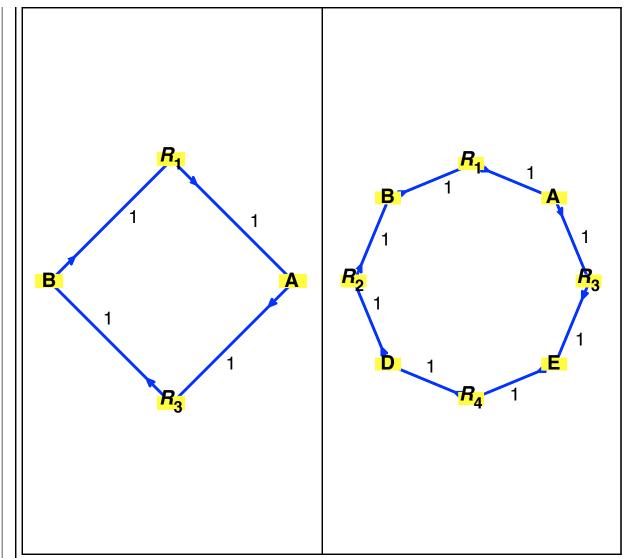


No. 45: injectiveEx3_2975.csv

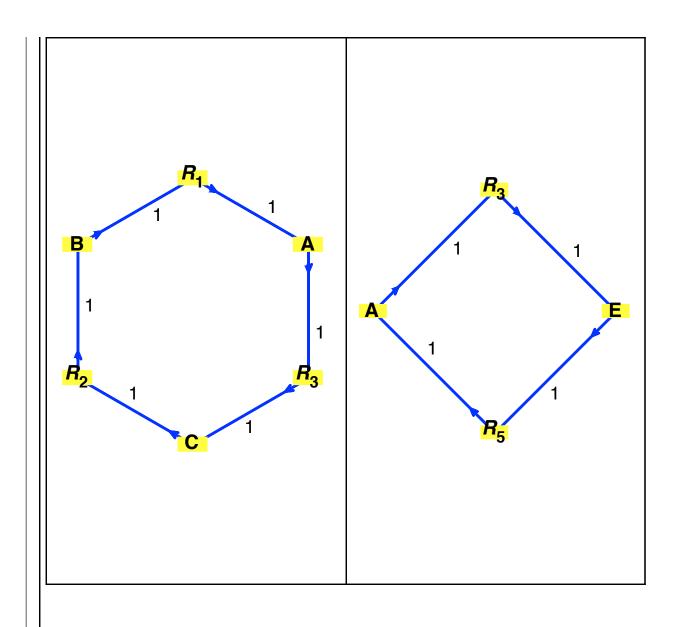


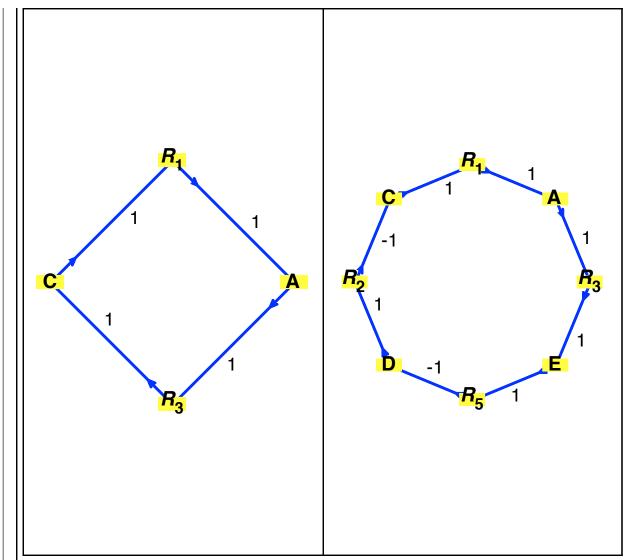


No. 46: injectiveEx3_2980.csv

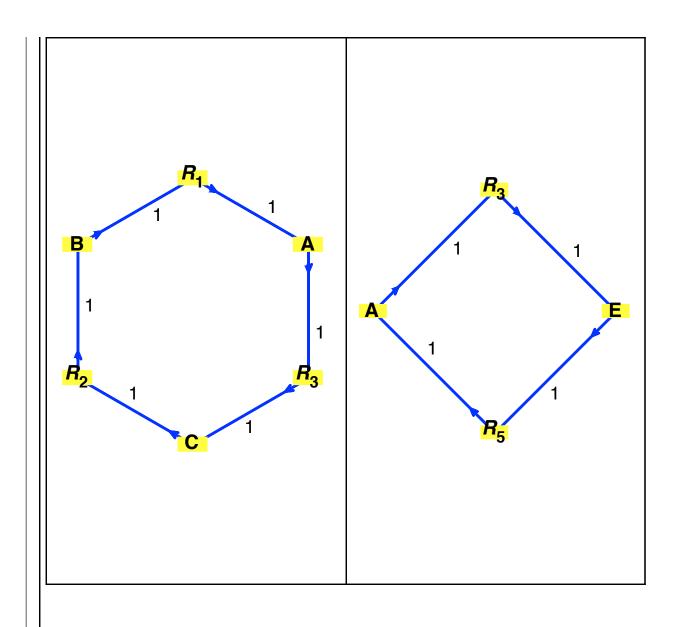


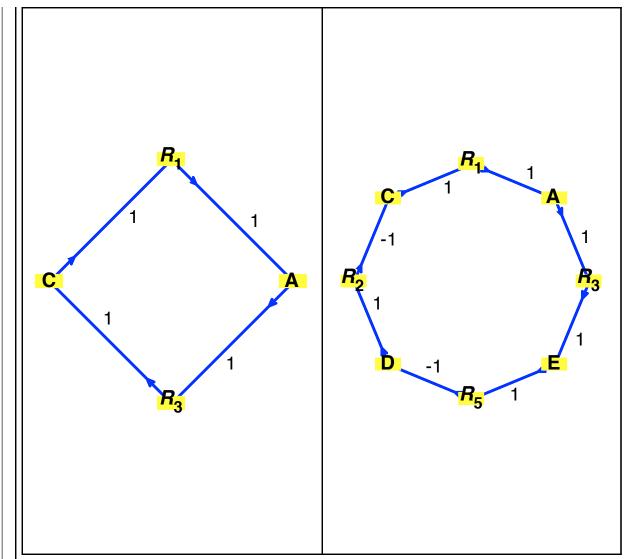
No. 47: injectiveEx3_2983.csv



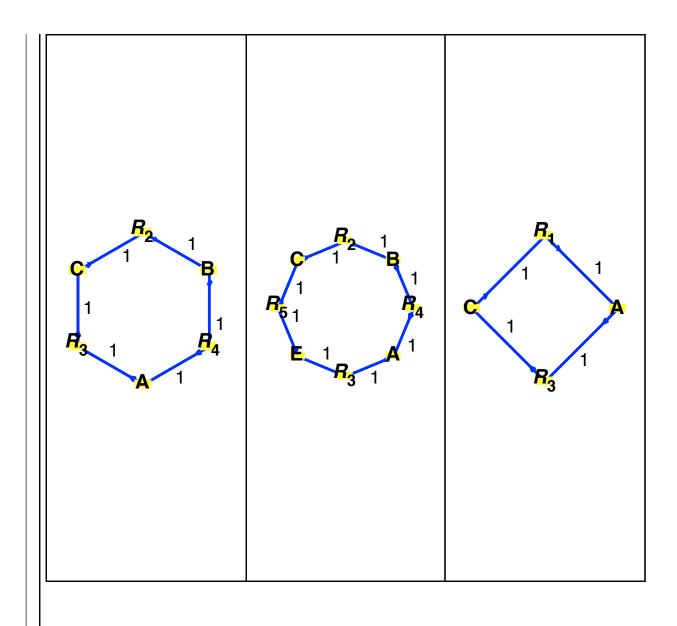


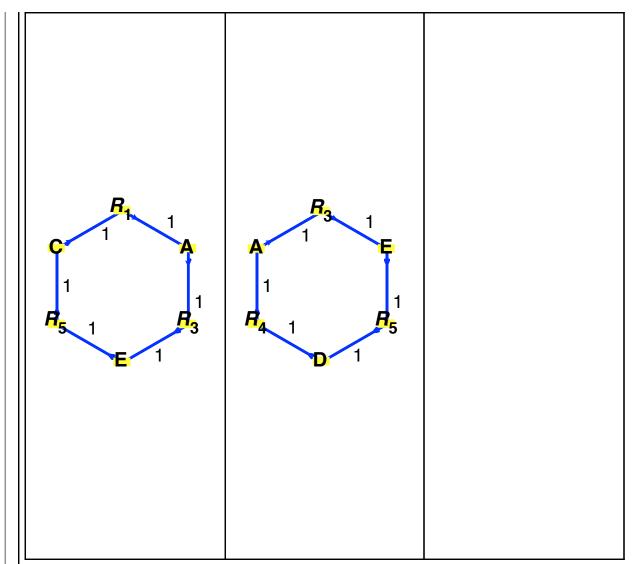
No. 48: injectiveEx3_2986.csv



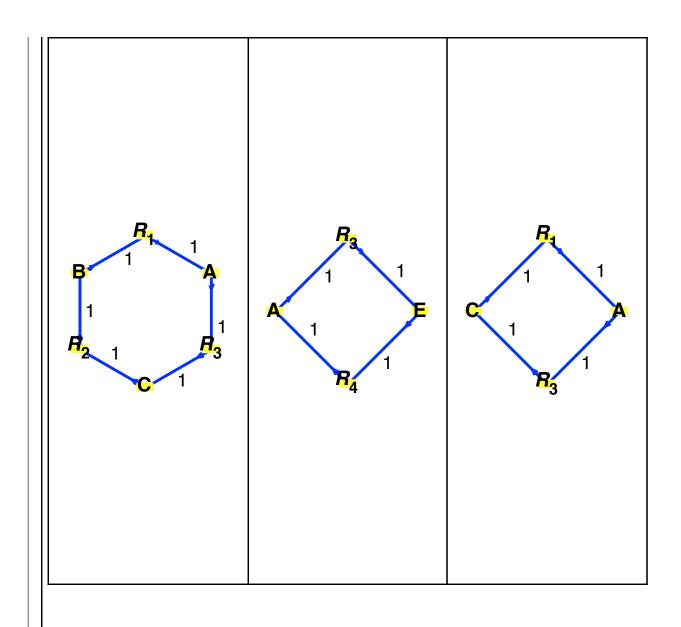


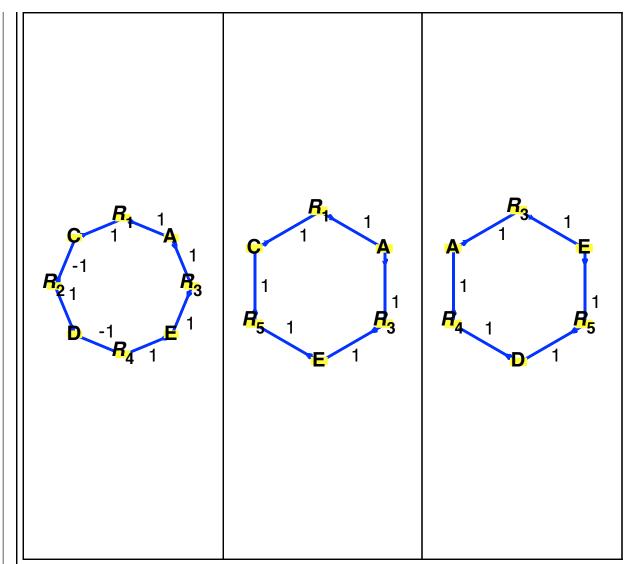
No. 49: injectiveEx3_2990.csv



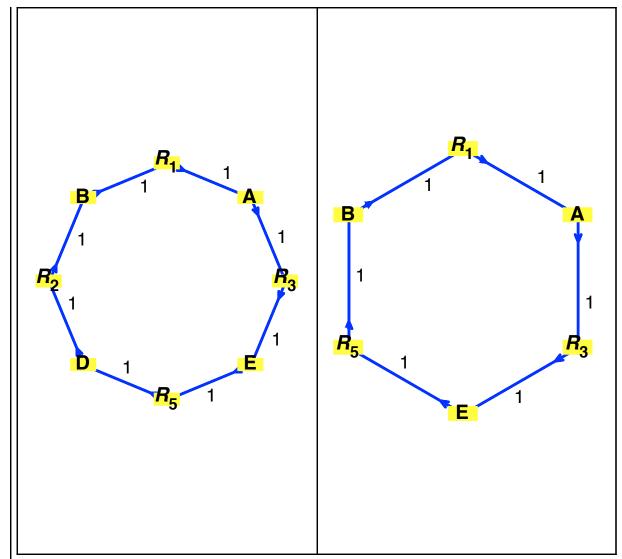


No. 50: injectiveEx3_2992.csv

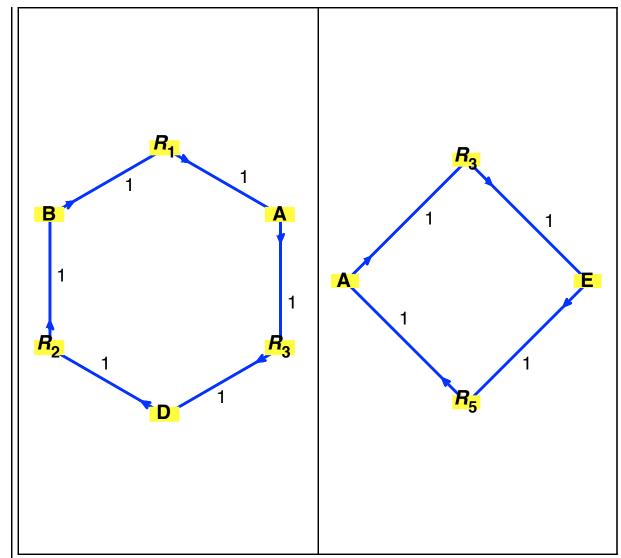




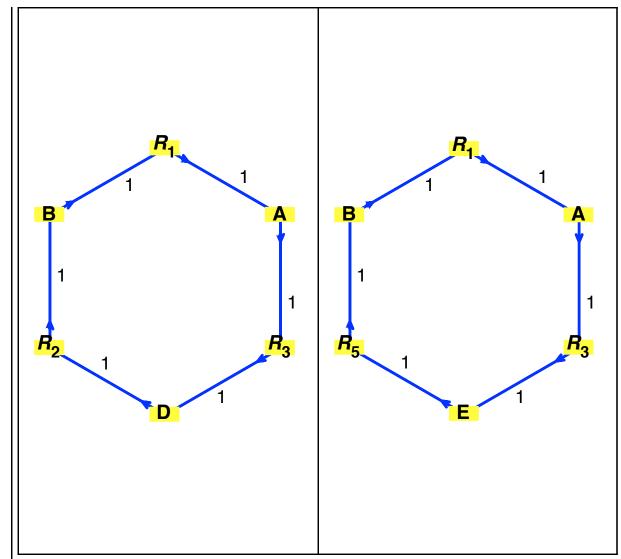
No. 51: injectiveEx3_2996.csv



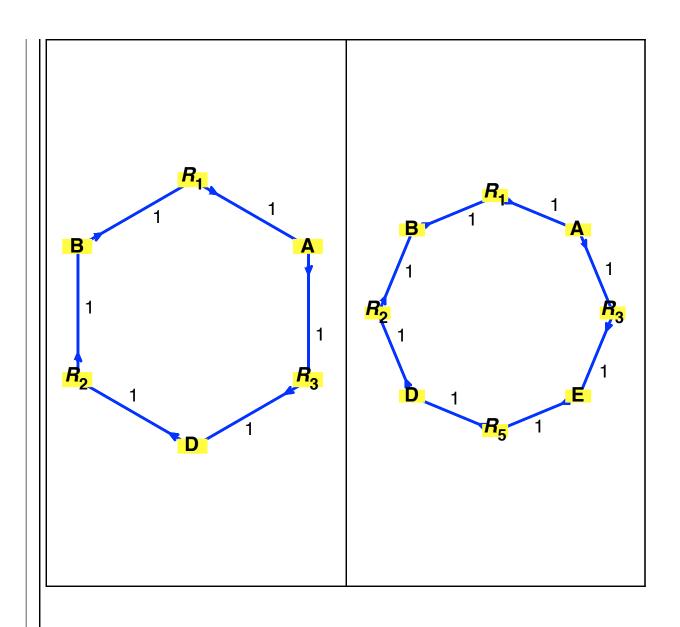
No. 52: injectiveEx3_3000.csv

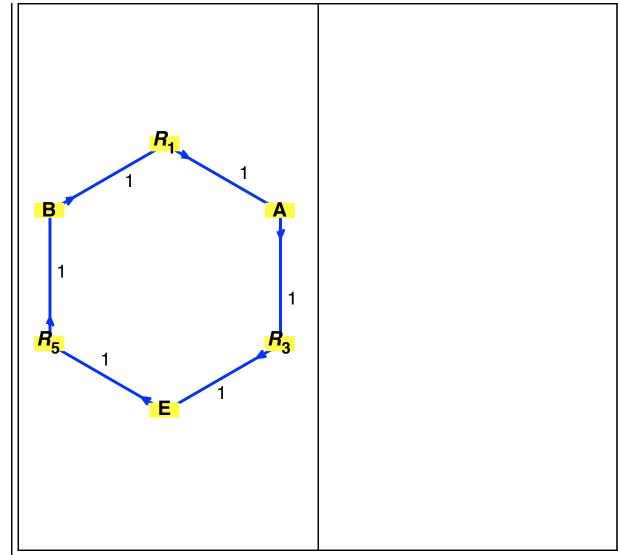


No. 53: injectiveEx3_3002.csv

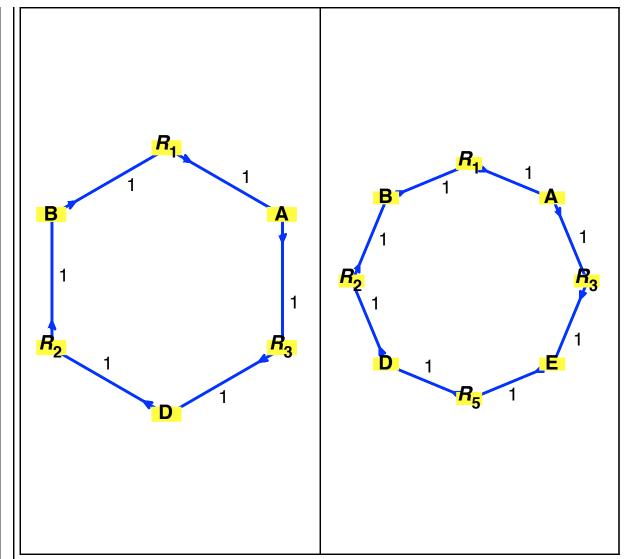


No. 54: injectiveEx3_3003.csv

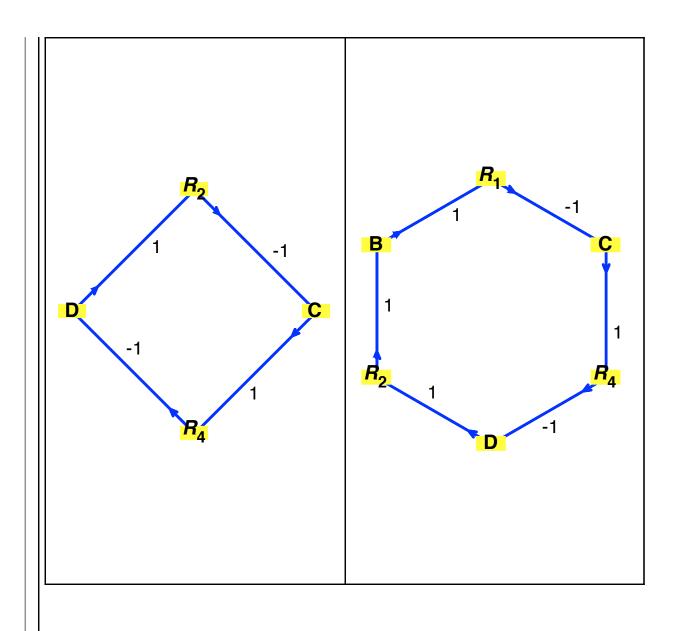


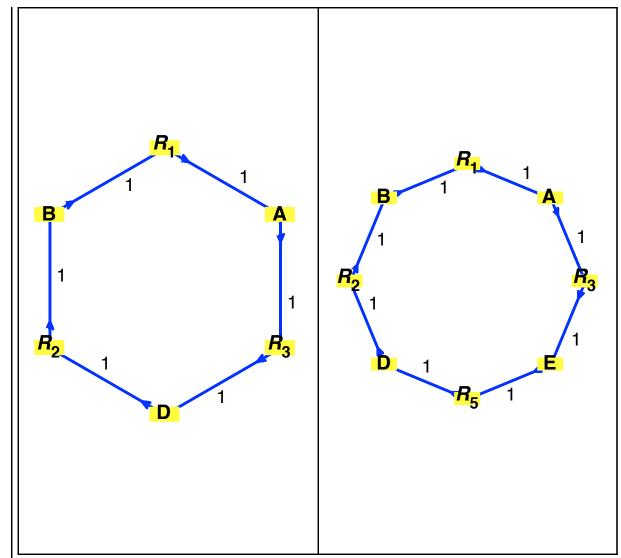


No. 55: injectiveEx3_3004.csv

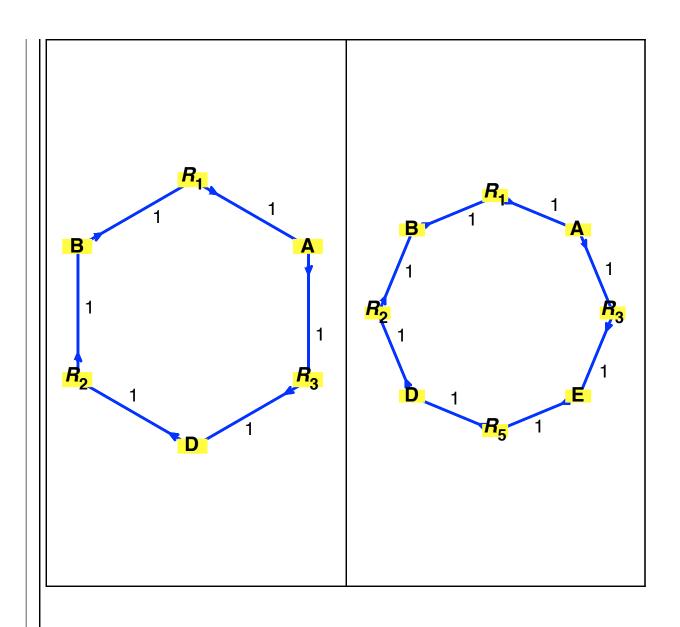


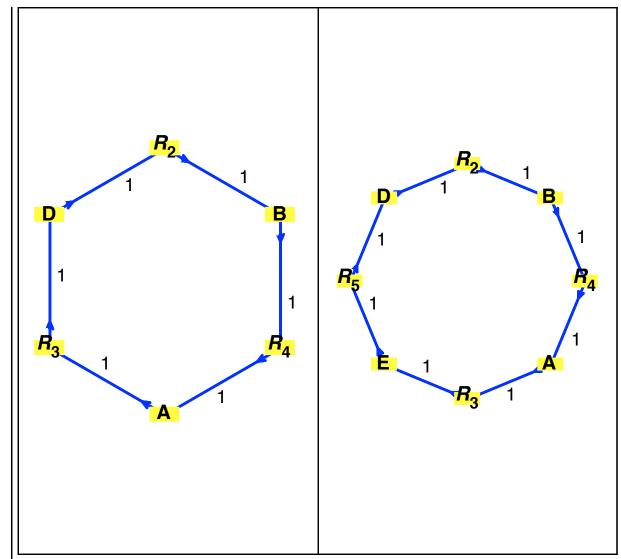
No. 56: injectiveEx3_3007.csv



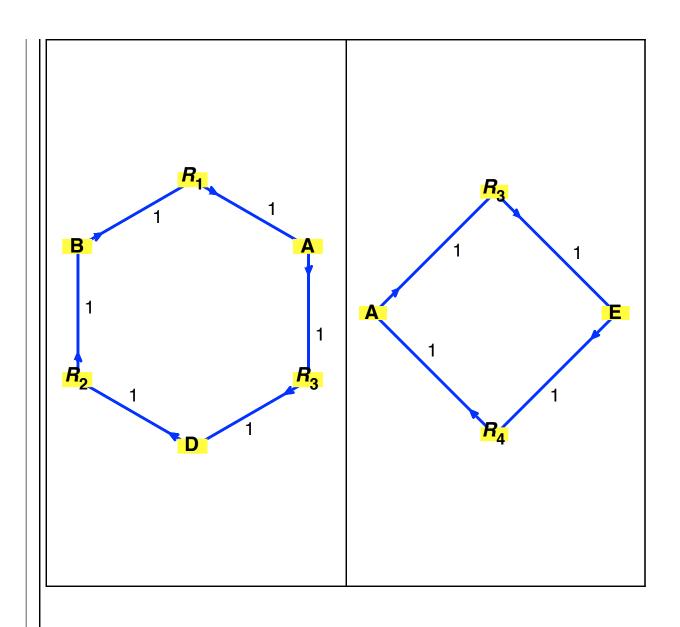


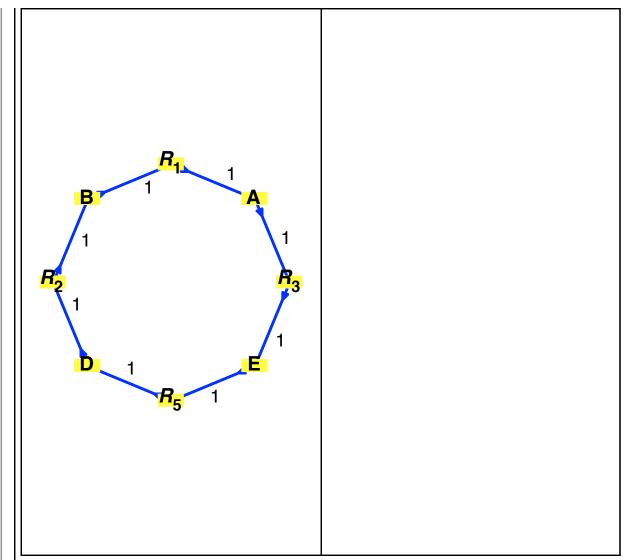
No. 57: injectiveEx3_3009.csv



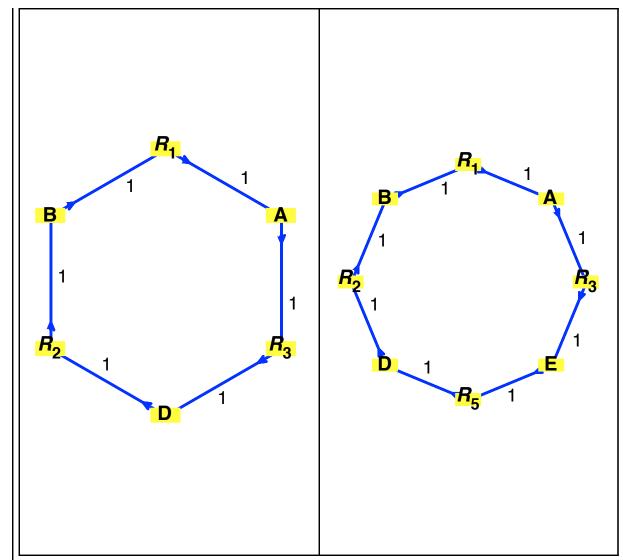


No. 58: injectiveEx3_3011.csv

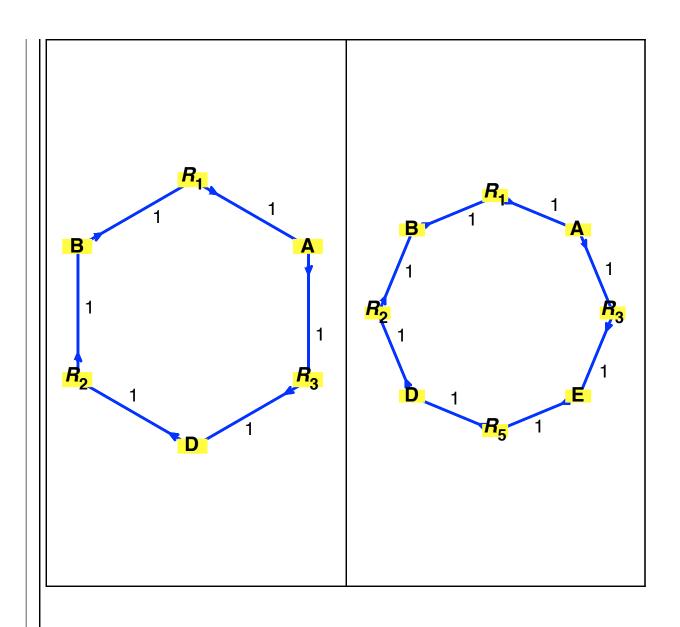


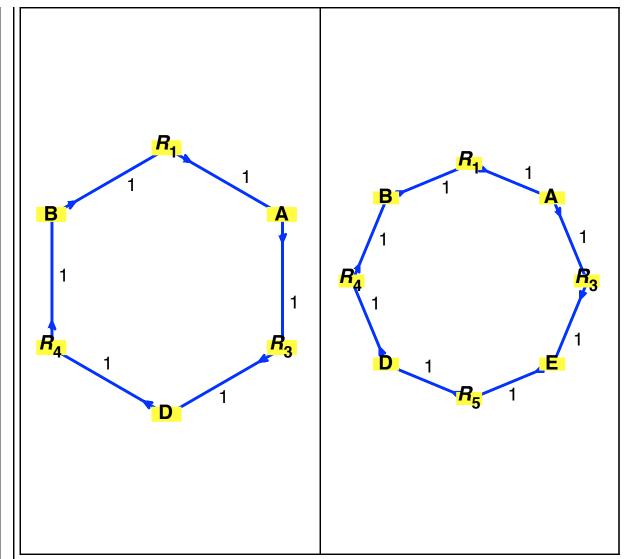


No. 59: injectiveEx3_3012.csv

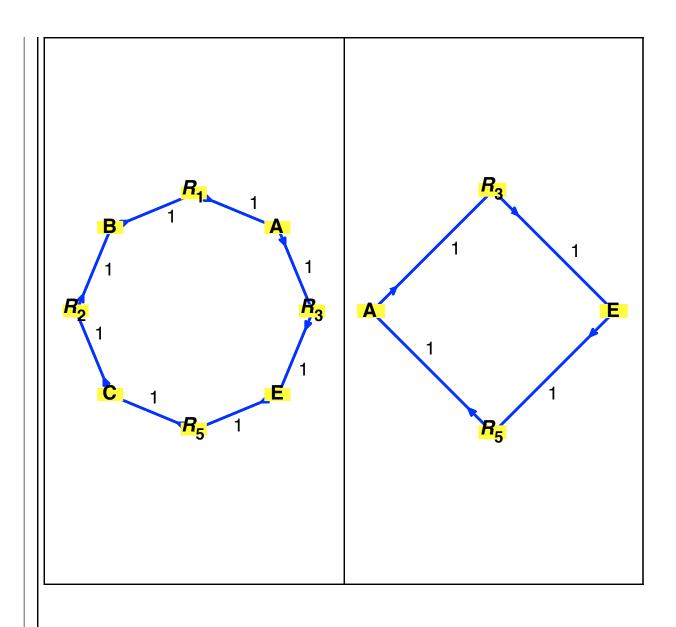


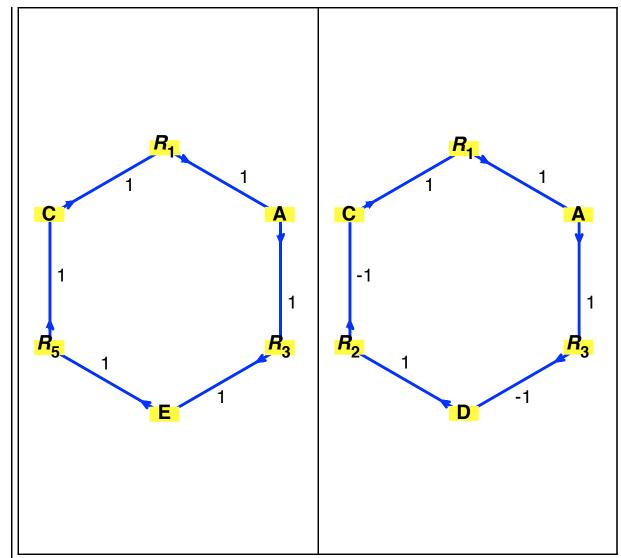
No. 60: injectiveEx3_3014.csv



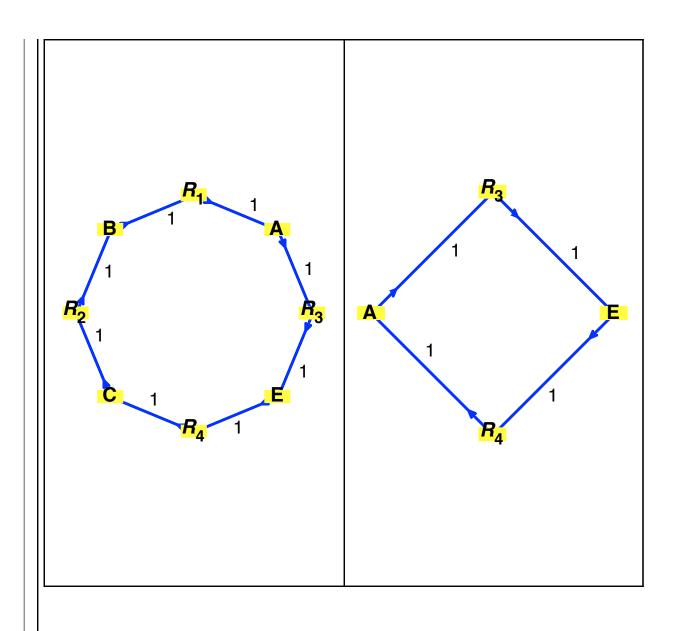


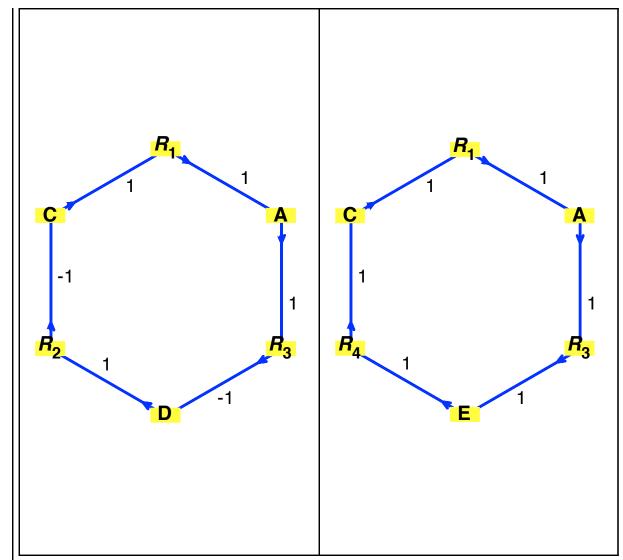
No. 61: injectiveEx3_3015.csv



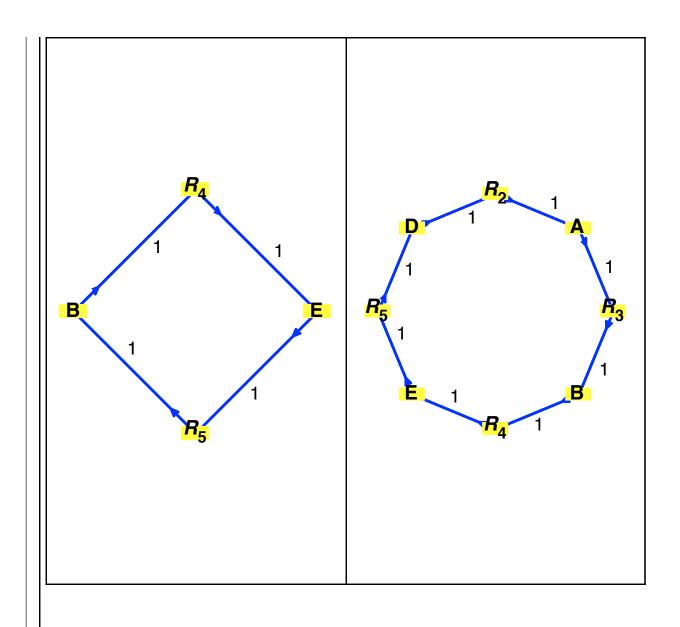


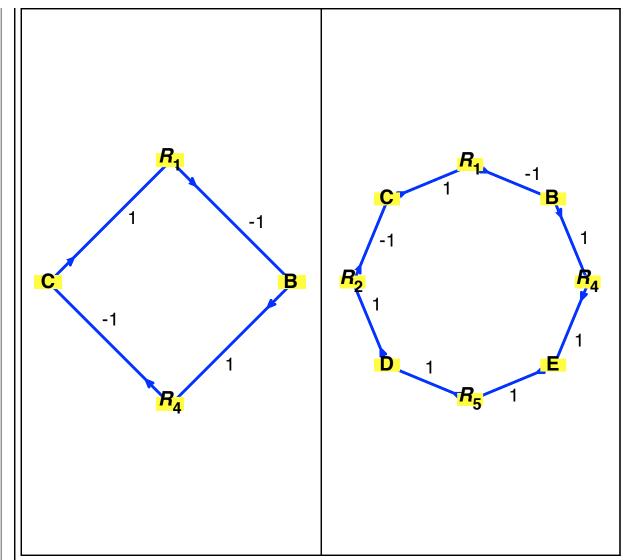
No. 62: injectiveEx3_3017.csv



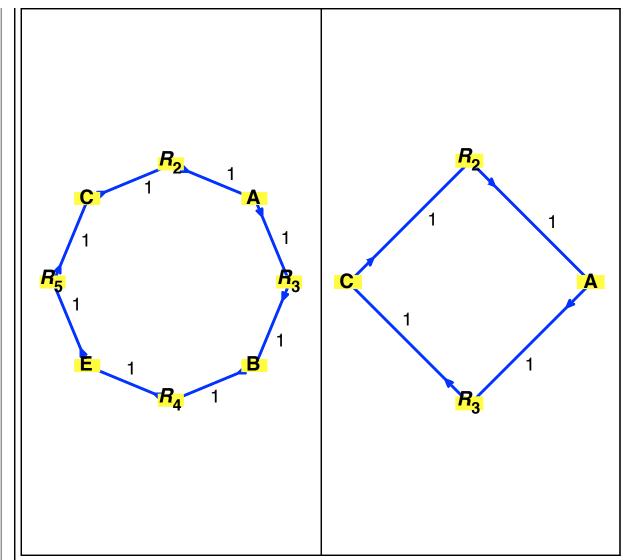


No. 63: injectiveEx3_3110.csv

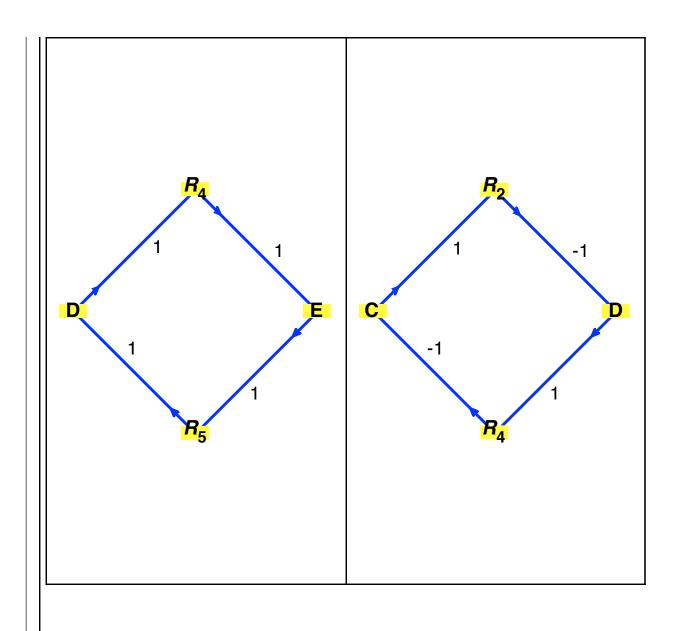


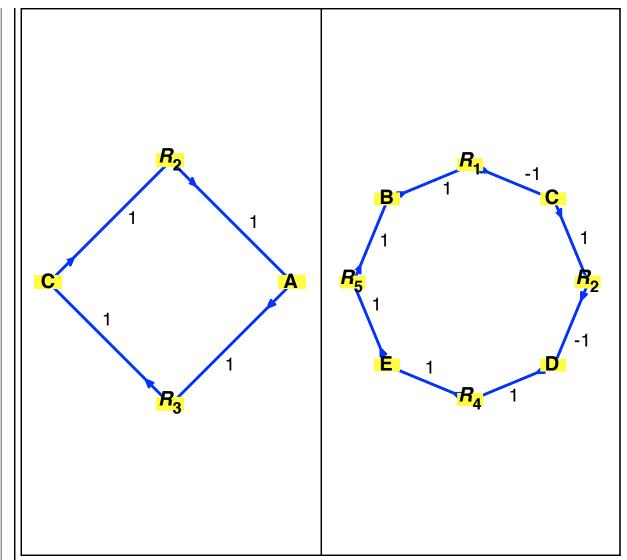


No. 64: injectiveEx3_3114.csv

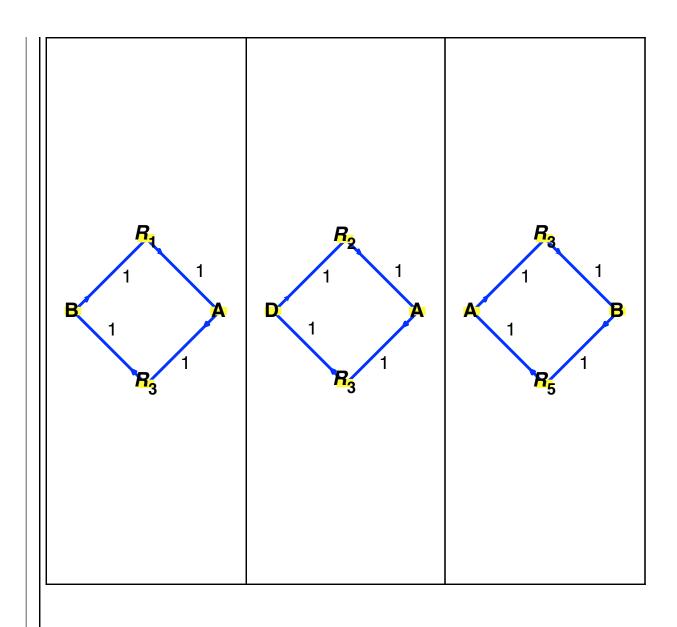


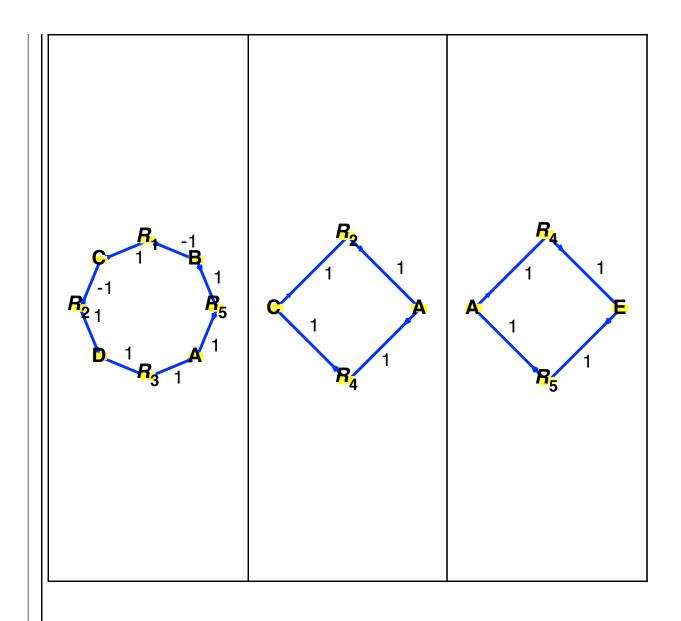
No. 65: injectiveEx3_3118.csv

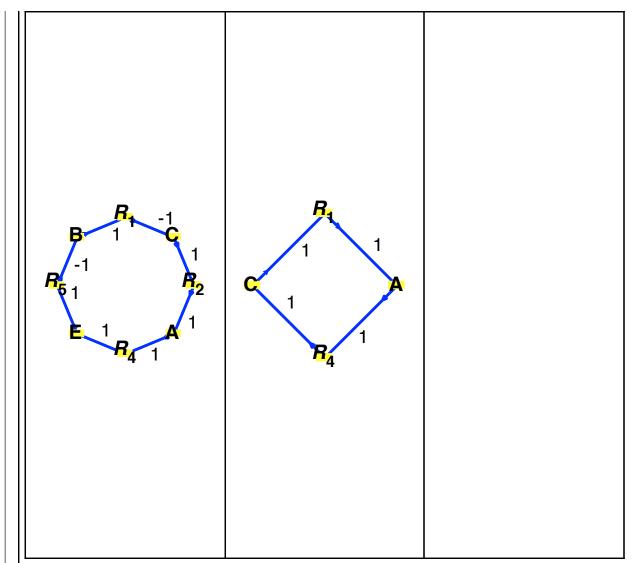




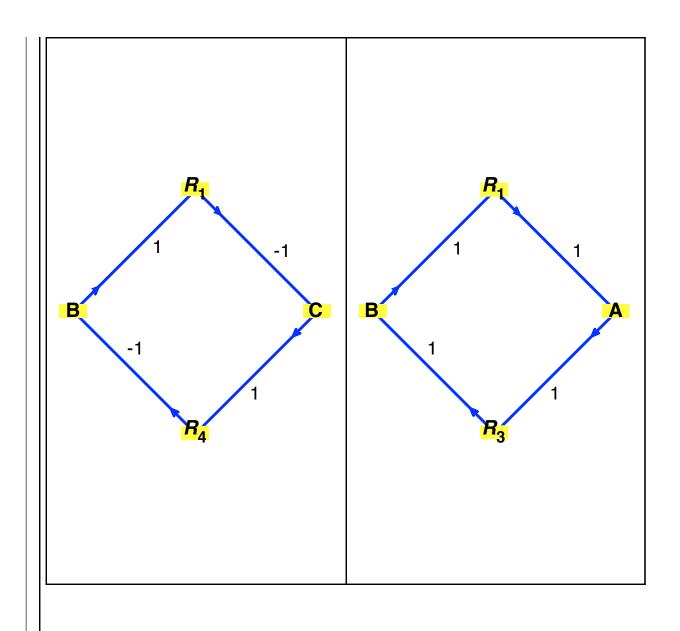
No. 66: injectiveEx3_3123.csv

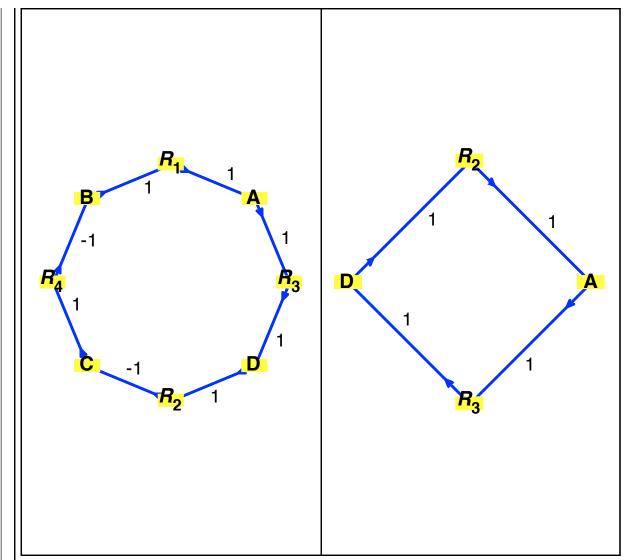




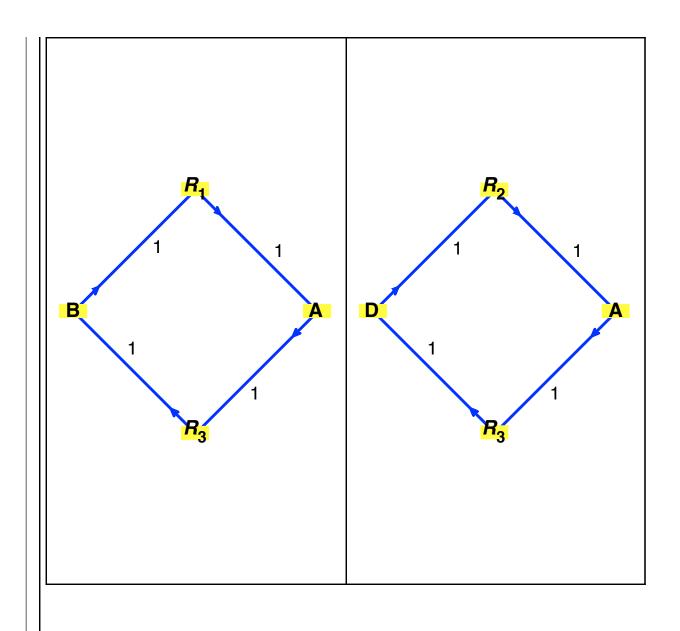


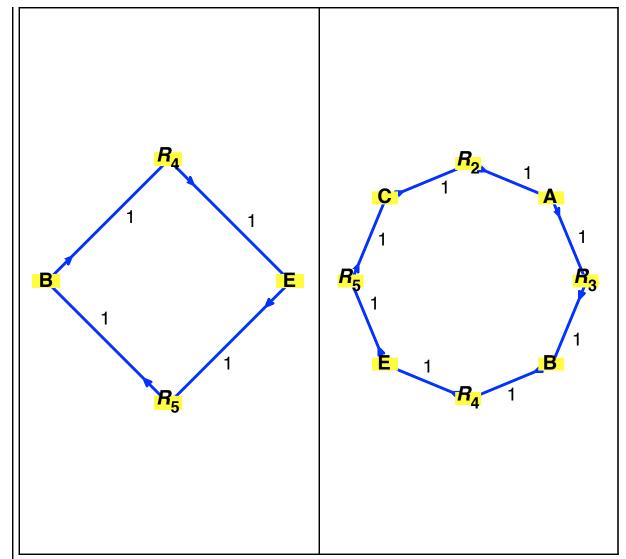
No. 67: injectiveEx3_3128.csv



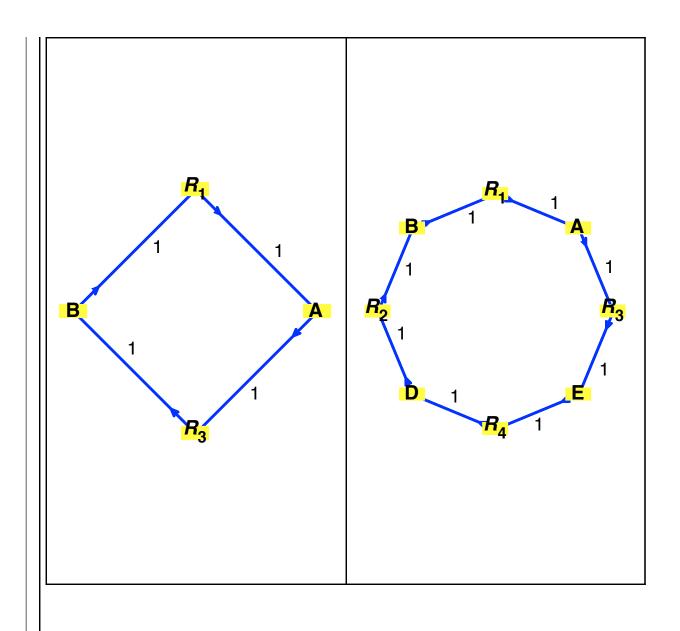


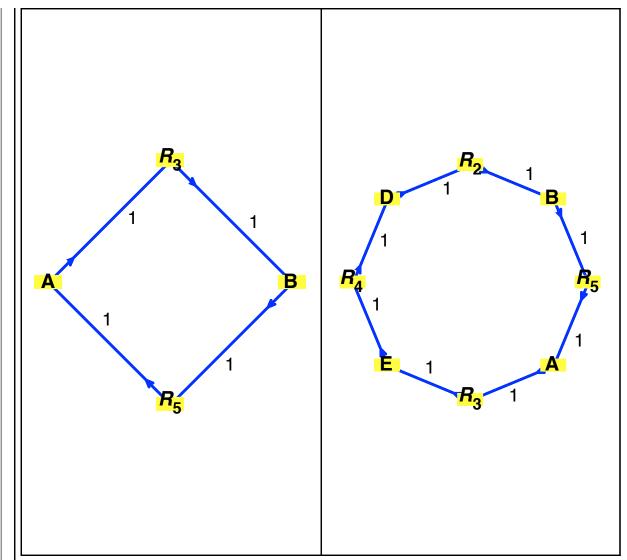
No. 68: injectiveEx3_3130.csv



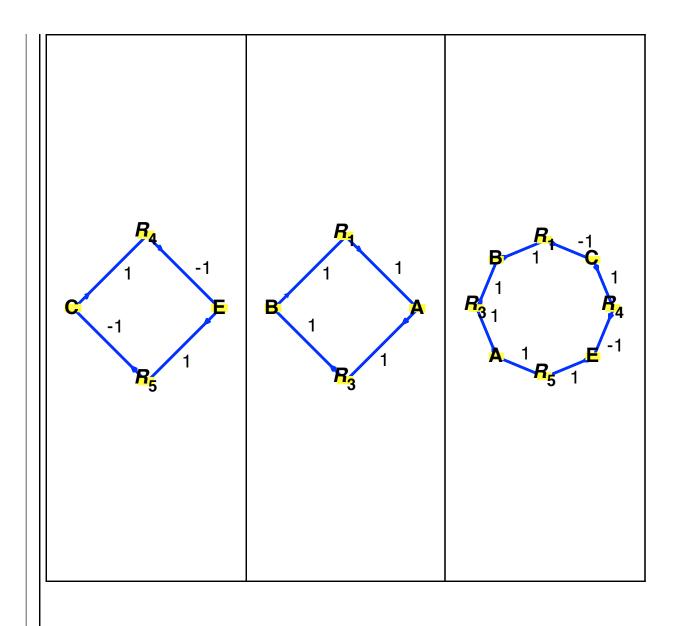


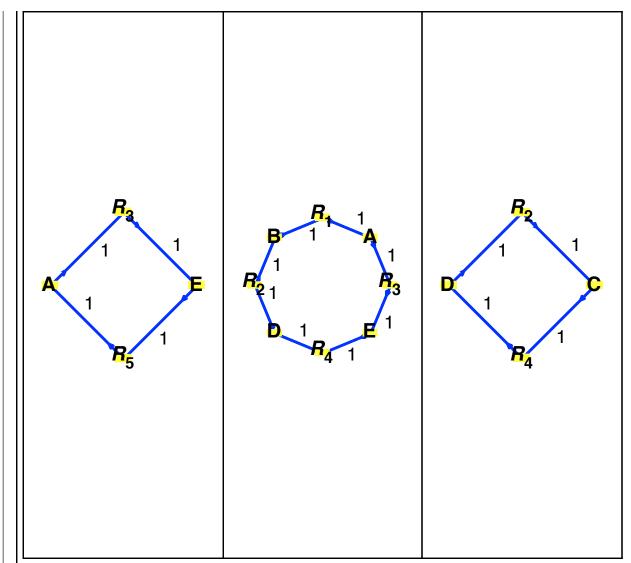
No. 69: injectiveEx3_3553.csv



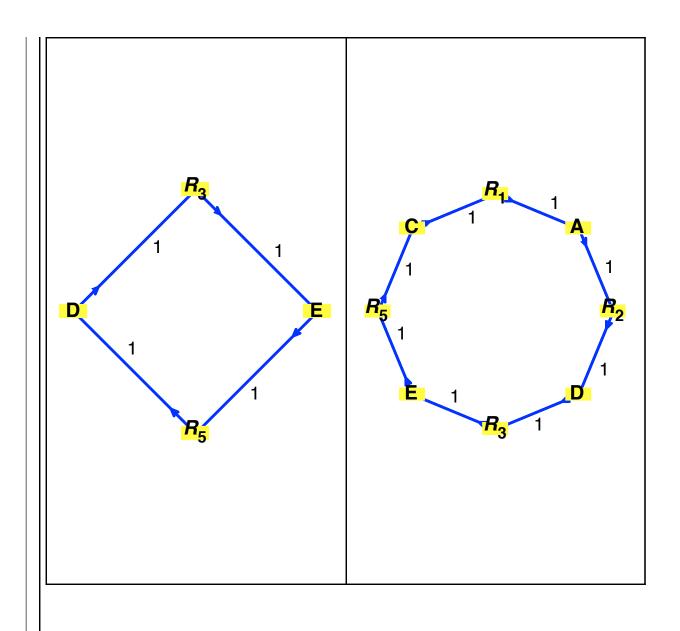


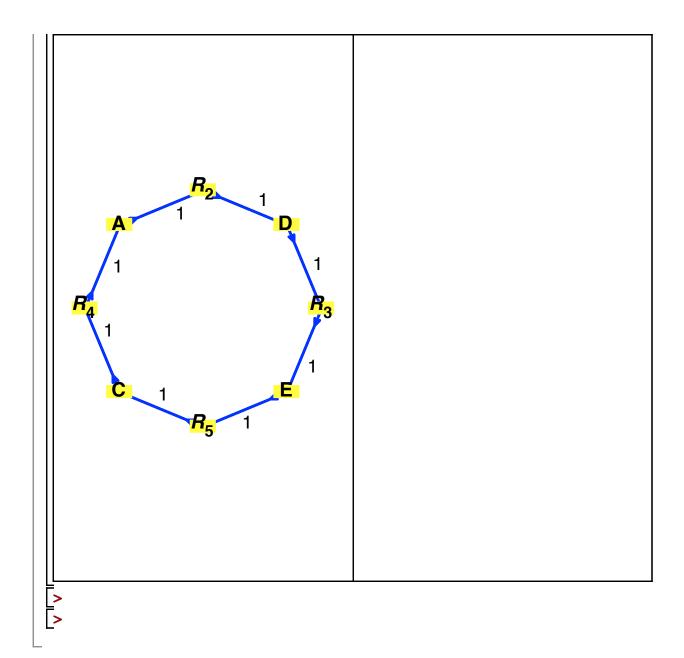
No. 70: injectiveEx3_3555.csv





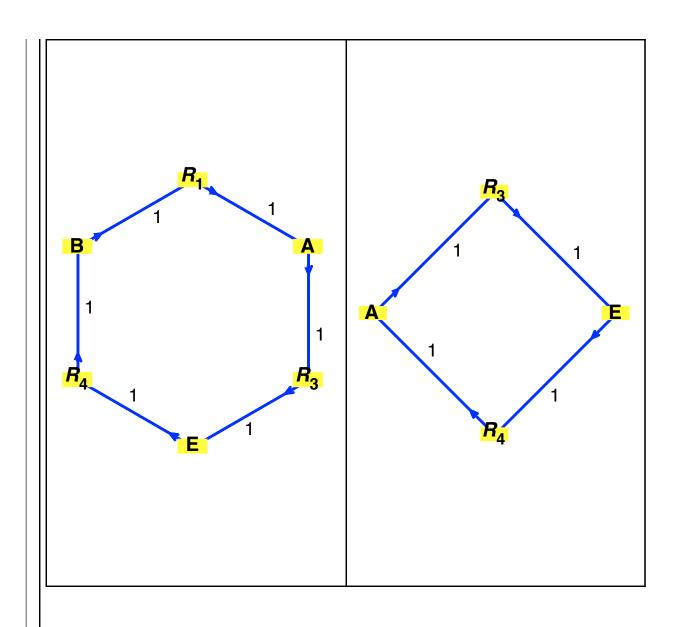
No. 71: injectiveEx3_4324.csv

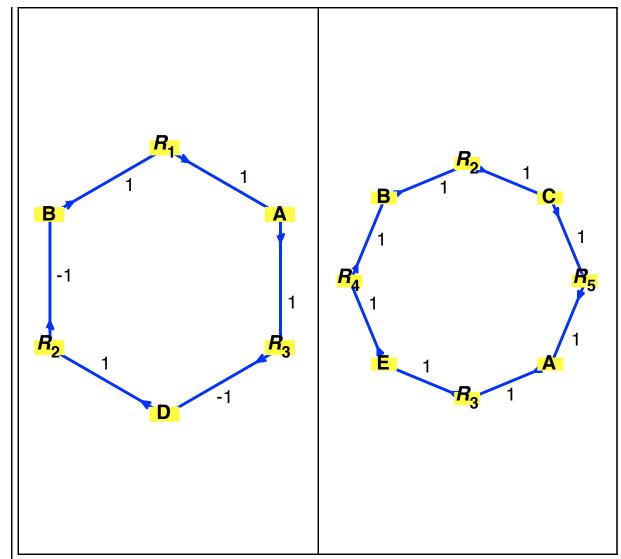




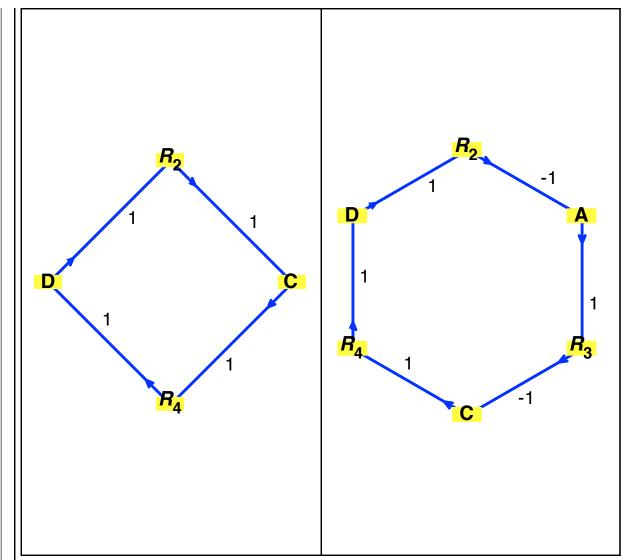
Multistationary: with competition loop, with intersecting loops

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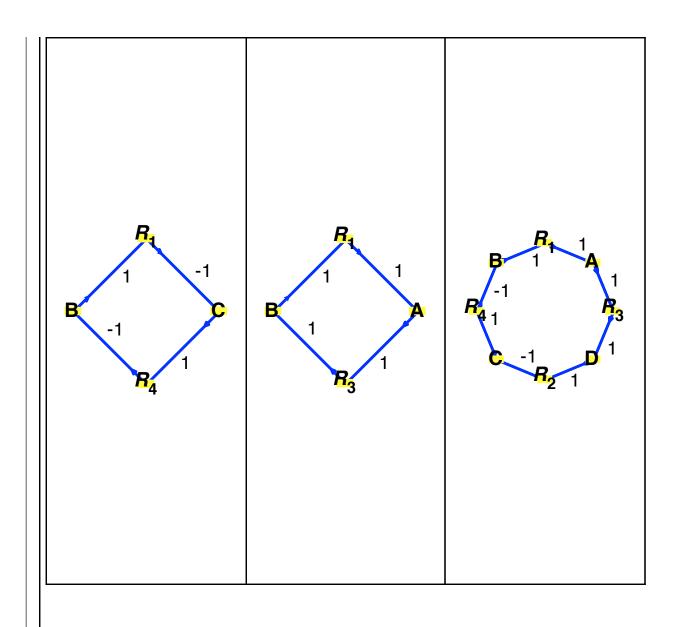


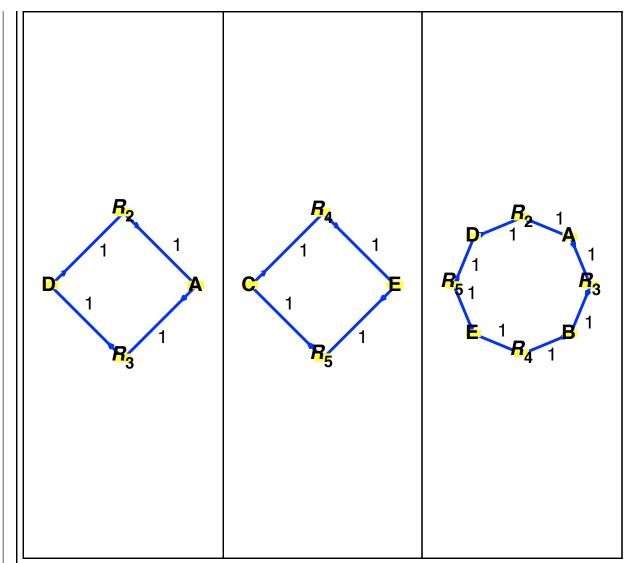


No. 2: injectiveEx0_1742.csv

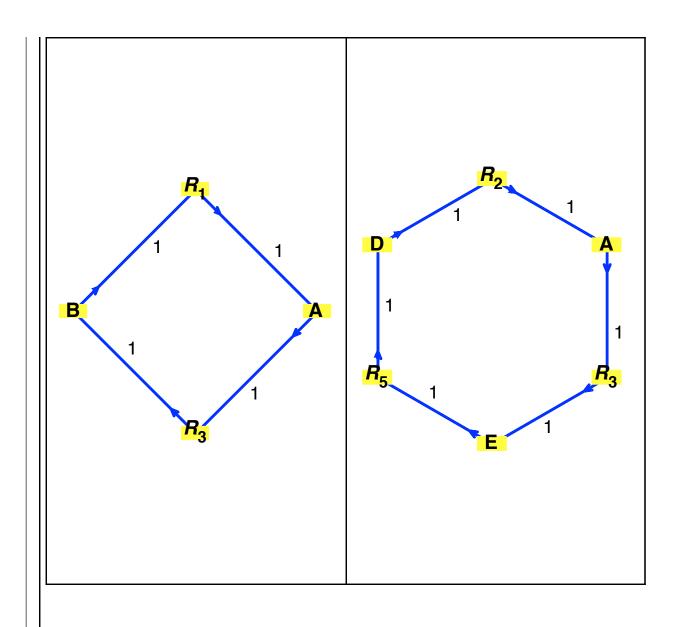


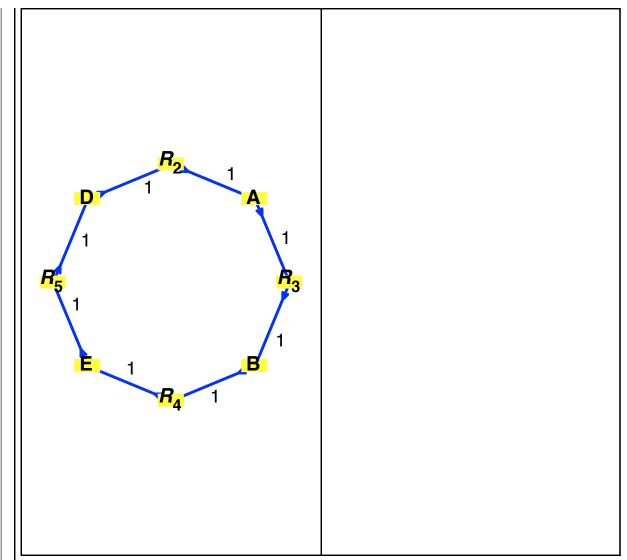
No. 3: injectiveEx0_3129.csv



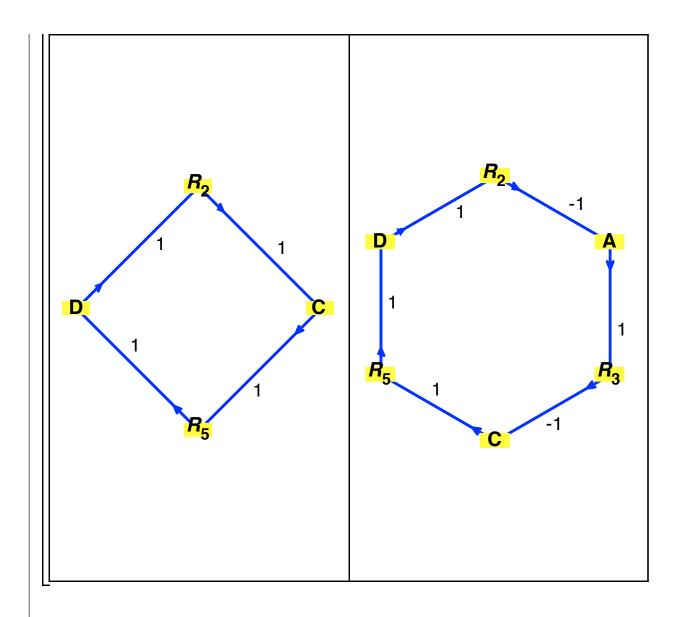


No. 4: injectiveEx0_3141.csv



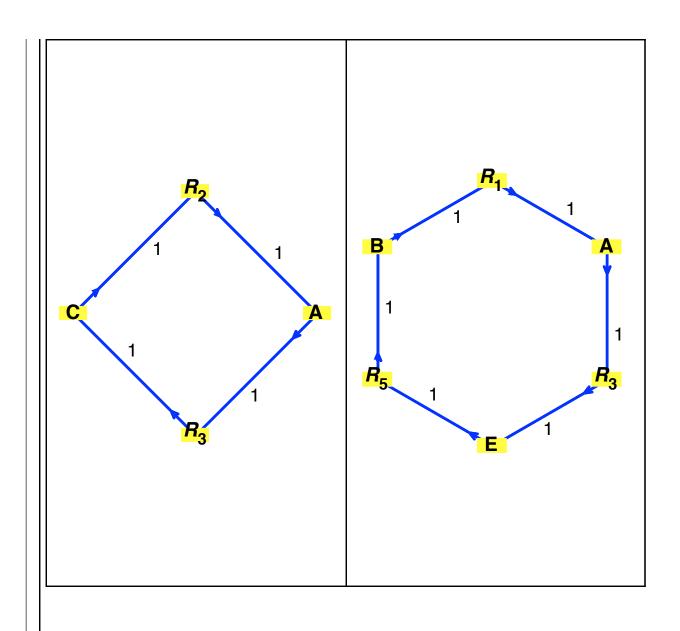


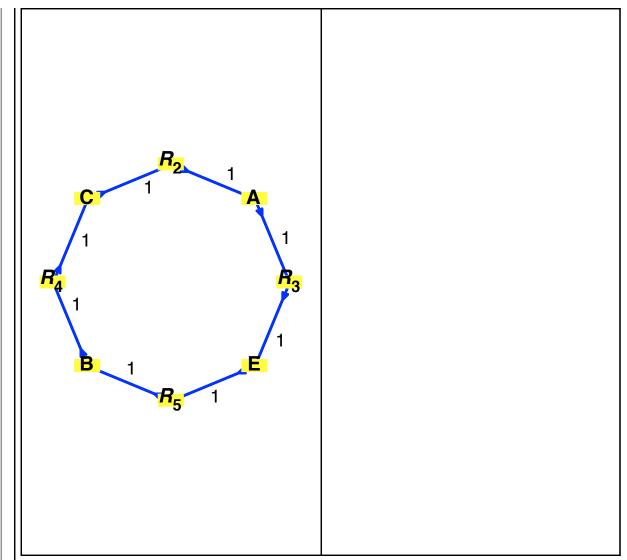
No. 5: injectiveEx2_1741.csv



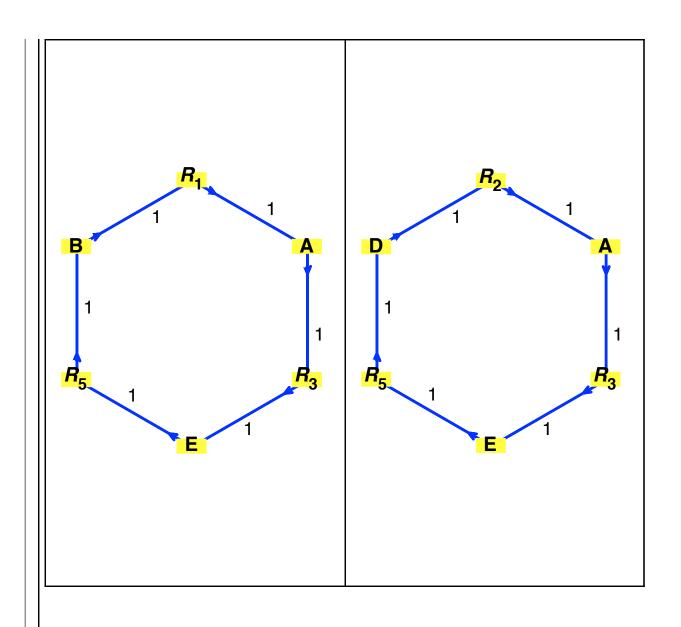
Multistationary: no competition loop, with intersecting loops

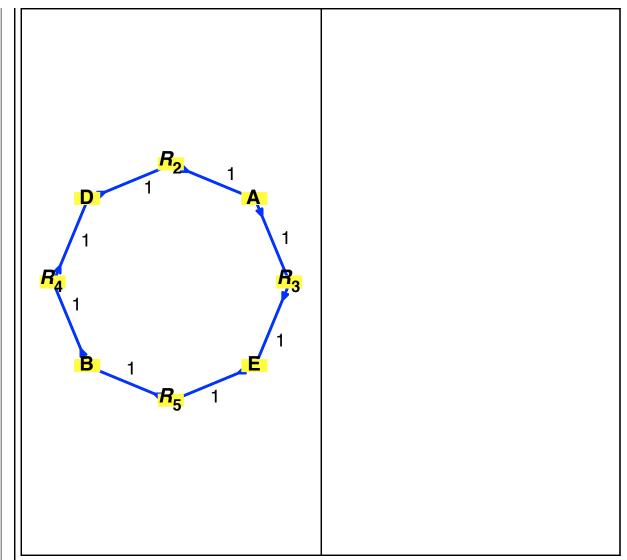
```
| wniquefolder := "5species/multistationary/unique_nocompetitionloop_intersectingloops" :
| uniquefiles := ListDirectory(uniquefolder) :
| m := nops(uniquefiles) :
| for i from 1 to m by 1 do uni := op(i, uniquefiles) : A := ImportMatrix(cat(uniquefolder, "/", uni)) : Z := findZ(A) : selected := findloops(A, Z) : printf("No. %d: %s \n", i, uni); print(drawloops(selected, speciesord)); end do:
| No. 1: injectiveEx0_1387.csv
```



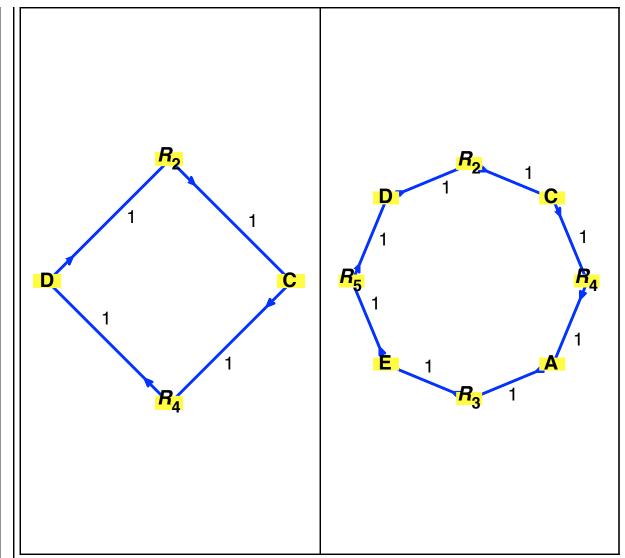


No. 2: injectiveEx0_1392.csv

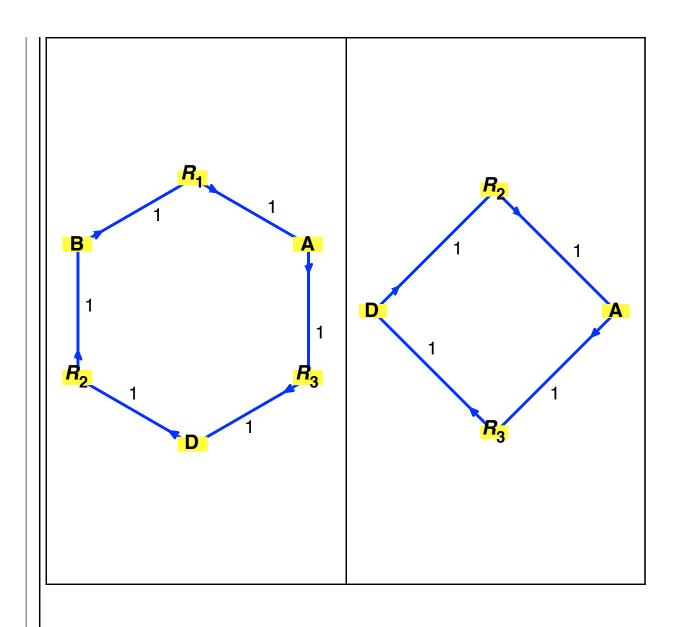


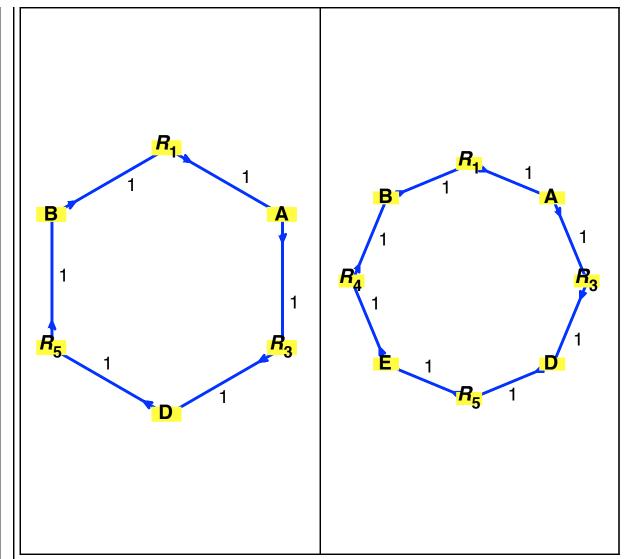


No. 3: injectiveEx0_1487.csv

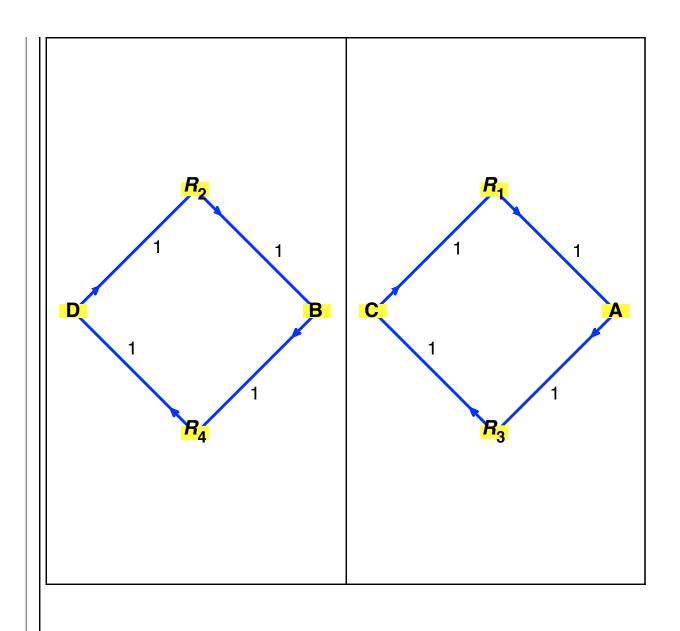


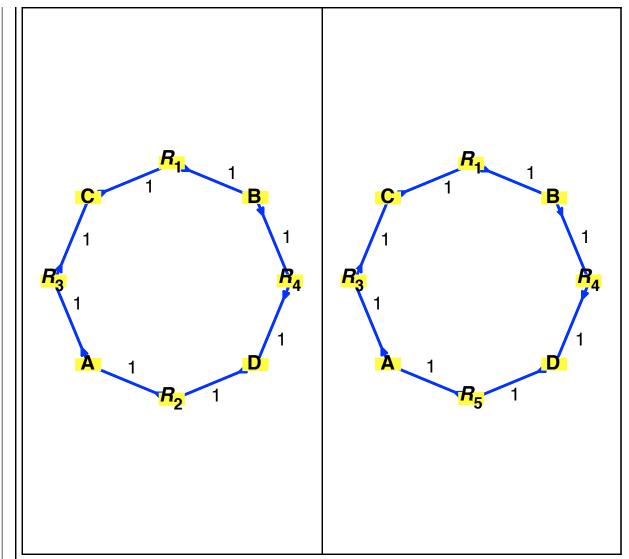
No. 4: injectiveEx0_3733.csv



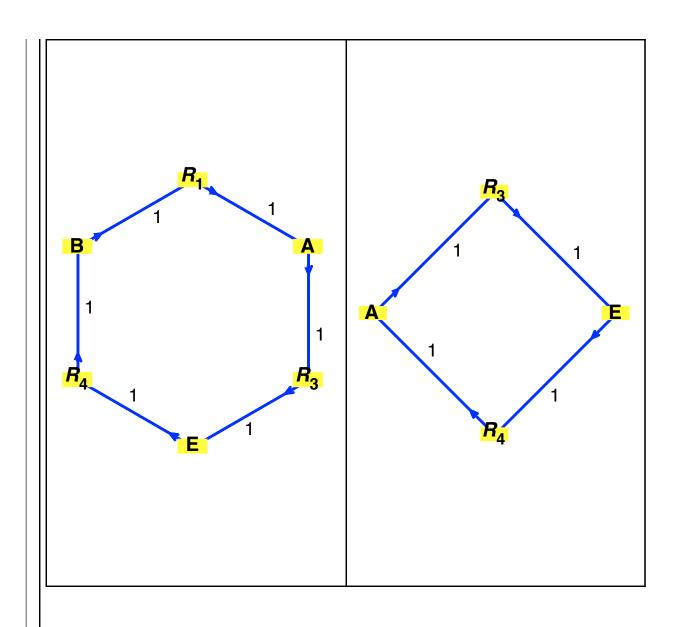


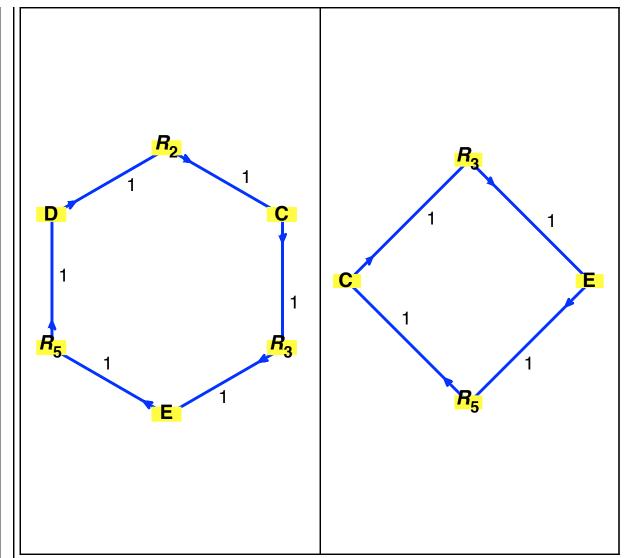
No. 5: injectiveEx0_5631.csv



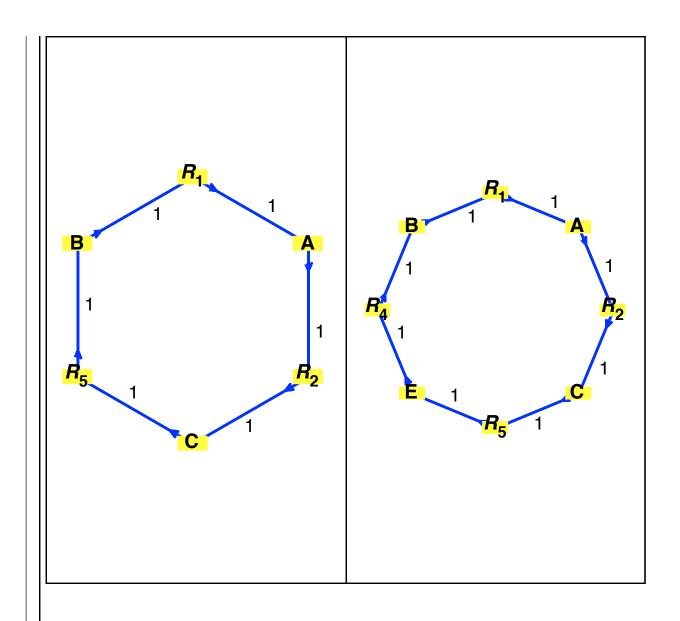


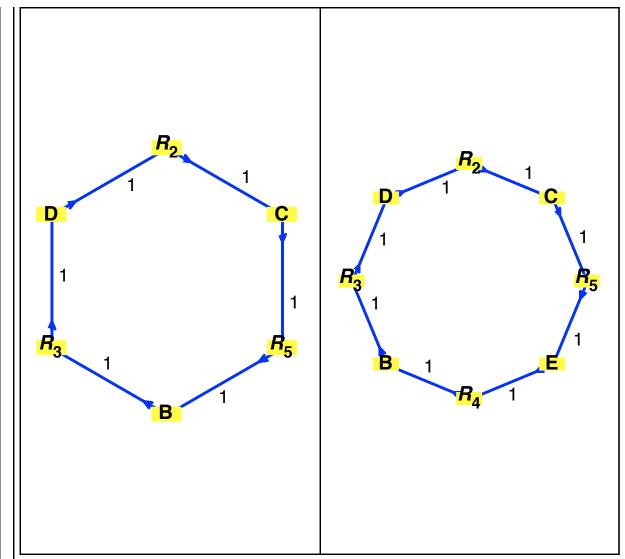
No. 6: injectiveEx2_146.cvs.csv



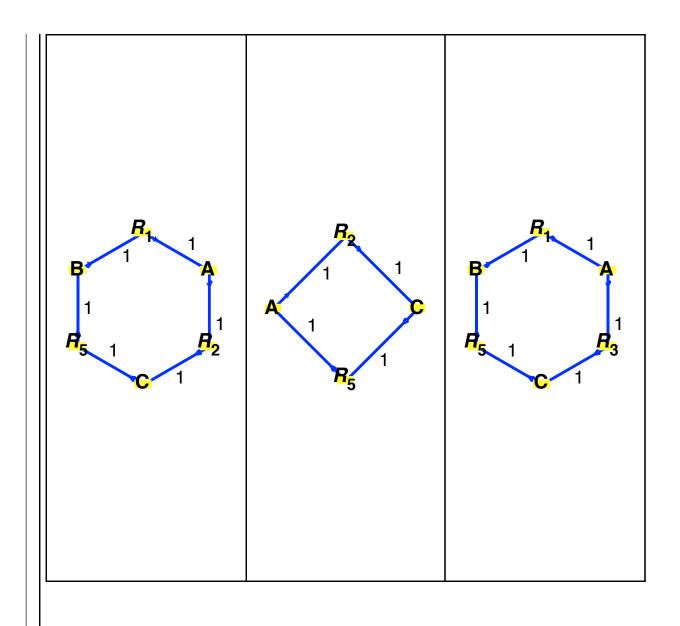


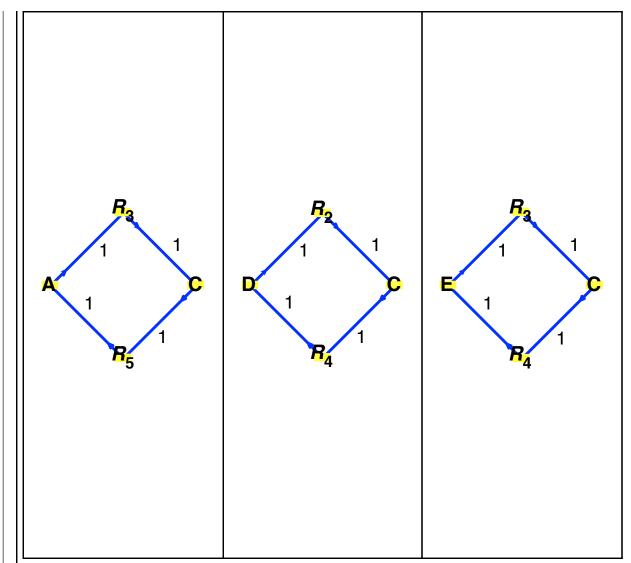
No. 7: injectiveEx2_1573.csv



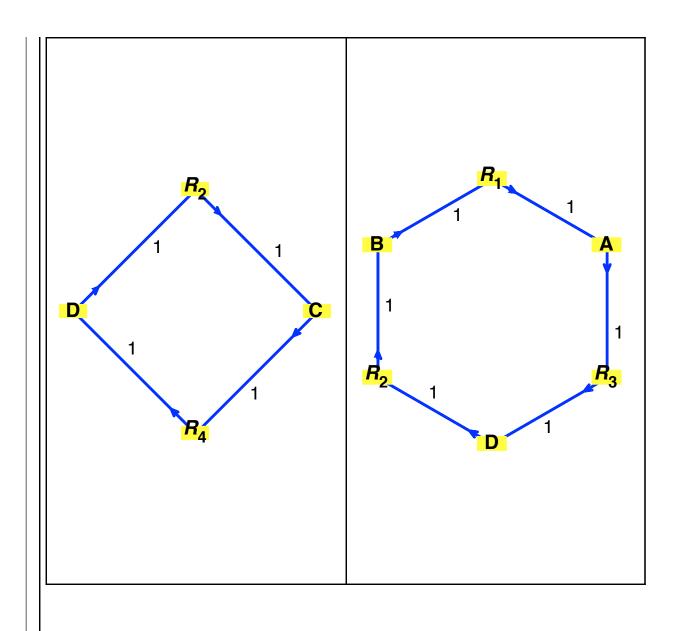


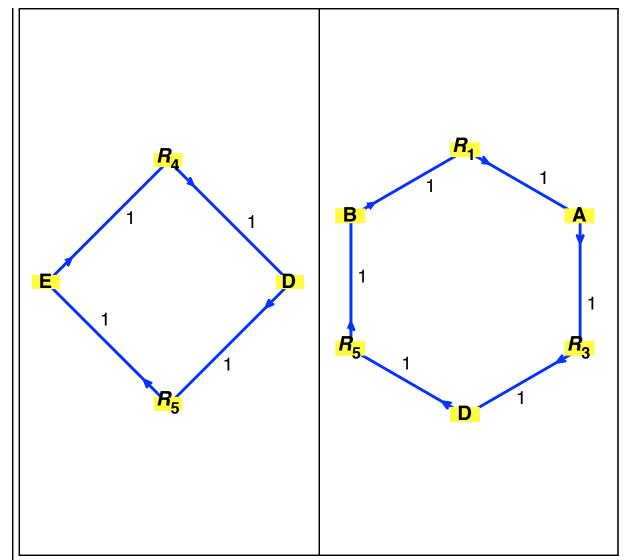
No. 8: injectiveEx2_1749.csv



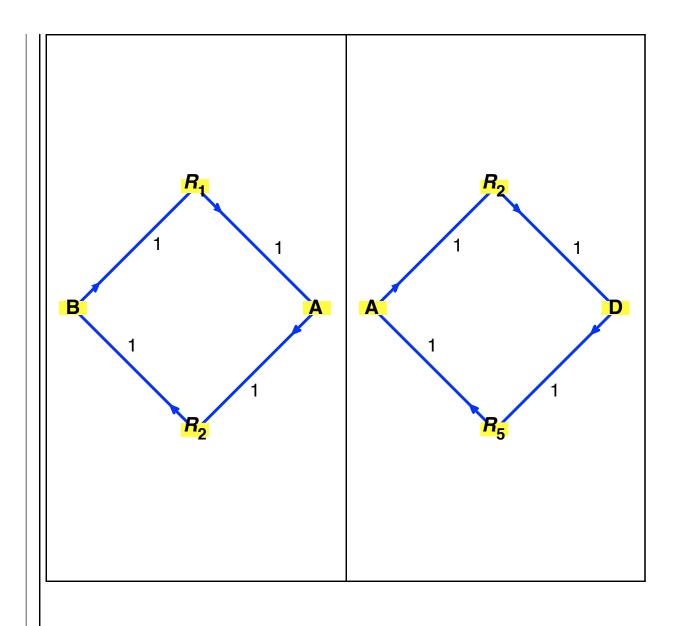


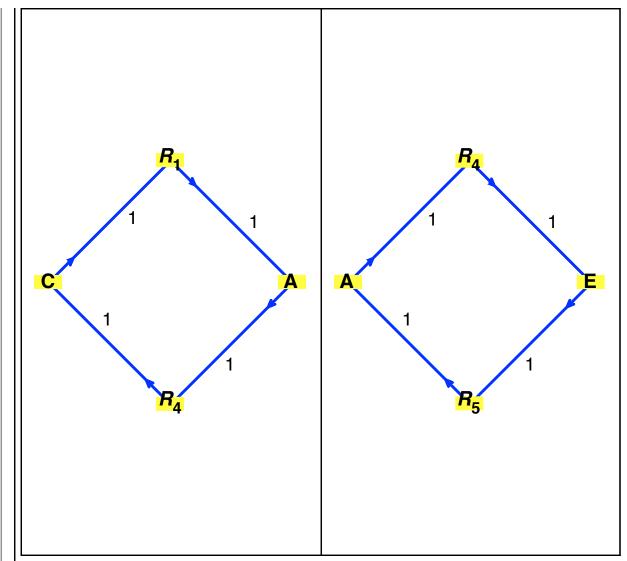
No. 9: injectiveEx2_2052.csv



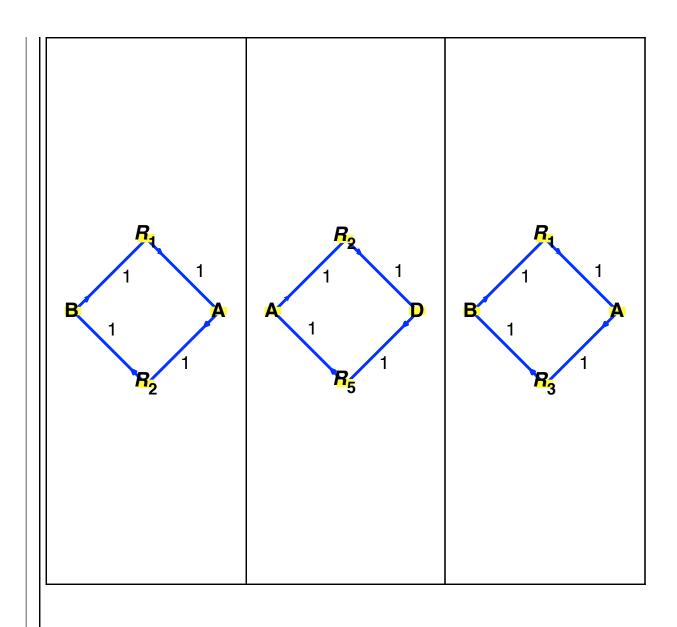


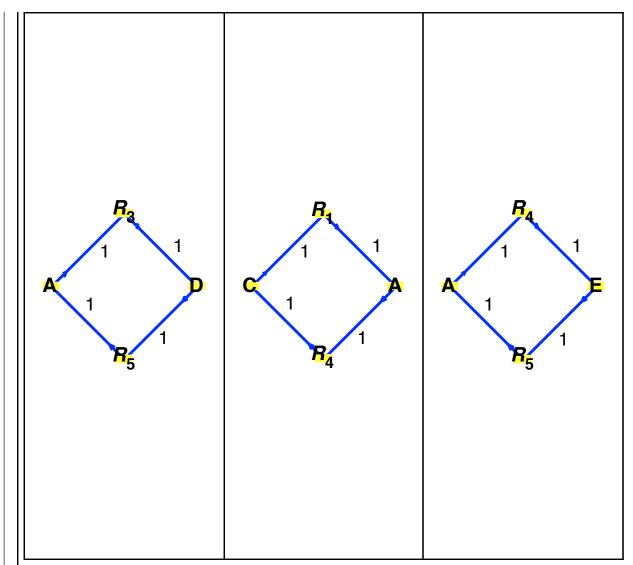
No. 10: injectiveEx2_4180.csv





No. 11: injectiveEx2_4203.csv WRONG.csv





No. 12: injectiveEx2_4242.csv WRONG.csv

