

## ▼ BPL\_TEST2\_Perfusion script with PyFMI ver 2.7.4

The key library PyFMI v2.7.4 is installed and downgrading is done Numpy v1.19.1. To simplify this we first install conda.

After the installation a small application BPL\_TEST2\_Perfusion is loaded and run. You can continue with this example if you like.

```
!lsb_release -a # Actual VM Ubuntu version used by Google
```

```
No LSB modules are available.
Distributor ID: Ubuntu
Description:    Ubuntu 18.04.6 LTS
Release:        18.04
Codename:       bionic
```

```
%env PYTHONPATH=
```



```
env: PYTHONPATH=
```

```
!wget https://repo.anaconda.com/miniconda/Miniconda3-py37_4.12.0-Linux-x86_64.sh
```

```
!chmod +x Miniconda3-py37_4.12.0-Linux-x86_64.sh
```

```
!bash ./Miniconda3-py37_4.12.0-Linux-x86_64.sh -b -f -p /usr/local
```

```
import sys
```

```
sys.path.append('/usr/local/lib/python3.7/site-packages/')
```

```
- ruamel_yaml==0.15.100=py37h27cfd23_0
- setuptools==61.2.0=py37h06a4308_0
- six==1.16.0=pyhd3eb1b0_1
- sqlite==3.38.2=hc218d9a_0
- tk==8.6.11=h1ccaba5_0
- tqdm==4.63.0=pyhd3eb1b0_0
- urllib3==1.26.8=pyhd3eb1b0_0
- wheel==0.37.1=pyhd3eb1b0_0
- xz==5.2.5=h7b6447c_0
- yaml==0.2.5=h7b6447c_0
- zlib==1.2.12=h7f8727e_1
```

The following NEW packages will be INSTALLED:

_libgcc_mutex	pkgs/main/linux-64::_libgcc_mutex-0.1-main
_openmp_mutex	pkgs/main/linux-64::_openmp_mutex-4.5-1_gnu
brotlipy	pkgs/main/linux-64::brotlipy-0.7.0-py37h27cfd23_1003
ca-certificates	pkgs/main/linux-64::ca-certificates-2022.3.29-h06a4308_1
certifi	pkgs/main/linux-64::certifi-2021.10.8-py37h06a4308_2
cffi	pkgs/main/linux-64::cffi-1.15.0-py37hd667e15_1
charset-normalizer	pkgs/main/noarch::charset-normalizer-2.0.4-pyhd3eb1b0_0
colorama	pkgs/main/noarch::colorama-0.4.4-pyhd3eb1b0_0
conda	pkgs/main/linux-64::conda-4.12.0-py37h06a4308_0
conda-content-tru~	pkgs/main/noarch::conda-content-trust-0.1.1-pyhd3eb1b0_0
conda-package-han~	pkgs/main/linux-64::conda-package-handling-1.8.1-py37h7f8
cryptography	pkgs/main/linux-64::cryptography-36.0.0-py37h9ce1e76_0

```

lana pkgs/main/noarch::lana-3.3-pyhd3eb1b0_0
ld_impl_linux-64 pkgs/main/linux-64::ld_impl_linux-64-2.35.1-h7274673_9
libffi pkgs/main/linux-64::libffi-3.3-he6710b0_2
libgcc-ng pkgs/main/linux-64::libgcc-ng-9.3.0-h5101ec6_17
libgomp pkgs/main/linux-64::libgomp-9.3.0-h5101ec6_17
libstdcxx-ng pkgs/main/linux-64::libstdcxx-ng-9.3.0-hd4cf53a_17
ncurses pkgs/main/linux-64::ncurses-6.3-h7f8727e_2
openssl pkgs/main/linux-64::openssl-1.1.1n-h7f8727e_0
pip pkgs/main/linux-64::pip-21.2.2-py37h06a4308_0
pycosat pkgs/main/linux-64::pycosat-0.6.3-py37h27cfd23_0

pycparser pkgs/main/noarch::pycparser-2.21-pyhd3eb1b0_0
pyopenssl pkgs/main/noarch::pyopenssl-22.0.0-pyhd3eb1b0_0
pysocks pkgs/main/linux-64::pysocks-1.7.1-py37_1
python pkgs/main/linux-64::python-3.7.13-h12debd9_0
readline pkgs/main/linux-64::readline-8.1.2-h7f8727e_1
requests pkgs/main/noarch::requests-2.27.1-pyhd3eb1b0_0
ruamel_yaml pkgs/main/linux-64::ruamel_yaml-0.15.100-py37h27cfd23_0
setuptools pkgs/main/linux-64::setuptools-61.2.0-py37h06a4308_0
six pkgs/main/noarch::six-1.16.0-pyhd3eb1b0_1
sqlite pkgs/main/linux-64::sqlite-3.38.2-hc218d9a_0
tk pkgs/main/linux-64::tk-8.6.11-h1ccaba5_0
tqdm pkgs/main/noarch::tqdm-4.63.0-pyhd3eb1b0_0
urllib3 pkgs/main/noarch::urllib3-1.26.8-pyhd3eb1b0_0
wheel pkgs/main/noarch::wheel-0.37.1-pyhd3eb1b0_0
xz pkgs/main/linux-64::xz-5.2.5-h7b6447c_0
yaml pkgs/main/linux-64::yaml-0.2.5-h7b6447c_0
zlib pkgs/main/linux-64::zlib-1.2.12-h7f8727e_1

```

Preparing transaction: done

Executing transaction: done

.....

```
!conda update -n base -c defaults conda --yes
```

```

colorama-0.4.4-pyhd3eb1b0_0
conda-content-trust-0.1.1-pyhd3eb1b0_0
six-1.16.0-pyhd3eb1b0_1

```

The following packages will be UPDATED:

```

_openmp_mutex 4.5-1_gnu --> 5.1-1_gnu
ca-certificates 2022.3.29-h06a4308_1 --> 2022.07.19-h06
certifi 2021.10.8-py37h06a4308_2 --> 2022.9.24-py37
cffi 1.15.0-py37hd667e15_1 --> 1.15.1-py37h74
conda 4.12.0-py37h06a4308_0 --> 22.9.0-py37h06
conda-package-handling 1.8.1-py37h7f8727e_0 --> 1.9.0-py37h5ee
cryptography 36.0.0-py37h9ce1e76_0 --> 37.0.1-py37h9c
ld_impl_linux-64 2.35.1-h7274673_9 --> 2.38-h1181459_
libgcc-ng 9.3.0-h5101ec6_17 --> 11.2.0-h123456
libgomp 9.3.0-h5101ec6_17 --> 11.2.0-h123456
libstdcxx-ng 9.3.0-hd4cf53a_17 --> 11.2.0-h123456
ncurses 6.3-h7f8727e_2 --> 6.3-h5eee18b_3
openssl 1.1.1n-h7f8727e_0 --> 1.1.1q-h7f8727
pip 21.2.2-py37h06a4308_0 --> 22.2.2-py37h06
requests pkgs/main/noarch::requests-2.27.1-pyh~ --> pkgs/main/linu
setuptools 61.2.0-py37h06a4308_0 --> 63.4.1-py37h06
sqlite 3.38.2-hc218d9a_0 --> 3.39.3-h508229
tk 8.6.11-h1ccaba5_0 --> 8.6.12-h1ccaba
tqdm pkgs/main/noarch::tqdm-4.63.0-pyhd3eb~ --> pkgs/main/linu
urllib3 pkgs/main/noarch::urllib3-1.26.8-pyhd3eb1b0_0 --> pkgs/main/linu

```

```

urllib3          pkgs/main/noarch::urllib3-1.26.8-py39~ --> pkgs/main/linux
xz                5.2.5-h7b6447c_0 --> 5.2.6-h5eee18k
zlib              1.2.12-h7f8727e_1 --> 1.2.12-h5eee18k

```

#### Downloading and Extracting Packages

```

xz-5.2.6          | 394 KB      | : 100% 1.0/1 [00:00<00:00, 7.43it/s]
ncurses-6.3       | 781 KB      | : 100% 1.0/1 [00:00<00:00, 3.67it/s]
certifi-2022.9.24 | 154 KB      | : 100% 1.0/1 [00:00<00:00, 16.77it/s]
conda-package-handl | 887 KB      | : 100% 1.0/1 [00:00<00:00, 13.55it/s]
tk-8.6.12         | 3.0 MB      | : 100% 1.0/1 [00:00<00:00, 6.13it/s]
ca-certificates-2022 | 124 KB      | : 100% 1.0/1 [00:00<00:00, 18.02it/s]

pip-22.2.2        | 2.3 MB      | : 100% 1.0/1 [00:00<00:00, 4.54it/s]
tqdm-4.64.1       | 126 KB      | : 100% 1.0/1 [00:00<00:00, 15.22it/s]
setuptools-63.4.1 | 1.1 MB      | : 100% 1.0/1 [00:00<00:00, 7.88it/s]
ld_impl_linux-64-2.3 | 654 KB      | : 100% 1.0/1 [00:00<00:00, 13.93it/s]
openssl-1.1.1q    | 2.5 MB      | : 100% 1.0/1 [00:00<00:00, 5.16it/s]
libgomp-11.2.0    | 474 KB      | : 100% 1.0/1 [00:00<00:00, 12.58it/s]
cryptography-37.0.1 | 1.3 MB      | : 100% 1.0/1 [00:00<00:00, 8.06it/s]
sqlite-3.39.3     | 1.1 MB      | : 100% 1.0/1 [00:00<00:00, 13.07it/s]
conda-22.9.0      | 878 KB      | : 100% 1.0/1 [00:00<00:00, 6.25it/s]
requests-2.28.1   | 92 KB       | : 100% 1.0/1 [00:00<00:00, 17.26it/s]
libstdcxx-ng-11.2.0 | 4.7 MB      | : 100% 1.0/1 [00:00<00:00, 5.71it/s]
cffi-1.15.1       | 227 KB      | : 100% 1.0/1 [00:00<00:00, 15.75it/s]
libgcc-ng-11.2.0  | 5.3 MB      | : 100% 1.0/1 [00:00<00:00, 4.19it/s]
toolz-0.11.2      | 49 KB       | : 100% 1.0/1 [00:00<00:00, 15.53it/s]
zlib-1.2.12       | 103 KB      | : 100% 1.0/1 [00:00<00:00, 17.23it/s]
_openmp_mutex-5.1 | 21 KB       | : 100% 1.0/1 [00:00<00:00, 17.10it/s]
urllib3-1.26.11   | 181 KB      | : 100% 1.0/1 [00:00<00:00, 15.52it/s]

```

Preparing transaction: done

Verifying transaction: done

Executing transaction: done

```
!conda --version
```

```
!python --version
```

```
conda 22.9.0
```

```
Python 3.7.13
```

```
!conda install -c conda-forge pyfmi==2.7.4 --yes # Install the key package
```

```

liblapack          conda-forge/linux-64::liblapack-0.3.0-15_linux64_openblas
libopenblas        conda-forge/linux-64::libopenblas-0.3.20-pthreads_h78a641
libxml2            conda-forge/linux-64::libxml2-2.9.12-h72842e0_0 None
libxslt            conda-forge/linux-64::libxslt-1.1.33-h15afd5d_2 None
lxml               conda-forge/linux-64::lxml-4.8.0-py37h540881e_2 None
metis             conda-forge/linux-64::metis-5.1.0-h58526e2_1006 None
mpfr              conda-forge/linux-64::mpfr-4.1.0-h9202a9a_1 None
numpy             conda-forge/linux-64::numpy-1.21.6-py37h976b520_0 None
pyfmi             conda-forge/linux-64::pyfmi-2.7.4-py37h161383b_0 None
python_abi        conda-forge/linux-64::python_abi-3.7-2_cp37m None
scipy             conda-forge/linux-64::scipy-1.7.3-py37hf2a6cf1_0 None
suitesparse       conda-forge/linux-64::suitesparse-5.10.1-h9e50725_1 None
sundials          conda-forge/linux-64::sundials-5.8.0-h558c624_0 None
tbb               conda-forge/linux-64::tbb-2021.5.0-h924138e_1 None

```

The following packages will be UPDATED:

```

ca-certificates    pkgs/main::ca-certificates-2022.07.19~ --> conda-forge::c
conda              pkgs/main::conda-22.9.0-h7b6447c_0 --> conda-forge::c

```

conda

pkgs/main::conda-22.9.0-pys/n06a4308\_0 --&gt; conda-forge::c

The following packages will be SUPERSEDED by a higher-priority channel:

```
certifi          pkgs/main/linux-64::certifi-2022.9.24~ --> conda-forge/nc
openssl          pkgs/main::openssl-1.1.1q-h7f8727e_0 --> conda-forge::c
```

#### Downloading and Extracting Packages

tbb-2021.5.0	1.9 MB	: 100% 1.0/1 [00:00<00:00, 2.23it/s]
sundials-5.8.0	1.0 MB	: 100% 1.0/1 [00:00<00:00, 3.27it/s]
libblas-3.9.0	12 KB	: 100% 1.0/1 [00:00<00:00, 26.02it/s]
fmilib-2.2.3	532 KB	: 100% 1.0/1 [00:00<00:00, 6.63it/s]
assimulo-3.2.9	2.6 MB	: 100% 1.0/1 [00:00<00:00, 1.77it/s]
libxslt-1.1.33	522 KB	: 100% 1.0/1 [00:00<00:00, 5.68it/s]
libgfortran5-12.1.0	1.8 MB	: 100% 1.0/1 [00:00<00:00, 2.76it/s]
libgfortran-ng-12.1.	23 KB	: 100% 1.0/1 [00:00<00:00, 26.59it/s]
openssl-1.1.1o	2.1 MB	: 100% 1.0/1 [00:00<00:00, 2.63it/s]
metis-5.1.0	4.1 MB	: 100% 1.0/1 [00:00<00:00, 1.28it/s]
liblapack-3.9.0	12 KB	: 100% 1.0/1 [00:00<00:00, 27.62it/s]
libxml2-2.9.12	772 KB	: 100% 1.0/1 [00:00<00:00, 6.09it/s]
libcbblas-3.9.0	12 KB	: 100% 1.0/1 [00:00<00:00, 29.07it/s]
icu-68.2	13.1 MB	: 100% 1.0/1 [00:02<00:00, 2.14s/it]
python_abi-3.7	4 KB	: 100% 1.0/1 [00:00<00:00, 31.95it/s]
libiconv-1.17	1.4 MB	: 100% 1.0/1 [00:00<00:00, 4.53it/s]
gmp-6.2.1	806 KB	: 100% 1.0/1 [00:00<00:00, 5.39it/s]
scipy-1.7.3	21.8 MB	: 100% 1.0/1 [00:03<00:00, 3.86s/it]
suitesparse-5.10.1	2.4 MB	: 100% 1.0/1 [00:00<00:00, 2.15it/s]
conda-22.9.0	960 KB	: 100% 1.0/1 [00:00<00:00, 3.66it/s]
certifi-2022.9.24	155 KB	: 100% 1.0/1 [00:00<00:00, 16.22it/s]
ca-certificates-2022	150 KB	: 100% 1.0/1 [00:00<00:00, 19.54it/s]
libopenblas-0.3.20	10.1 MB	: 100% 1.0/1 [00:01<00:00, 1.83s/it]
pyfmi-2.7.4	12.4 MB	: 100% 1.0/1 [00:01<00:00, 1.54s/it]
lxml-4.8.0	1.4 MB	: 100% 1.0/1 [00:00<00:00, 3.24it/s]
numpy-1.21.6	6.1 MB	: 100% 1.0/1 [00:01<00:00, 1.35s/it]
mpfr-4.1.0	2.6 MB	: 100% 1.0/1 [00:00<00:00, 2.33it/s]

Preparing transaction: done

Verifying transaction: done

Executing transaction: done

Retrieving notices: ...working... done

```
!conda install numpy=1.19.1 --yes # Need to downgrade numpy
```

Collecting package metadata (current\_repodata.json): done

Solving environment: failed with initial frozen solve. Retrying with flexible

Collecting package metadata (repodata.json): done

Solving environment: done

## Package Plan ##

environment location: /usr/local

added / updated specs:

- numpy=1.19.1

The following packages will be downloaded:

package	build
---------	-------

```

-----|-----
blas-1.0                | openblas                | 46 KB
numpy-1.19.1            | py37h30dfecb_0         | 21 KB
numpy-base-1.19.1      | py37h75fe3a5_0         | 4.1 MB
-----|-----
Total:                  | 4.2 MB

```

The following NEW packages will be INSTALLED:

```

blas                pkgs/main/linux-64::blas-1.0-openblas None
numpy-base         pkgs/main/linux-64::numpy-base-1.19.1-py37h75fe3a5_0 None

```

The following packages will be UPDATED:

```

openssl            conda-forge::openssl-1.1.1o-h166bdaf_0 --> pkgs/main::ope

```

The following packages will be SUPERSEDED by a higher-priority channel:

```

ca-certificates    conda-forge::ca-certificates-2022.9.2~ --> pkgs/main::ca-
certifi            conda-forge/noarch::certifi-2022.9.24~ --> pkgs/main/linu
conda              conda-forge::conda-22.9.0-py37h89c186~ --> pkgs/main::cor
numpy              conda-forge::numpy-1.21.6-py37h976b52~ --> pkgs/main::num

```

Downloading and Extracting Packages

```

blas-1.0           | 46 KB      | : 100% 1.0/1 [00:00<00:00, 10.22it/s]
numpy-1.19.1       | 21 KB      | : 100% 1.0/1 [00:00<00:00, 10.29it/s]
numpy-base-1.19.1 | 4.1 MB     | : 100% 1.0/1 [00:00<00:00, 3.06it/s]
Preparing transaction: done
Verifying transaction: done
Executing transaction: done
Retrieving notices: ...working... done

```

## ▼ Notes of BPL\_TEST2\_Perfusion

This notebook explore perfusion cultivation in comparison with ordinary continuous cultivation (chemostat) and use comparable settings to earlier notebook. Further you see here examples of interaction with the simplified commands `par()`, `init()`, `simu()` etc as well as direct interaction with the FMU which is called "model" here. The last simulation is always available in the workspace and called "sim\_res". Note that `describe()` brings mainly up from descriptive information from the Modelica code from the FMU but is complemented by some information given in the Python setup file.

Now specific installation run a simulation and notebook for that Start with connecting to Github. Then upload the two files:

- FMU - BPL\_TEST2\_Perfusion\_linux\_jm\_cs.fmu
- Setup-file - BPL\_TEST2\_Perfusion\_explore.py

```
# Filter out DeprecationWarnings for 'np.float as alias' is needed - wish I could m
```

```

import warnings
warnings.filterwarnings("ignore")

%%bash
git clone https://github.com/janpeter19/BPL_TEST2_Perfusion

    Cloning into 'BPL_TEST2_Perfusion'...

%cd BPL_TEST2_Perfusion

/content/BPL_TEST2_Perfusion

run -i BPL_TEST2_Perfusion_explore.py

Linux - run FMU pre-comiled JModelica 2.4

Model for bioreactor has been setup. Key commands:
- par()      - change of parameters and initial values
- init()     - change initial values only
- simu()     - simulate and plot
- newplot()  - make a new plot
- show()     - show plot from previous simulation
- disp()     - display parameters and initial values from the last simulation
- describe() - describe culture, broth, parameters, variables with values /

Note that both disp() and describe() takes values from the last simulation

Brief information about a command by help(), eg help(simu)
Key system information is listed with the command system_info()

%matplotlib inline
plt.rcParams['figure.figsize'] = [25/2.54, 20/2.54]

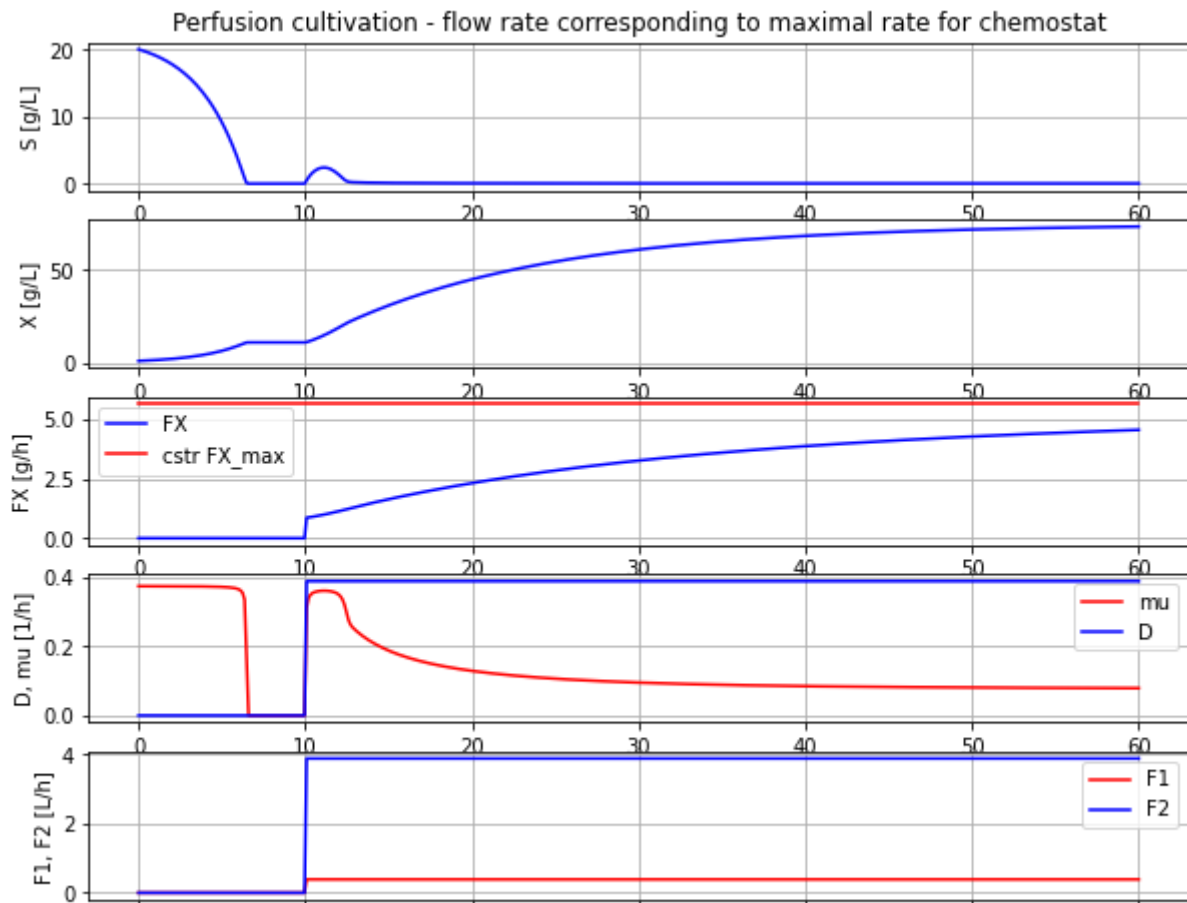
# Process parameters used throughout
par(Y=0.5, qSmax=0.75, Ks=0.1)                # Culture
par(filter_eps=0.10, filter_alpha_X=0.02, filter_alpha_S=0.10) # Filter
par(S_in=30.0)                                # Inlet substrate
init(V_0=1.0, VX_0=1.0)                      # Process initial
eps = parDict['filter_eps']                   # Pump schedule

# Simulation of process with flow rate plot to wash-out for chemostat

init(VS_0=20)                                # Process initial
par(pump1_t1=10, pump2_t1=10)                 # Pump schedule - recycle
par(pump1_F1=2.5*0.155, pump2_F1=2.5*0.155/eps)
par(pump1_t2=940, pump2_t2=940, pump1_t3=950, pump2_t3=950, pump1_t4=960, pump2_t4=960)

newplot(title='Perfusion cultivation - flow rate corresponding to maximal rate for
simu(60)

```



```
# Concentration factor of the filter
c=model.get('filter.retentate.c[1]')[0]/model.get('filter.inlet.c[1]')[0]
print('Conc factor of perfusion filter =', np.round(c,3))
```

Conc factor of perfusion filter = 1.369

```
c_data=sim_res['filter.retentate.c[1]']/sim_res['filter.inlet.c[1]']
print('Conc factor variation', np.round(min(c_data[151:]), 3), np.round(max(c_data[
```

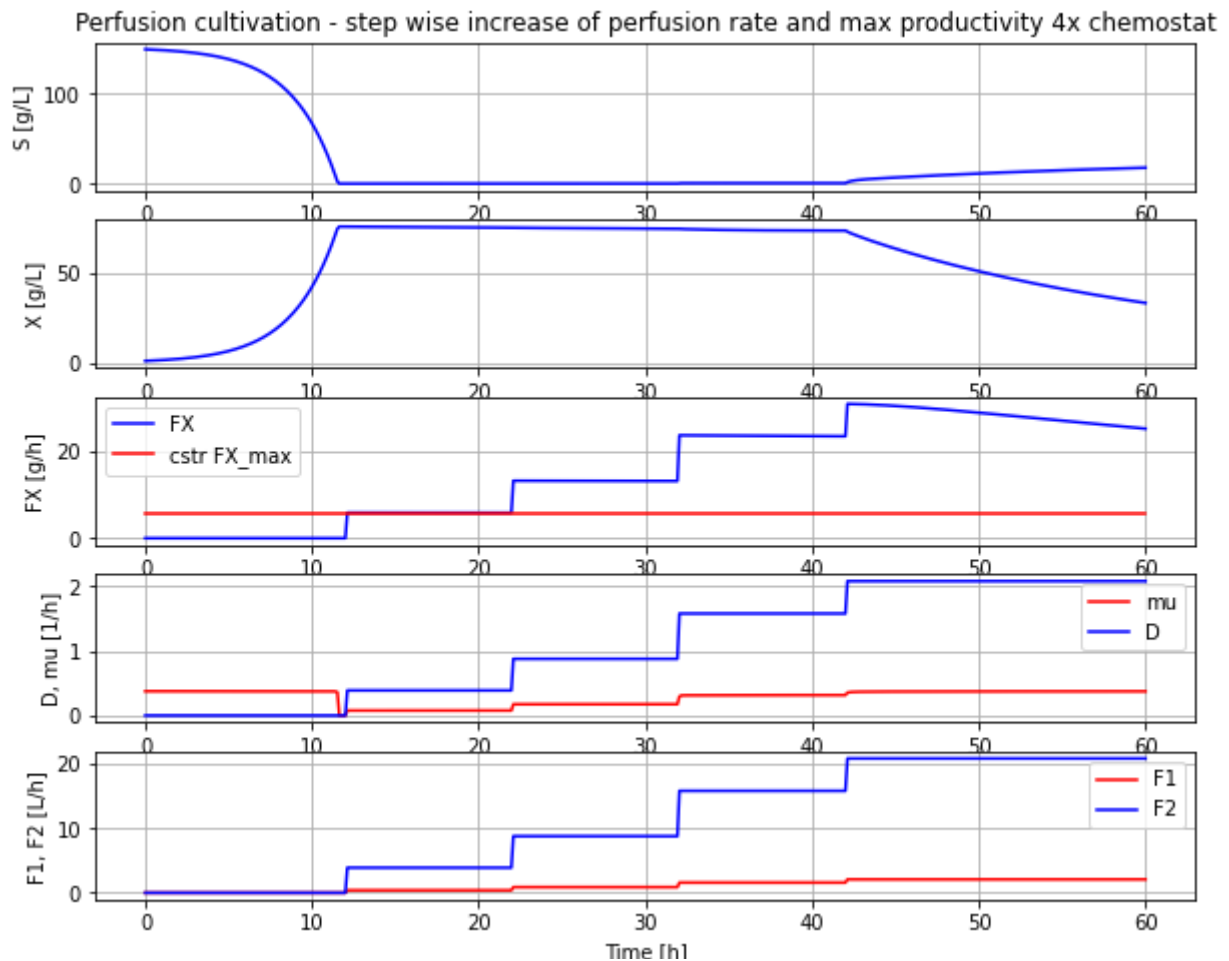
Conc factor variation 1.369 1.649

```
# Simulation of process with step-wise increase of perfusion rate until wash-out.
# This means that re-circulation rate change at the same time as the perfusion rate
```

```
init(VS_0=150) # Process initial varied

par(pump1_t1=12, pump2_t1=12) # Pump schedule - recycle
par(pump1_F1=2.5*0.155, pump2_F1=2.5*0.155/eps)
par(pump1_t2=22, pump2_t2=22)
par(pump1_F2=2.5*0.35, pump2_F2=2.5*0.35/eps)
par(pump1_t3=32, pump2_t3=32)
par(pump1_F3=2.5*0.63, pump2_F3=2.5*0.63/eps)
par(pump1_t4=42, pump2_t4=42)
par(pump1_F4=2.5*0.83, pump2_F4=2.5*0.83/eps)

newplot(title='Perfusion cultivation - step wise increase of perfusion rate and max
simu(60)
```



```
# Simulation without a plot and just to check typical values at high production rat
simu(40)
c_data=sim_res['filter.retentate.c[1]']/sim_res['filter.inlet.c[1]']
print('Conc factor variation', np.round(min(c_data[190:]), 3), 'to', np.round(max(c

Conc factor variation 1.075 to 1.087
```

```
describe('cstrProdMax')
```

```
Calculate from the model maximal chemostat productivity  $FX\_max$  : 5.625 [ g/h ]
```

```
# The maximal biomass productivity before washout is obtained aroudn 40 hours
np.round(model.get('harvesttank.inlet.F')[0]*model.get('harvesttank.inlet.c[1]')[0]

23.5
```

```
# Thus perfusion (with this filter) brings a productivity improvement of about
np.round(23.5/5.6,1)
```

```
4.2
```

```
# Finally we check the filter flow rates at time 40 hour - note the negative sign f
model.get('filter.inlet.F')[0]
```

```
15.749999999999998
```



```
model.get('filter.filtrate.F')[0]
```

```
-1.575
```

```
model.get('filter.retentate.F')[0]
```

```
-14.174999999999999
```

## ▼ Summary

- The perfusion filter had a concentration factor of cells around 1.08 and re-cycling flow was set to a factor 10 higher than the perfusion rate and changed when perfusion rate was change to keep the ratio factor 10.
- The first simulation showed that by cell retention using perfusion filter the process could be run at a perfusion flow rate at the maximal flow rate possible for corresponding chemostat culture and cell concentration increased steadily.
- The second simulation showed that with a proper startup cell concentration, the cell concentration remained constant when perfusion rate increased in a similar way as what we see in a chemostat.
- The second simulation also showed that biomass productivity in this case was increased by a factor 4.2 compared to chemostat.
- If the perfusion rate increased to higher levels washout started but the decrease of cell concentration was slow.

Some of you who read this may have your perfusion experience with CHO-cultures. For such cultures the cell concentration do increase with increase of perfusion rate and there are understood reasons for that. But for this simplified process as well as microbial processes they typically keep cell concentration constant when flow rate is chaged, and that under quite wide conditions. I will try come back to this phenomena in a later notebook.

```
# List of components in the process setup and also a couple of other things like li
describe('parts')
```

```
['bioreactor', 'bioreactor.culture', 'D', 'feedtank', 'filter', 'harvesttank',
```

```
system_info()
```

```
System information
```

```
-OS: Linux
```

```
-Python: 3.7.14
```

```
-PyFMI: 2.7.4
```

```
-FMU by: JModelica.org
```

```
-FMI: 2.0
```

```
-Type: FMUModelCS2
```

```
-Name: BPL_TEST2.Perfusion
```

```
-Generated: 2022-06-01T18:47:10
```

-Description: Bioprocess Library version 2.0.9  
-Interaction: FMU-explore ver 0.9.1

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