Experiments for fitting stochastic COVID model

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Create pomp object with data

Not sure if all initial states have to be set to 1 or greater so do that for testing purposes. REMEMBER TO UNDO.

Code model. Notice we change the parameterizatio of beta factor so beta_redfactor> 0 rather than beta_red_factor > 1. Assume negative binomial observation model with fit parameter. NEED TO SWITCH TO EULER MULTINOMIAL MODEL AT LINE 129

```
## measurement model C snippets
rmeas <- "
  cases = rnbinom_mu(theta, rho * C);
dmeas <- "
 lik = dnbinom_mu(cases, theta, rho * C, give_log);
## initializer
rinit <- "
 S = 10600000;
 E1 = 35;
 E2 = 35:
 E3 = 35;
 E4 = 35;
 E5 = 35;
 E6 = 35;
 I1 = 14;
 I2 = 14;
  I3 = 14;
 I4 = 14;
 Iu1 = 111;
 Iu2 = 111;
 Iu3 = 111;
```

```
Iu4 = 111;
 C = 1;
 Ru = 1;
## rprocess
covid_step_C <- "</pre>
 double trans[17]; //C indexes at 0, I hate that so I'm making things 1 bigger and start with index 1,
 double Epresymptom;
 double Idetected;
 double Iundetected;
 double foi; //force of infection
 double gamma; // rate of transition through I compartments
 double detect_frac; //fraction of those that get eventually diagnosed
 Epresymptom = E1+E2+E3+E4+E5+E6; //all pre-symptomatic
 Idetected = I1+I2+I3+I4;
                                  //all symptomatic that wil be detected
 //force of infection
 //time dependent transmission, multiplied by different groups
 //each group can have its own transmission rate
 //t_int1 days after simulation start, an intervention reduces transmission rate by some factor
 //t_int1 is new, not in original code. There it was assumed to be the same as t_int2
 if (t<=t_int1)</pre>
       foi = beta_d*Idetected + beta_u*Iundetected + beta_e*Epresymptom;
 else
       foi = (beta_red_factor+1)*(beta_d*Idetected + beta_u*Iundetected + beta_e*Epresymptom);
 //time-dependent rate of movement through infected and detected classes
 //t_int2 days after simulation start, the time at which individuals are diagnosed and thus the time s
 //t_int2 is caled z in original code
 if (t<t_int2) //if time is less then intervention time, duration spent in I is given by 1/gamma_u, ot
     gamma = gamma_u;
 else
     gamma = gamma_d;
 //time dependent fraction of those that move into detected category at the end of the E phase
 //t_int3 days after simulation start, the fraction detected (those that move into I instead of Iu aft
 //note that both higher fraction detected and faster rate of detection speed up arrival of individual
 //t_int3 is called w in original code, detect_frac is called q/q0/q1 in the original code
 if (t<t_int3)</pre>
   detect_frac = detect_frac_0;
   detect_frac = detect_frac_1;
 // define all transmission rates
 trans[1] = rbinom(S,1-exp(-foi*dt));
                                                 //transition from S to E
 trans[2] = rbinom(E1,1-exp(-sigma*dt));
                                                // transition between E compartments 1/2
```

```
trans[3] = rbinom(E2,1-exp(-sigma*dt));
                                                                                         // transition between E compartments
   trans[4] = rbinom(E3,1-exp(-sigma*dt));
                                                                                         // transition between E compartments
   trans[5] = rbinom(E4,1-exp(-sigma*dt));
                                                                                         // transition between E compartments 4/5
   trans[6] = rbinom(E5,1-exp(-sigma*dt));
                                                                                         // transition between E compartments 5/6
   trans[7] = rbinom(E6,(1-exp(-sigma*dt))*detect_frac);
                                                                                                                 // transition between E6 compartment
   trans[8] = rbinom(E6,(1-exp(-sigma*dt))*(1-detect_frac));
                                                                                                                        // transition between E6 compartm
   trans[10] = rbinom(I2,1-exp(-gamma*dt));
trans[11] = rbinom(I3,1-exp(-gamma*dt));
                                                                                       // transition between I compartments 2/3
                                                                                          // transition between I compartments 3/4
   trans[12] = rbinom(I4,1-exp(-gamma*dt));
                                                                                         // transition between I compartments and C
   trans[13] = rbinom(Iu1,1-exp(-gamma_u*dt));
                                                                                                // transition between Iu compartments 1/2
   trans[14] = rbinom(Iu2,1-exp(-gamma_u*dt));
                                                                                               // transition between Iu compartments 2/3
   trans[15] = rbinom(Iu3,1-exp(-gamma_u*dt));
                                                                                              // transition between Iu compartments 3/4
   trans[16] = rbinom(Iu4,1-exp(-gamma_u*dt));
                                                                                               // transition between Iu compartments and Ru
   // define all transmissions for each compartment
   S -= trans[1];
   E1 += trans[1] - trans[2];
   E2 += trans[2] - trans[3];
   E3 += trans[3] - trans[4];
   E4 += trans[4] - trans[5];
   E5 += trans[5] - trans[6];
   E6 += trans[6] - trans[7] - trans[8];
   I1 += trans[7] - trans[9];
   I2 += trans[9] - trans[10];
   I3 += trans[10] - trans[11];
   I4 += trans[11] - trans[12];
   Iu1 += trans[8] - trans[13];
   Iu2 += trans[13] - trans[14];
   Iu3 += trans[14] - trans[15];
   Iu4 += trans[15] - trans[16];
   C += trans[12]; //detected cases, assumed to be isolated and not further contribute to transmission
   Ru += trans[16]; //undetected cases that recover, assumed to not further contribute to transmission
varnames = c("S", "E1", "E2", "E3", "E4", "E5", "E6", "I1", "I2", "I3", "I4", "Iu1", "Iu2", "Iu3", "Iu4", "Iu5", "
#parameter and variable names
parnames1 = c("beta_d", "beta_u", "beta_e", "beta_red_factor", "t_int1", "t_int2", "t_int3", "gamma_u",
#initial conditions of state variables are also parameters
parnames2 = c("E1_0", "E2_0", "E3_0", "E4_0", "E5_0", "E6_0", "I1_0", "I2_0", "I3_0", "I4_0", "Iu1_0",
parnames = c(parnames1,parnames2)
inivals = c(S_0 = 10600000, E1_0 = 35, E2_0 = 35, E3_0 = 35, E4_0 = 35, E5_0 = 35, E6_0 = 35, I1_0 = 14,
Ntot = sum(inivals)
#values for parameters. beta is scaled by population size here instead of inside the simulation functio
parvals = c(beta_d = 5e-7, beta_u = 0.25/Ntot, beta_e = 0.1/Ntot, beta_red_factor = 0.5, t_int1 = 12, t
```

```
pomp(
    data= covid_ga_data[1:25,],
                                                                        # have to remove the NA to fit model, i.e. can only fit to the current
    times=seq(1,25,by=1), t0=0,
  # covar=covariate_table(birthdat, times="time"), # save this as a model for including covariates in th
    dmeasure = Csnippet(dmeas),
    rmeasure = Csnippet(rmeas),
    rinit=Csnippet(rinit),
    rprocess = euler(
         step.fun = Csnippet(covid_step_C),
        delta.t = 1/20
    ),
    partrans=parameter_trans(
        log=c("beta_d", "beta_u", "beta_e", "beta_red_factor", "t_int1", "t_int2", "t_int3", "gamma_u", "ga
        logit=c("detect_frac_0", "detect_frac_1")
    ),
    statenames = varnames,
    obsnames=c("cases"),
    paramnames = parnames,
    accumvars = c("C"),
    params = c(parvals, inivals)
) -> covid_ga_model
Set up parallelization
#ncores <- detectCores()</pre>
registerDoParallel()
Fit with MIF.
registerDoRNG(03292020)
## Warning: executing %dopar% sequentially: no parallel backend registered
theta.guess <- theta.true <- coef(covid_ga_model)</pre>
estpars <- c("beta_d", "beta_u", "beta_e", "beta_red_factor", "gamma_u", "gamma_d", "detect_frac_0")</pre>
foreach (i = 1:10, .combine = c) %dopar% { #Inspect from multiple, randomly chosen starting points
    theta.guess <- theta.true
    theta.guess[estpars] <- rlnorm(n = length(estpars),</pre>
           meanlog = log(theta.guess[estpars]), sdlog = 1)
    mif2(covid_ga_model, Nmif = 500, params = theta.guess,
    Np = 2000, cooling.fraction = 0.5,
    rw.sd = rw.sd(beta_d = 0.02, beta_u = 0.02, beta_e = 0.02, beta_red_factor = 0.02, gamma_u = 0
} -> mifs
# Use particle filter to get the likelihood at the end of MIF run
pf1 <- foreach(mf = mifs, .combine = c) %dopar% {</pre>
    pf <- replicate(n = 10, logLik(pfilter(mf, Np = 10000)))</pre>
    logmeanexp(pf)
# Pick the best parameter set
mf1 <- mifs[[which.max(pf1)]]</pre>
theta.mif <- coef(mf1)
```

Plotting

```
mifs %>%
  traces() %>%
  melt() %>%
  filter(variable=="loglik" | variable=="beta_d" | variable=="beta_u" | variable=="beta_e" | variable==
  ggplot(aes(x=iteration,y=value,group=L1,color=L1))+
  facet_wrap(~variable,scales="free_y")+
  guides(color=FALSE)
                   loglik
                                                     beta d
                                                                                      beta u
       0 -
                                                                      4e-06 -
                                   2.0e-05 -
                                                                      3e-06 -
   -2000
                                   1.5e-05 -
                                                                      2e-06 -
                                   1.0e-05 -
   -4000
                                                                      1e-06 -
                                   5.0e-06 -
   -6000
                                   0.0e+00 -
                                                                      0e+00 -
                   beta e
                                                 beta_red_factor
                                                                                    gamma_u
                                                                         1.6
   4e-08 -
                                          4 -
                                                                         1.2
   3e-08
                                          3
                                                                         8.0
  2e-08
                                          2 -
   1e-08
                                          1 -
                                                                         0.4 -
                                          0
                                                                                100 200 300 400 500
                 gamma_d
                                                  detect_frac_0
                                        0.8 -
      12 -
                                        0.6 -
       8 -
                                        0.4 -
       4 -
                                        0.2
       0
                                        0.0
             100 200 300 400 500
                                                100 200 300 400 500
                                            ò
                                                   iteration
```

Here we set up Bayesian analysis. The first code chunk records various attempts to set up the priors. None of these worked, but not really sure why – leaving here for the time being in case I want to come back.

```
# hyperparams <- list(min = theta.mif[1:31]*0.99, max = theta.mif[1:31]*1.01)
# hyperparams$min[15:31] <- ceiling(hyperparams$min[15:31] ) # integerize state variables
# hyperparams_est <- list(min=hyperparams*min[estpars], max=hyperparams*min[estpars]) # just the param
# covid.dprior <- function (params, ..., log) {</pre>
    f \leftarrow sum(dunif(c(parvals, inivals), min = hyperparams_est\$min, max = hyperparams_est\$max, log = TRU
#
    if (log) f else exp(f)
# }
# covid.dprior <- function(params, ..., log=TRUE){</pre>
      f \leftarrow dlnorm(params["beta_d"], meanlog=log(5.828e-7), sdlog=1, log=TRUE) +
#
#
        dunif(params["beta_red_factor"], 0, 1, log=TRUE) +
        dgamma(params["gamma_u"], scale=0.25, shape=8, log=TRUE) +
#
        dunif(params["detect_frac_0"], min=0, max=0.3, log=TRUE)
      if (log) f else exp(f)
```

```
# }
#
# covid_ga_model <- pomp(covid_ga_model, dprior = covid.dprior, paramnames=c("beta_d", "beta_red_factor")</pre>
```

Here we set up a pMCMC closely following the tutorial at https://kingaa.github.io/pomp/vignettes/oaxaca. html#particle_mcmc. Note: I tried to set the a gamma ditributed prior for γ_u but received errors using unnamed arguments, so switched to lognormal. As written, dmeas returning illegal values, perhaps pecause euler multnomial misspectified at lines 129 and 130.

```
registerDoRNG(03292020)
priorDens <- "
  lik = dlnorm(beta_d, log(2e-6), 1, 1) +
    dlnorm(beta_u,log(5e-8),1,1)+
    dlnorm(beta_e,log(5e-8),1,1)+
    dunif(beta_red_factor,0.01,1,1)+
    dlnorm(gamma_u, log(0.5), 1, 1)+
    dlnorm(gamma_u, log(0.5), 1, 1)+
    dunif(detect_frac_0, 0.01, 0.6, 1);
  if (!give_log) lik = exp(lik);
tic <- Sys.time()</pre>
                    # set a timer
# for the mcmc proposal subroutine
rw.sd <- rep(0.01, length(estpars))</pre>
                                        # set noise level in parameter random walk; used only when propo
names(rw.sd) <- estpars</pre>
n <- 10
          # number of mcmc chains
foreach(i = 1:n, .combine = c) %dopar% {
  pmcmc(pomp(mf1, dprior = Csnippet(priorDens),
             paramnames=c("beta_d", "beta_u", "beta_e", "beta_red_factor", "gamma_u", "gamma_d", "detec
        Nmcmc = 500, Np = 1000,
        #proposal = mun.rw.adaptive(rw.sd, scale.start=1, shape.start=1))
                                                                                  # alternate noise mode
        proposal = mvn.diag.rw(rw.sd))
                                            # alternate noise model
} -> pmcmc1
toc <- Sys.time()</pre>
print(toc-tic)
plot(pmcmc1)
```

The following is a placeholder for fiting via ABC. CODE NOT WORKING.

```
foreach (i = 1:10, .combine = c) %dopar% {
  theta.guess <- theta.true
  theta.guess[estpars] <- rlnorm(n = length(estpars),
      meanlog = log(theta.guess[estpars]), sdlog = 1)

# the following lines need to be rewritten for abc
mif2(covid_ga_model, Nmif = 50, params = theta.guess,
Np = 2000, cooling.fraction = 0.7,
rw.sd = rw.sd(beta_d = 0.02))</pre>
```

```
abc(pomp)
} -> abc
```

Here is a fully worked example from pomp that demonstrates MIF, synthetic likelihood (a la Wood 2010), and pMCMC.

```
loc <- url("https://kingaa.github.io/pomp/vignettes/parus.csv")</pre>
dat <- read.csv(loc)</pre>
head(dat)
parus <- pomp(dat,times="year",t0=1959)</pre>
stochStep <- Csnippet("</pre>
 N = r*N*exp(-c*N+rnorm(0,sigma));
")
pomp(
  parus,
  rprocess=discrete time(step.fun=stochStep,delta.t=1),
  rinit=Csnippet("N = N_0;"),
  paramnames=c("r","c","sigma","N_0"),
  statenames=c("N")
) -> parus
rmeas <- Csnippet("pop = rpois(phi*N);")</pre>
dmeas <- Csnippet("lik = dpois(pop,phi*N,give_log);")</pre>
pomp(parus,
     rmeasure=rmeas,
     dmeasure=dmeas,
     statenames=c("N"),
     paramnames=c("phi")
) -> parus
coef(parus) \leftarrow c(N_0=2,r=20,c=1,sigma=0.1,phi=200)
mif2(parus, Nmif=30, Np=1000,
     cooling.fraction.50=0.8, cooling.type="geometric",
     rw.sd=rw.sd(r=0.02,sigma=0.02,phi=0.02,N_0=ivp(0.1))
) -> mf
plot(mf)
pf <- replicate(5, pfilter(mf,Np=1000))</pre>
11 <- sapply(pf,logLik)</pre>
logmeanexp(11,se=TRUE)
probe(mf, nsim=200,
      probes=list(
        mean=probe.mean("pop"),
        sd=probe.sd("pop"),
        probe.acf("pop",transform=sqrt,lags=c(1,2)),
        probe.quantile("pop",prob=c(0.2,0.8))
      )) -> pb
```

```
plot(pb)
probe_objfun(pb, nsim=200, est = c("N_0","r"),
 seed = 669237763L) -> pm
subplex(par=coef(pm,c("N_0","r")),fn=pm) -> fit
pm(fit$par)
summary(pm)
priorDens <- "
 lik = dnorm(sigma, 0.2, 1, 1) +
    dnorm(phi,200,100,1)+
    dexp(r,0.1,1);
 if (!give_log) lik = exp(lik);
pmcmc(pomp(mf, dprior=Csnippet(priorDens),
           paramnames=c("sigma","phi","r")),
     Nmcmc = 500, Np = 1000,
     proposal = mvn.diag.rw(
       rw.sd=c(N_0=0.1, sigma=0.02, r=0.02, phi=0.02)
      )) -> pmh
plot(pmh,pars=c("loglik","log.prior","N_0","sigma","r","phi"))
```