

# Fit mutant with binomial aging model

*h qin*

*June 28 2017 - 31 May, 2018*

```
rm(list=ls())  
library('flexsurv')
```

```
## Loading required package: survival
```

```
library('stringr')  
source("../lifespan.r")
```

The RLS data used here are from Kaerberlein et al 2004, Plos Genetics <http://journals.plos.org/plosbiology/article?id=10.1371/journal.pbio.0020296>

The original data set is at <https://doi.org/10.1371/journal.pbio.0020296.sd002>

```
tb = read.csv("kaeberlein04_selected_strains.csv")  
  
my.strains = names(tb);  
report = data.frame(my.strains)  
report$samplesize = NA; report$R=NA; report$t0=NA; report$n=NA; report$G=NA; #report$longfilename=NA;  
#write.csv(tb, "kaeberlein04_selected_strains.csv", row.names = FALSE, quote = FALSE)
```

## Fit all RLS data sets by strains

```
for( i in 1:length(report[,1])){  
  my.data = tb[,i]  
  my.data = my.data[! is.na(my.data)]  
  report$samplesize[i] = length(my.data)  
  
  GompFlex = flexsurvreg(formula = Surv(tb[,i]) ~ 1, dist = 'gompertz')  
  WeibFlex = flexsurvreg(formula = Surv(tb[,i]) ~ 1, dist = 'weibull')  
  
  report$avgLS[i] = mean(tb[,i], na.rm=T)  
  report$stdLS[i] = sd(tb[,i], na.rm = T)  
  report$CV[i] = report$stdLS[i] / report$avgLS[i]  
  
  report$GompGFlex[i] = GompFlex$res[1,1]  
  report$GompRFlex[i] = GompFlex$res[2,1]  
  report$GompLogLikFlex[i] = round(GompFlex$loglik, 1)  
  report$GompAICFlex[i] = round(GompFlex$AIC)  
  
  report$WeibShapeFlex[i] = WeibFlex$res[1,1]  
  report$WeibRateFlex[i] = WeibFlex$res[2,1]  
  report$WeibLogLikFlex[i] = round(WeibFlex$loglik, 1)  
  report$WeibAICFlex[i] = round(WeibFlex$AIC)  
  
  #set initial values  
  Rhat = report$GompRFlex[i]; # 'i' was missing. a bug costed HQ a whole afternoon.  
  Ghat = report$GompGFlex[i];
```

```

nhat = 6;
t0= (nhat-1)/Ghat;
fitBinom = optim ( c(Rhat, t0, nhat), llh.binomialMortality.single.run,
  lifespan=tb[,i],
  #method='SANN') #SANN needs control
  method="L-BFGS-B",
  lower=c(1E-10, 1, 1), upper=c(1,200,20) );
report[i, c("R", "t0", "n")] = fitBinom$par[1:3]
report$G[i] = (report$n[i] - 1)/report$t0[i]
}

```

Show the results

report

```

##          my.strains samplesize      R      t0      n
## 1      fig4b.BY4742.2glucose      60 0.002388227 35.59111 7.703055
## 2 fig4b.by4742.SIR2.ox.2glucose      60 0.003068054 66.82089 8.067874
## 3          fig2a.sir2      90 0.002656909 16.80162 8.109348
## 4          fig2b.sir2      90 0.002656909 16.80162 8.109348
## 5          fig1b.BY4742      250 0.004796452 58.10834 8.109395
## 6          fig1b.fob1      140 0.003056547 71.37470 7.695271
## 7          fig1b.hxk2      120 0.005736777 103.59300 7.578456
## 8          fig1b.fob1.hxk2      160 0.005902597 120.65937 6.023612
##      G      avgLS      stdLS      CV GompGFlex GompRFlex
## 1 0.18833511 26.06667 7.557389 0.2899254 0.13946781 0.002283890
## 2 0.10577341 34.60000 10.834972 0.3131495 0.07468834 0.004042203
## 3 0.42313480 13.96667 3.491402 0.2499810 0.27970246 0.003555048
## 4 0.42313480 13.96667 3.491402 0.2499810 0.27970246 0.003555048
## 5 0.12234725 26.62400 9.418144 0.3537464 0.08544660 0.006577615
## 6 0.09380453 37.75000 13.405746 0.3551191 0.06987148 0.003424637
## 7 0.06350290 36.73333 16.338197 0.4447785 0.04823602 0.006725196
## 8 0.04163466 48.28125 21.337707 0.4419460 0.04143860 0.004249514
##      GompLogLikFlex GompAICFlex WeibShapeFlex WeibRateFlex WeibLogLikFlex
## 1      -206.8      418      3.925269      28.75636      -206.2
## 2      -235.6      475      3.369961      38.51039      -227.8
## 3      -246.0      496      4.431406      15.29449      -240.4
## 4      -246.0      496      4.431406      15.29449      -240.4
## 5      -937.3     1879      3.024640      29.77138      -914.0
## 6      -565.5     1135      3.139805      42.25834      -559.6
## 7      -507.6     1019      2.397501      41.42400      -502.2
## 8      -713.6     1431      2.441562      54.41078      -714.3
##      WeibAICFlex
## 1      416
## 2      460
## 3      485
## 4      485
## 5     1832
## 6     1123
## 7     1008
## 8     1433

```

Output

```
write.csv(report, file = 'sandbox/_report_kaeberlein04_fit.csv', row.names = FALSE)
```

## Overlay histogram with probability density function of binomial aging model

see <http://hongqinlab.blogspot.com/2013/12/binomial-mortality-model.html>

$$m = R \left( 1 + t/t_0 \right)^{(n-1)} s = \exp \left( (R t_0/n) (1 - (1+t/t_0)^n) \right)$$

$$pdf = sm$$

It seems binomial model aging is a reasonable fit whenevern Gompertz model is a reasonable fit.

```
# (x,y) are legend positions for plots
x = c(0, 0, 0, 0, 40, 55, 50, 5);
y = c(0.06, 0.06, 0.1, 0.1, 0.04, 0.03, 0.03, 0.03);

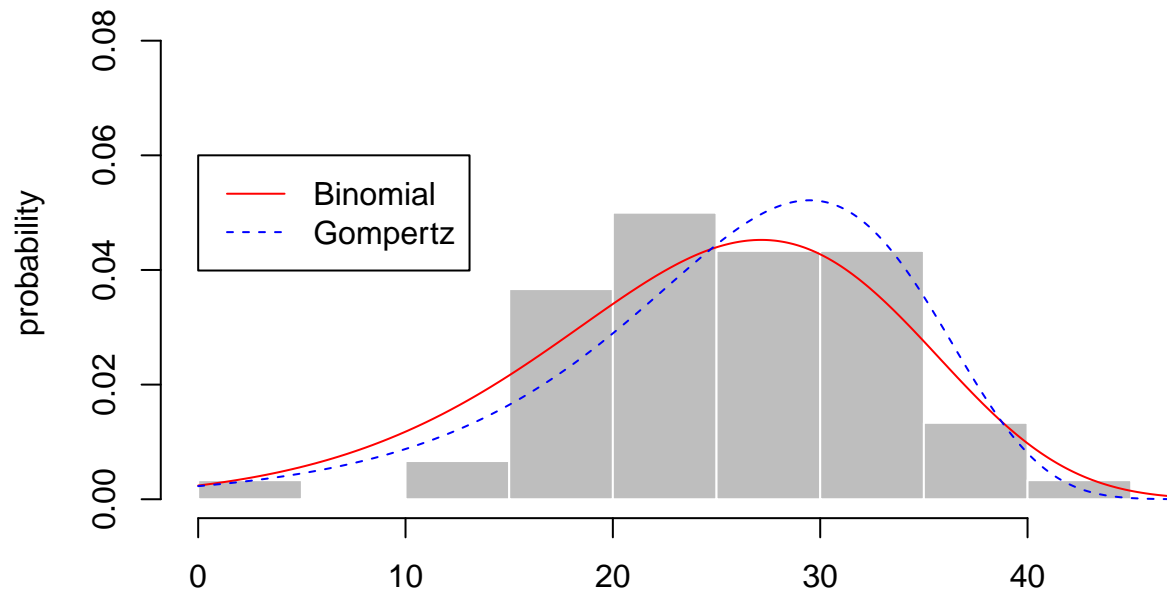
for( i in 1:length(report[,1])){
  my.data = tb[,i]
  my.data = my.data[! is.na(my.data)]

  h= hist(my.data, br=max(my.data)/2, plot = FALSE);
  hist(my.data, probability = TRUE, col='gray', border='white', xlab='RLS', ylab='probability',
        main=my.strains[i], ylim=c(0, max(h$density)*1.1), xlim=c(0, max(h$mids)*1.1))
  #plot( h$density ~ h$mids, main=my.strains[i], xlab="RLS", ylab="density")
  #par(new=TRUE);
  t= seq(0, max(h$mids)*1.2, by=0.1)
  s = exp( (report$R[i]* report$t0[i]/report$n[i])*(1 - (1+t/report$t0[i])^report$n[i]) ) );
  m = report$R[i]*(1 + t/ report$t0[i])^(report$n[i] -1 )
  pdf = s*m
  lines( pdf ~ t, col='red', lty=1)

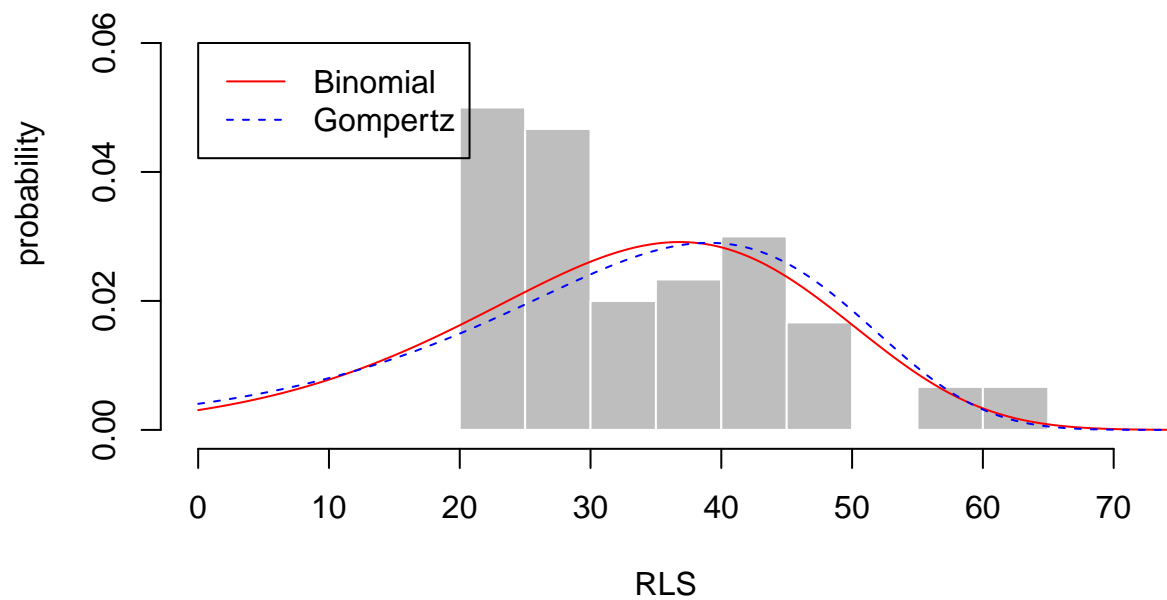
  s.g = G.s( c(report$GompRFlex[i], report$GompGFlex[i]), t );
  m.g = report$GompRFlex[i]*exp(report$GompGFlex[i]*t)
  pdf.g = s.g * m.g
  lines( pdf.g ~ t, col="blue", lty=2)

  legend ( x[i], y[i], c("Binomial", "Gompertz"), col=c("red", "blue"), lty=c(1,2) )
}
```

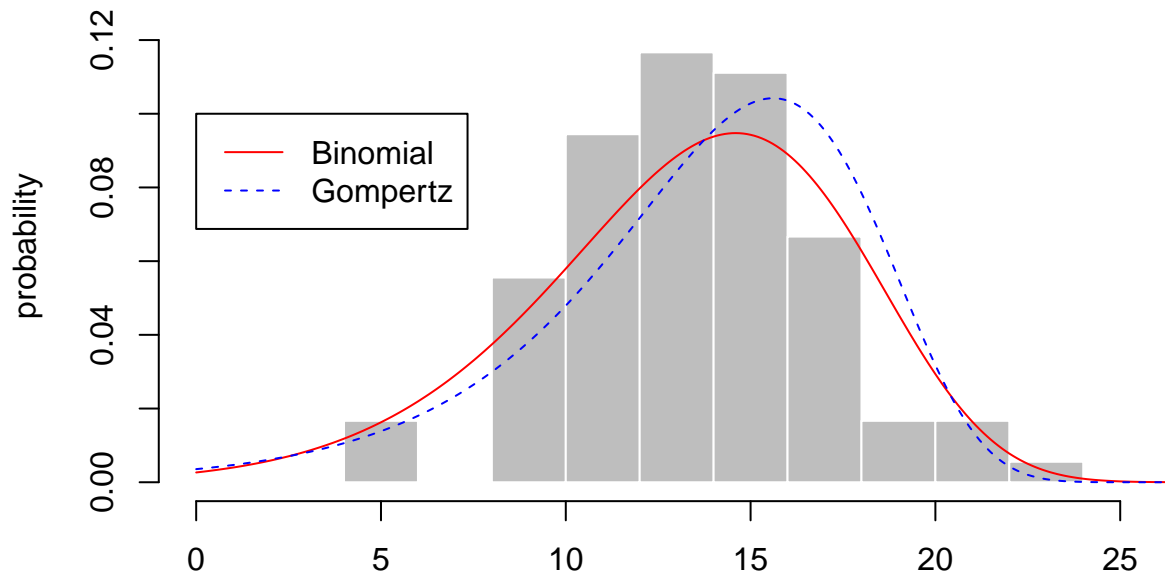
**fig4b.BY4742.2glucose**



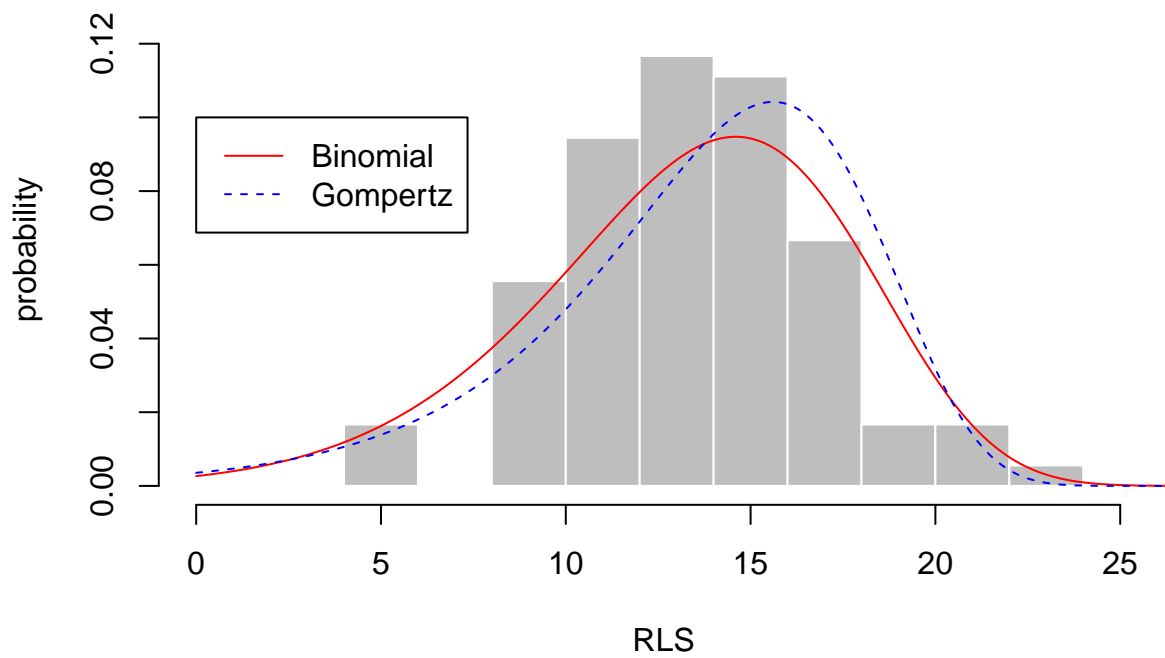
**fig4b.by4742.SIR2.ox.2glucose**



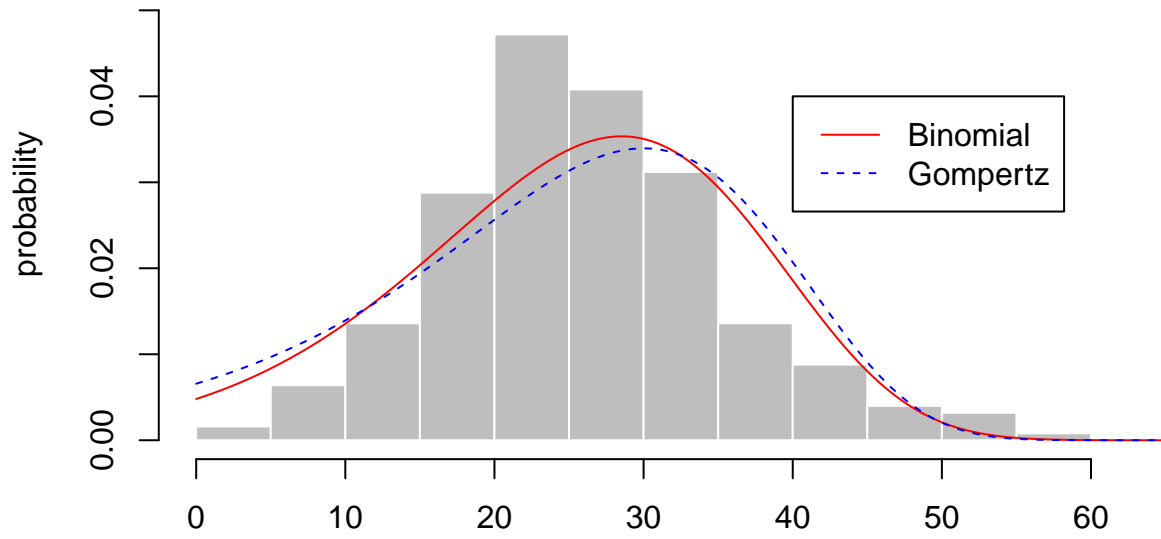
**fig2a.sir2**



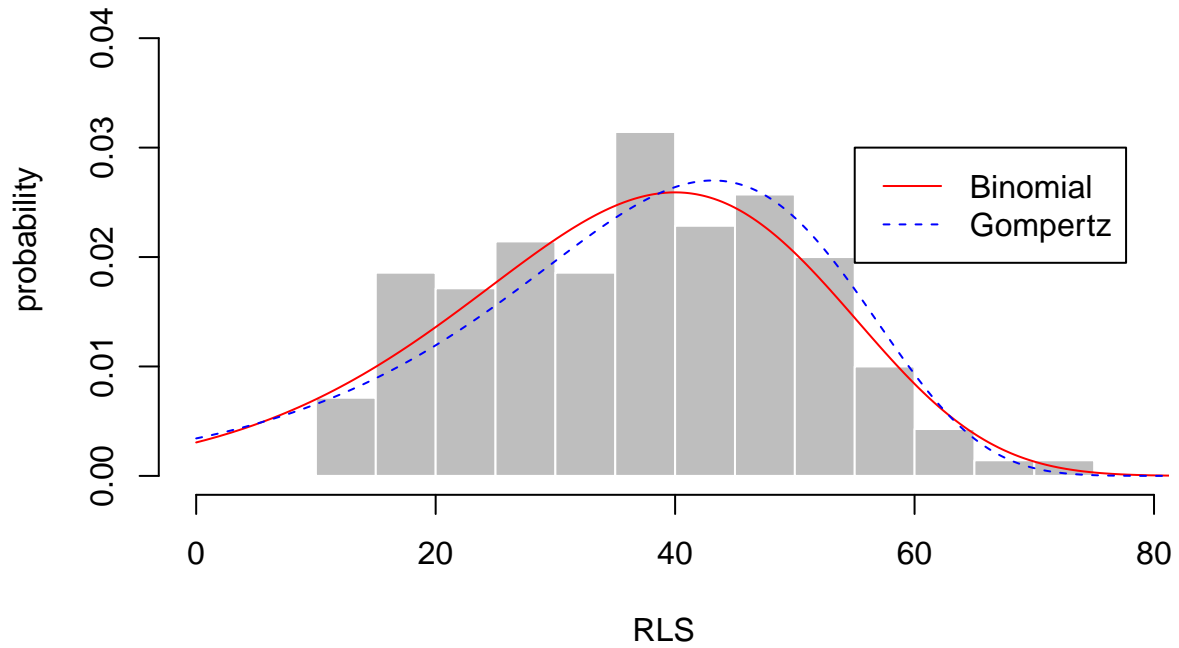
RLS  
**fig2b.sir2**



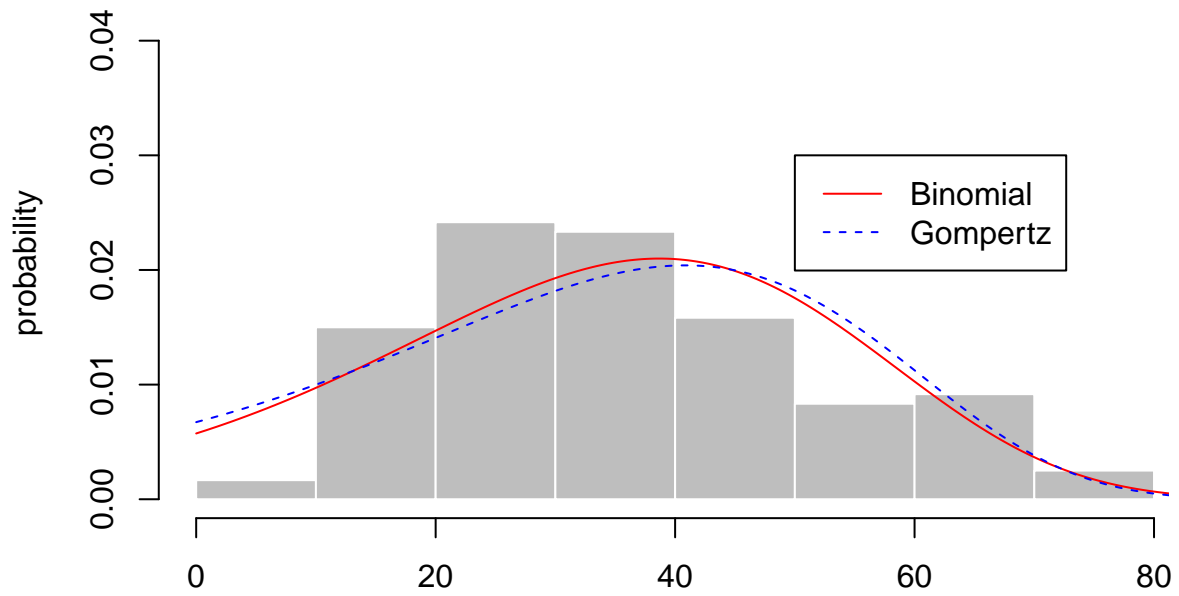
**fig1b.BY4742**



RLS  
**fig1b.fob1**



**fig1b.hxk2**



RLS  
**fig1b.fob1.hxk2**

