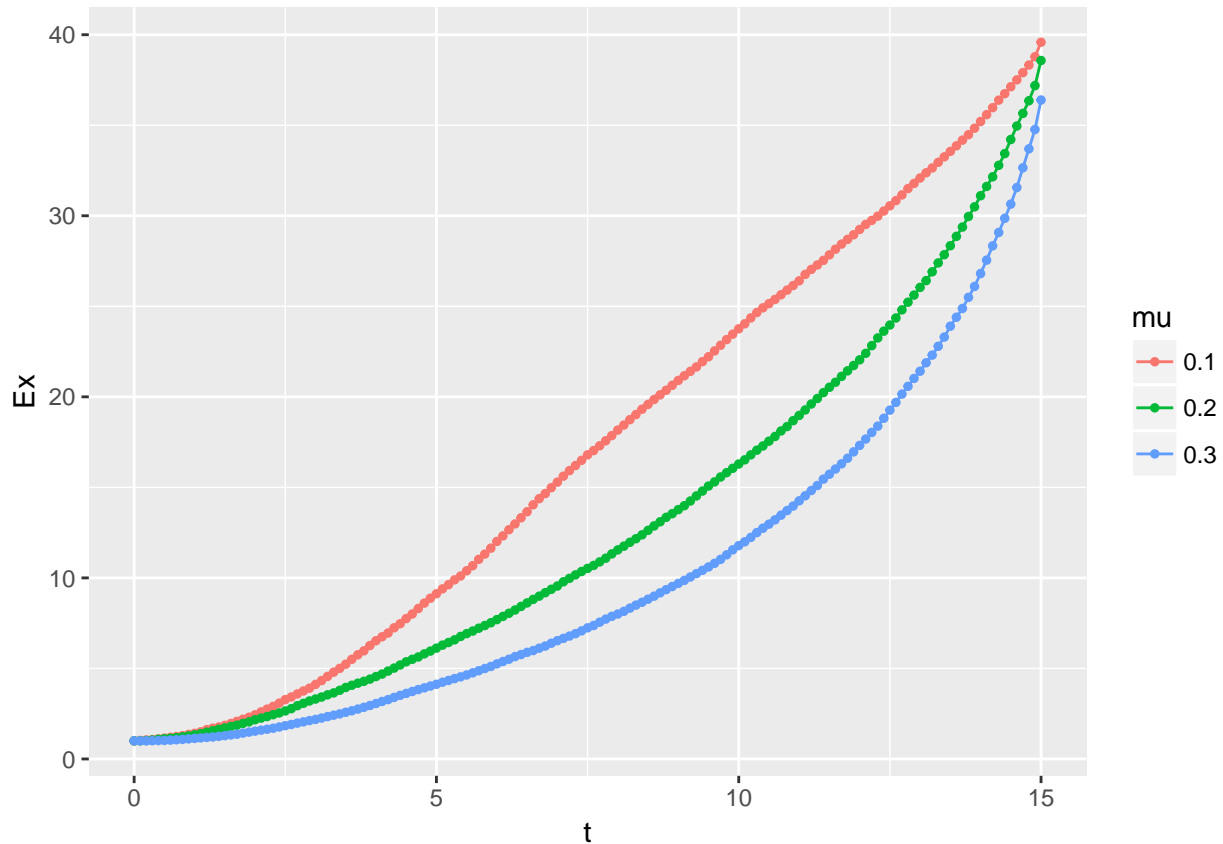


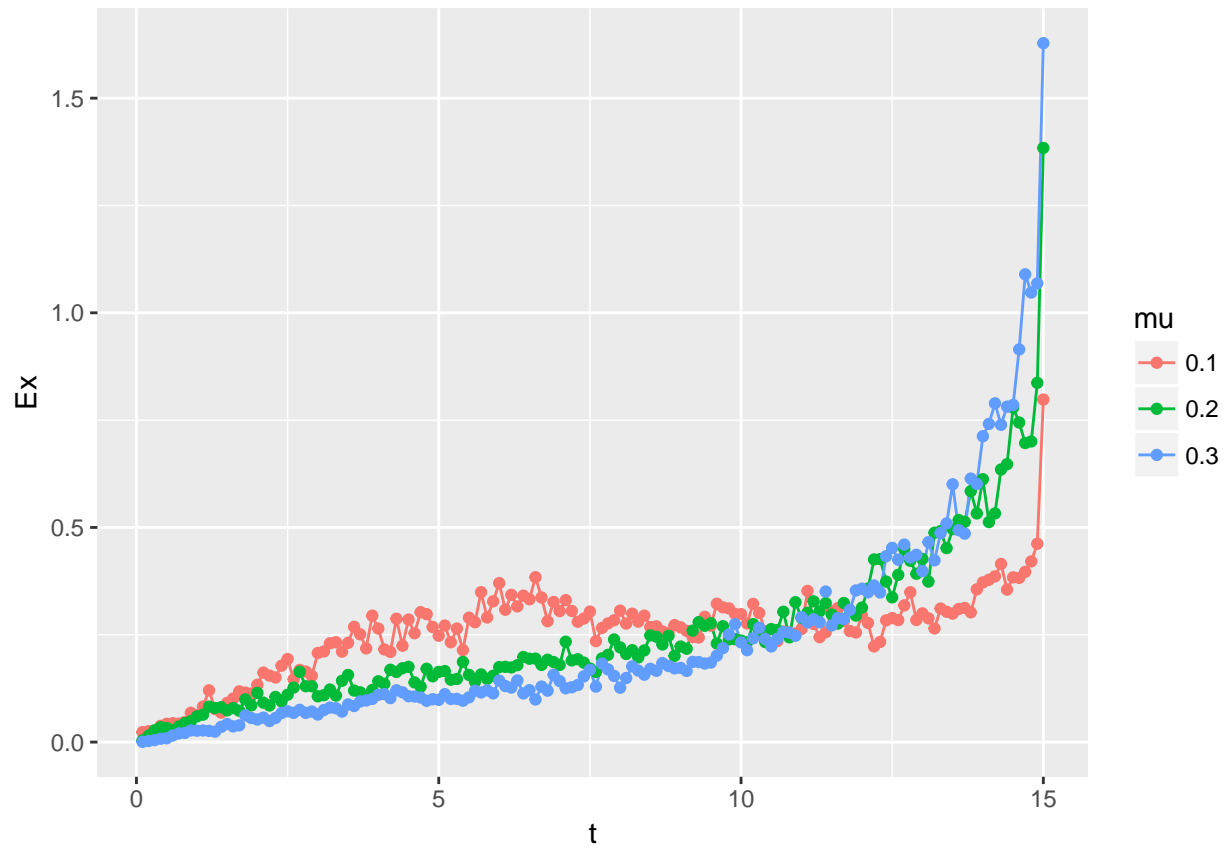
What extant species can tell us about extinction?

Because extinct species are rarely included on phylogenetic trees, we are interested on investigate the information that extant species contains about extinction rates. On the plot below we can see the expected Ltt plot, of extant-species only trees, for 3 different extinction rates

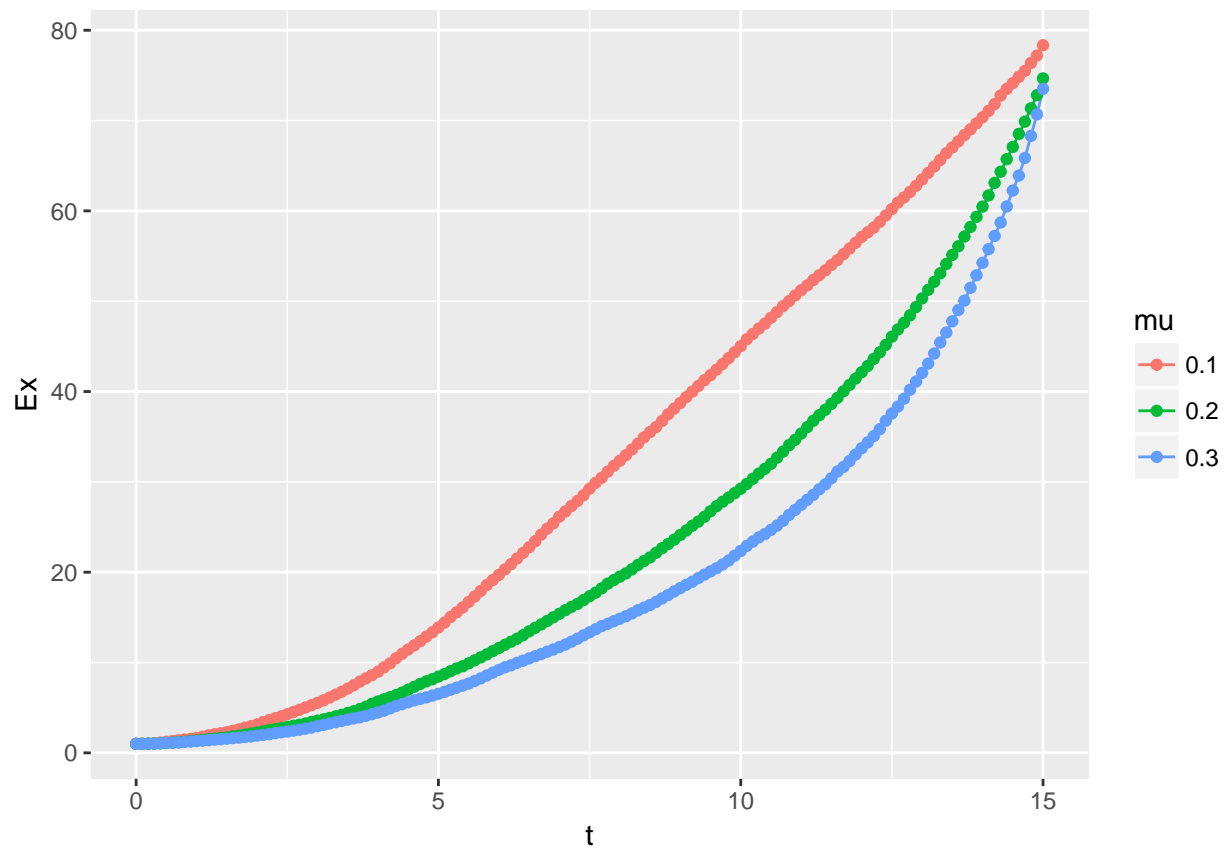


we can see a clear difference on Ltt plots of extant species, smaller extinction rates tends to grow faster on the beginning whereas higher extinction rate seems to have a slow grow on the beginning.

It seems also that is a matter of the first derivate, we can look at that also

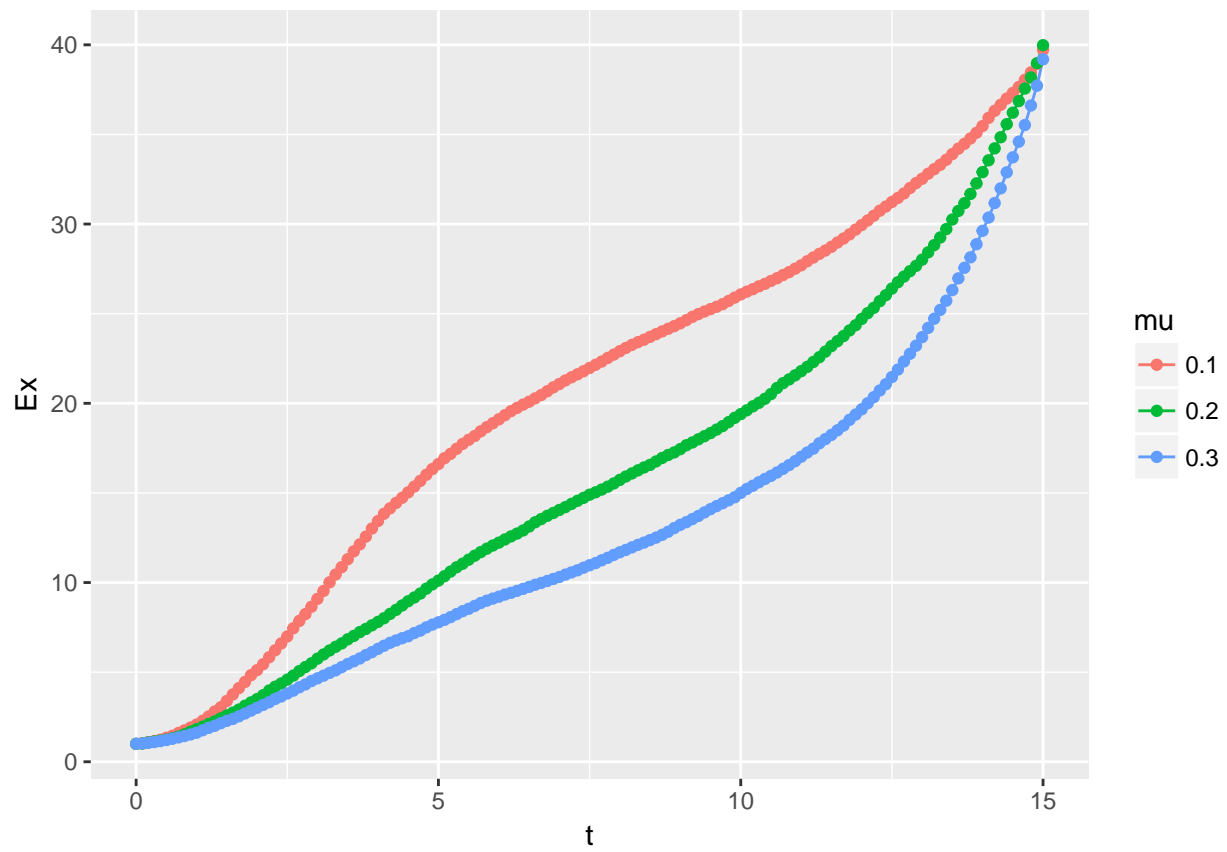


Now we do it again, but with $K = 80$ rather than $K = 40$ in order to check some influence on the K parameter

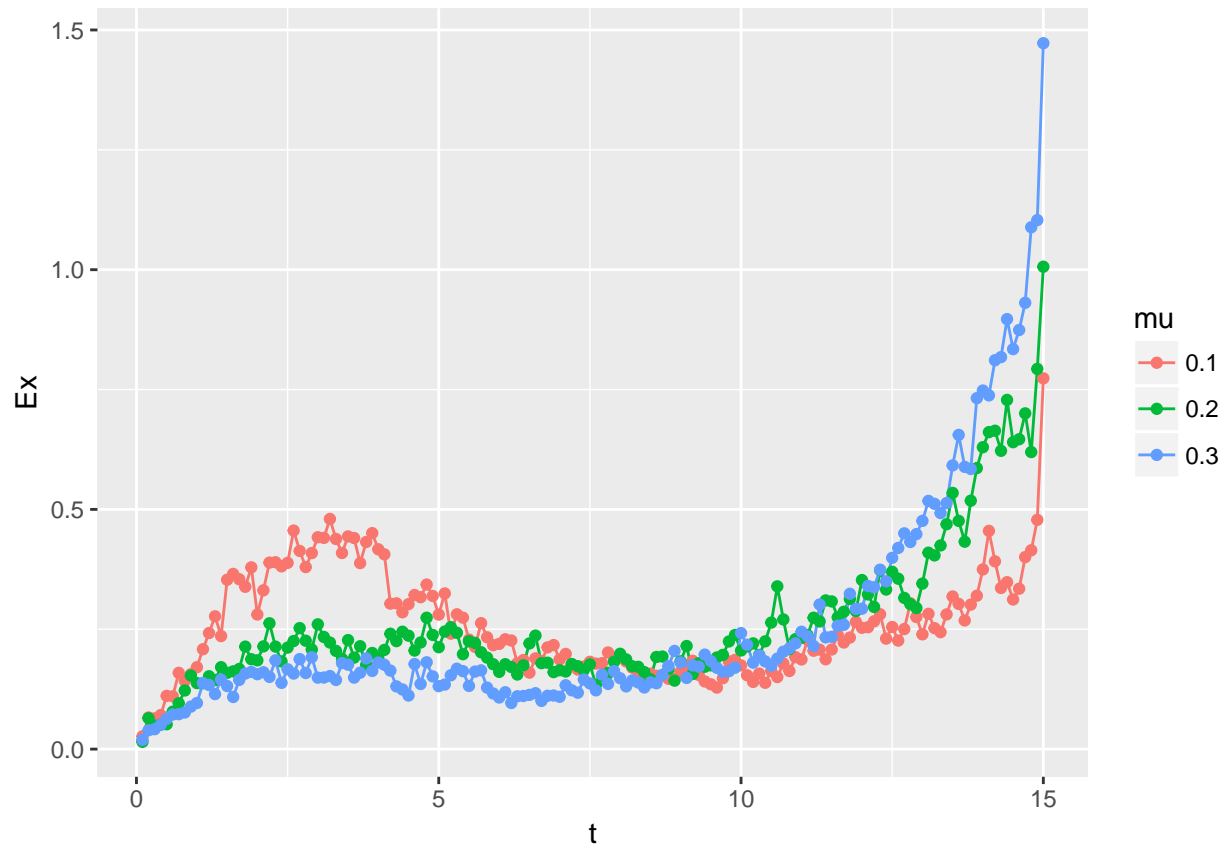


It seems it is just a change on scale. Now, what about λ ?

We set $\lambda = 1.2$ rather than $\lambda = 0.8$ and we see again the ltt plot

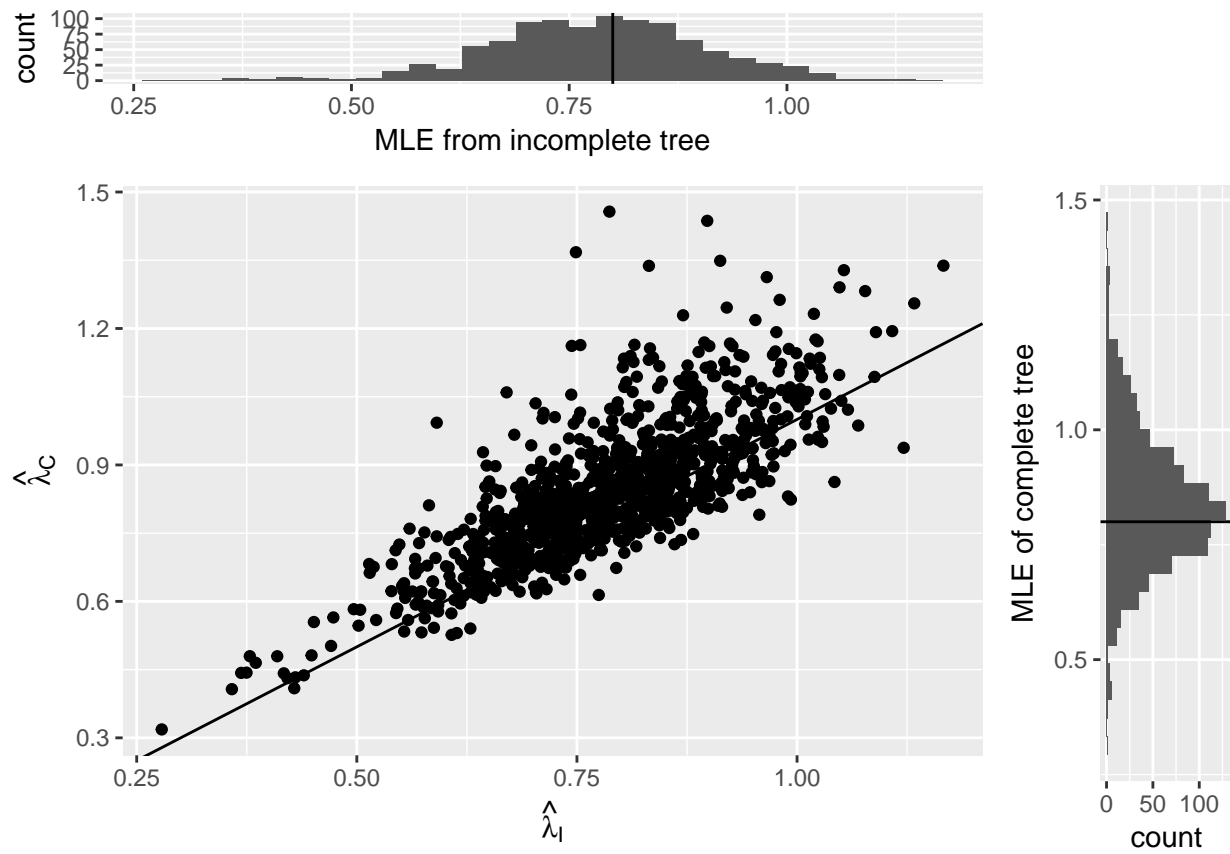


and we check the derivative again

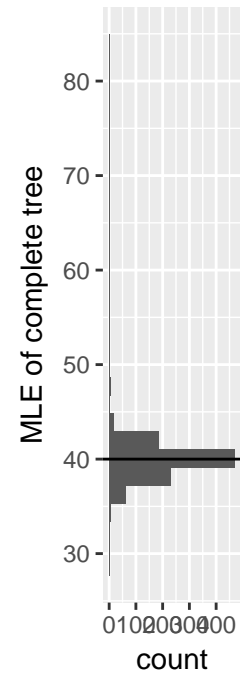
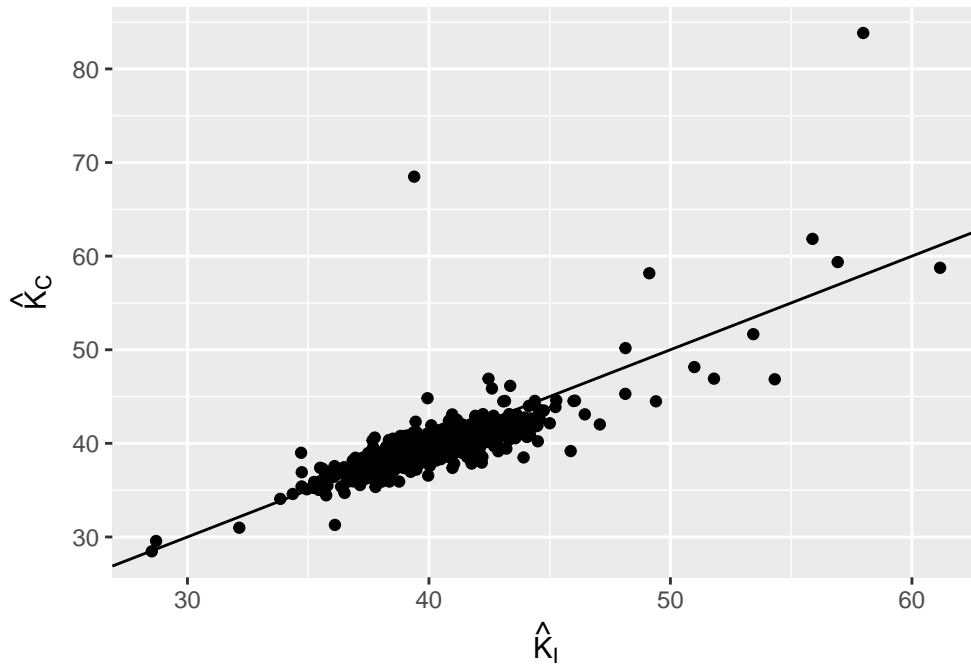
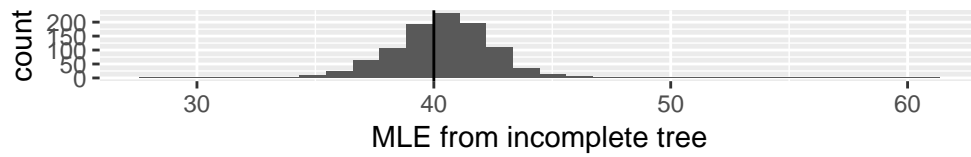


2 parameter estimation (fixing μ)

```
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
```



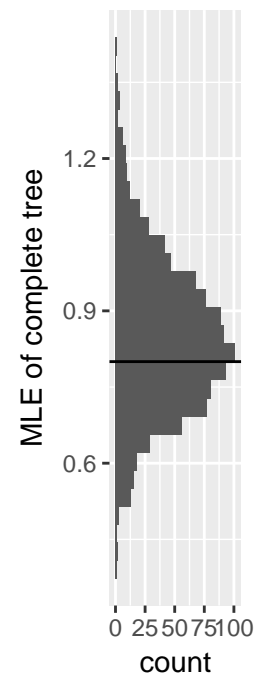
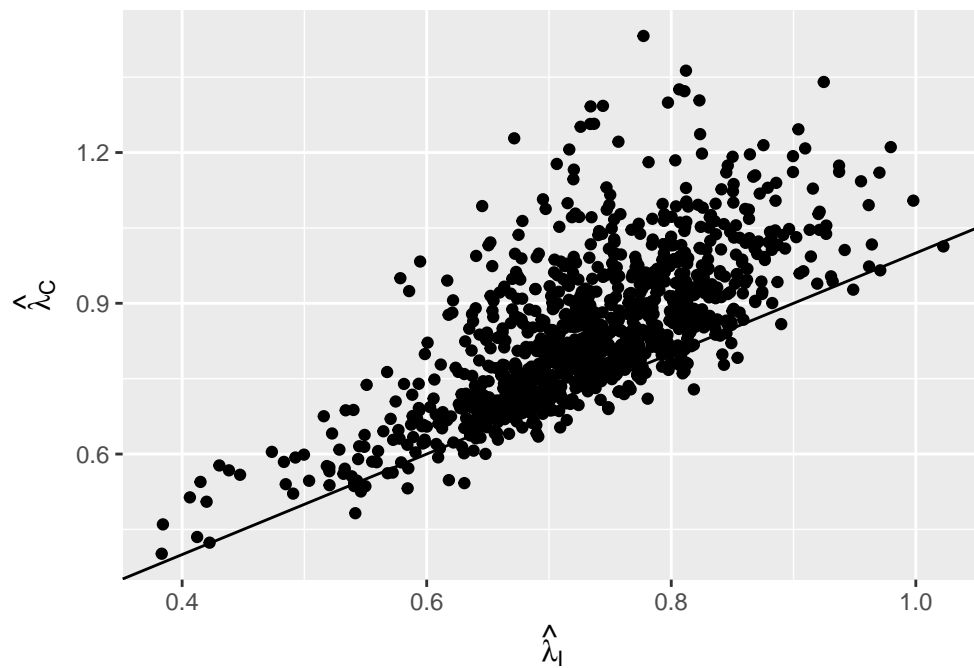
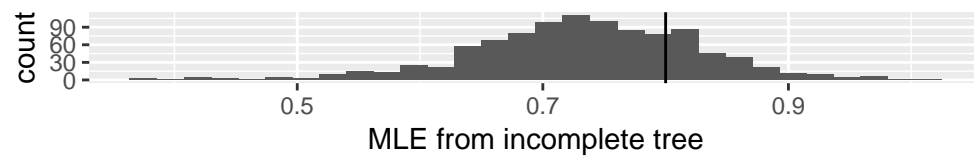
```
## [1] "0.007000000000000001 proportion of data was excluded for vizualization purposes"
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
```



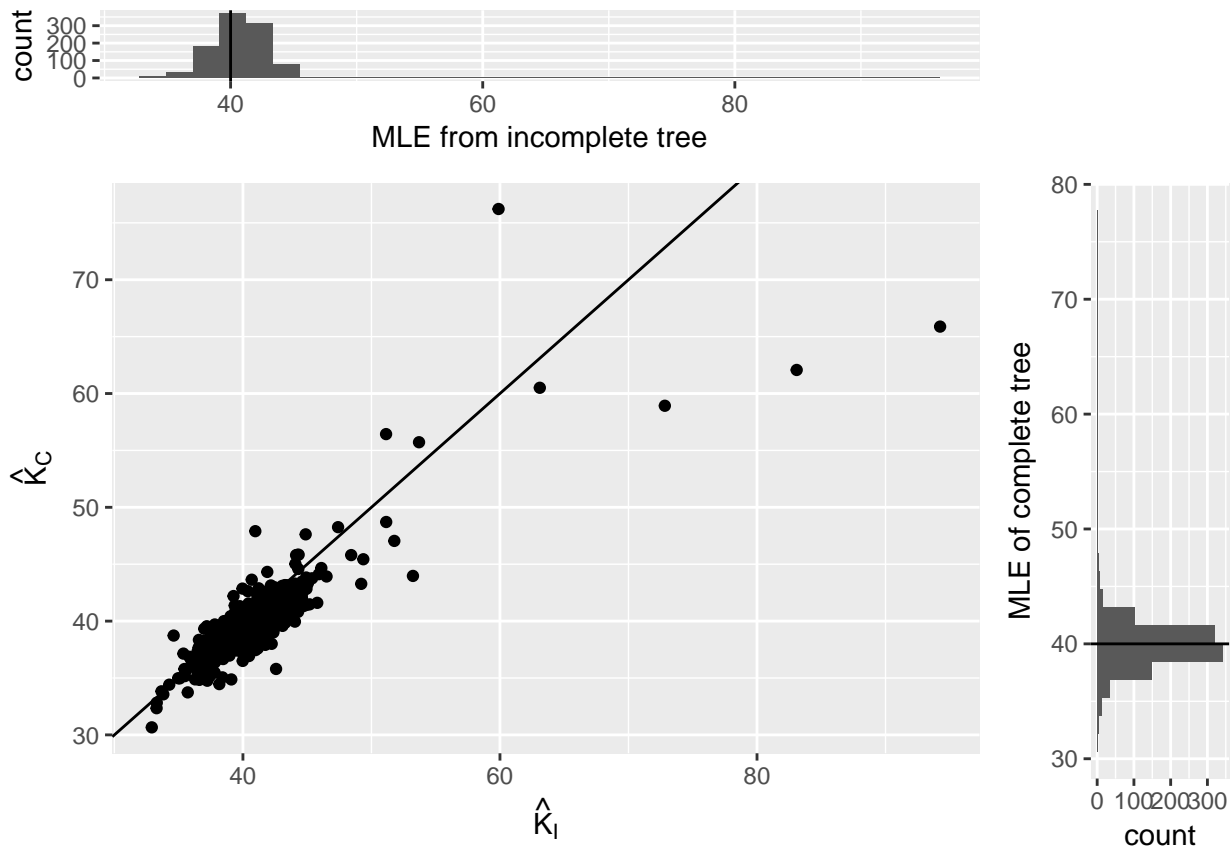
```
## Warning in proc.time() - p: longer object length is not a multiple of
## shorter object length

##      user      system    elapsed
## 366.612742 -0.148922 330.979588

## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
```



```
## [1] "0.005 proportion of data was excluded for vizualization purposes"
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
```

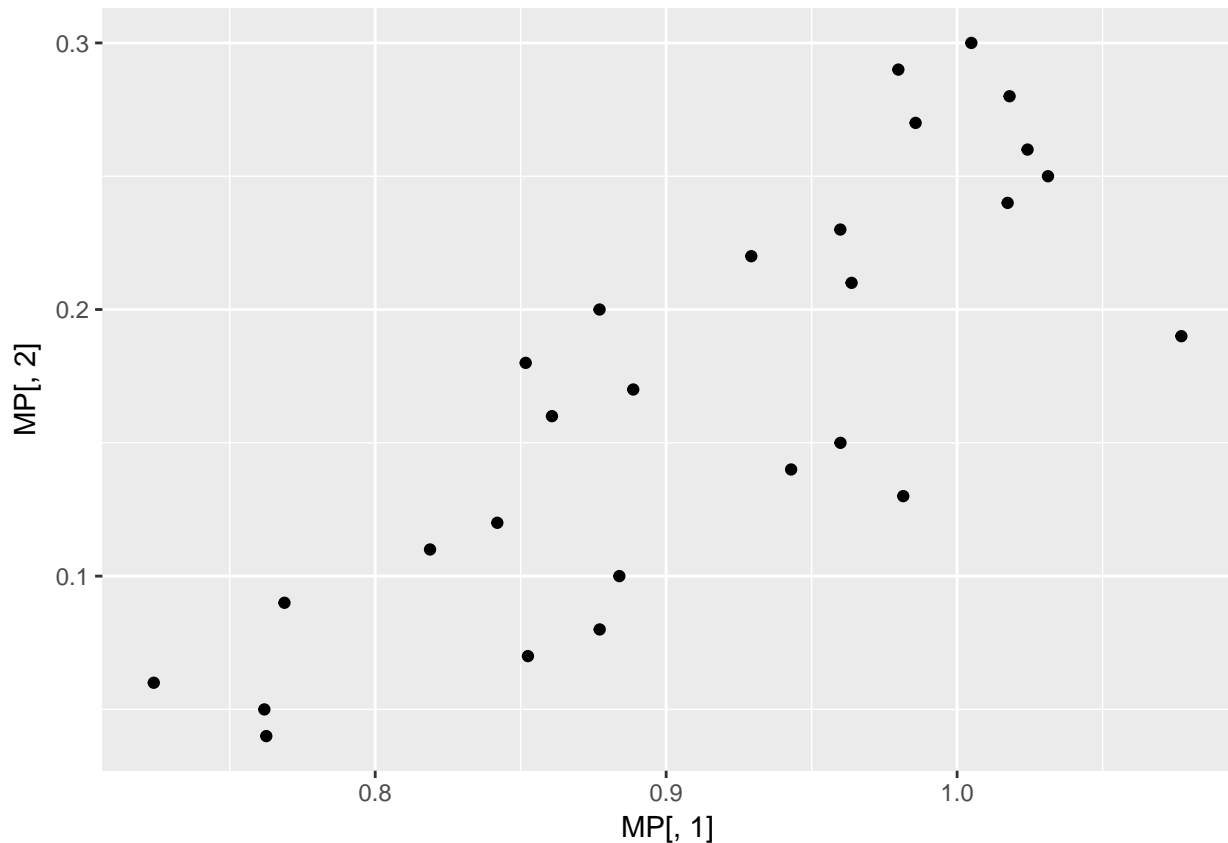
```
## Warning in proc.time() - p: longer object length is not a multiple of
## shorter object length
```

```
##          user      system    elapsed
## 2244.75775482   -0.09718191 2206.84986884
```

Ok. The estimations are fine.

Now let's try for a grid of μ

```
mu0 = seq(0.04,0.3,by=0.01)
s = sim_phyl(seed=3)
p <- subplex(par = c(2,0.2,60), fn = llik, n = s$n, E = s$E, t = s$wt)$par
wt = (s$newick.extant.p)$wt
MP = matrix(nrow=length(mu0), ncol=3)
n_trees = 10
for(i in 1:length(mu0)){
  mu = mu0[i]
  trees = sim_srt(wt=wt, pars=c(p[1],mu,p[3]), parallel = F, n_trees = n_trees)
  pars = subplex(par = c(2,60), fn = llik_st, setoftrees = trees, mu = mu, impSAM = FALSE)$par
  MP[i,] = c(pars[1],mu,pars[2])
}
qplot(MP[,1],MP[,2])
```



```
lm1 = lm(MP[,2] ~ MP[,1])
summary(lm1)
```

```
##
## Call:
## lm(formula = MP[, 2] ~ MP[, 1])
##
## Residuals:
```

	Min	1Q	Median	3Q	Max
	-0.090752	-0.039528	0.005664	0.033584	0.074825

```
##
## Coefficients:
```

	Estimate	Std. Error	t value	Pr(> t)
(Intercept)	-0.44537	0.08967	-4.967	4.06e-05 ***
MP[, 1]	0.67416	0.09773	6.898	3.13e-07 ***

```
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.0475 on 25 degrees of freedom
## Multiple R-squared:  0.6556, Adjusted R-squared:  0.6418
## F-statistic: 47.59 on 1 and 25 DF,  p-value: 3.131e-07

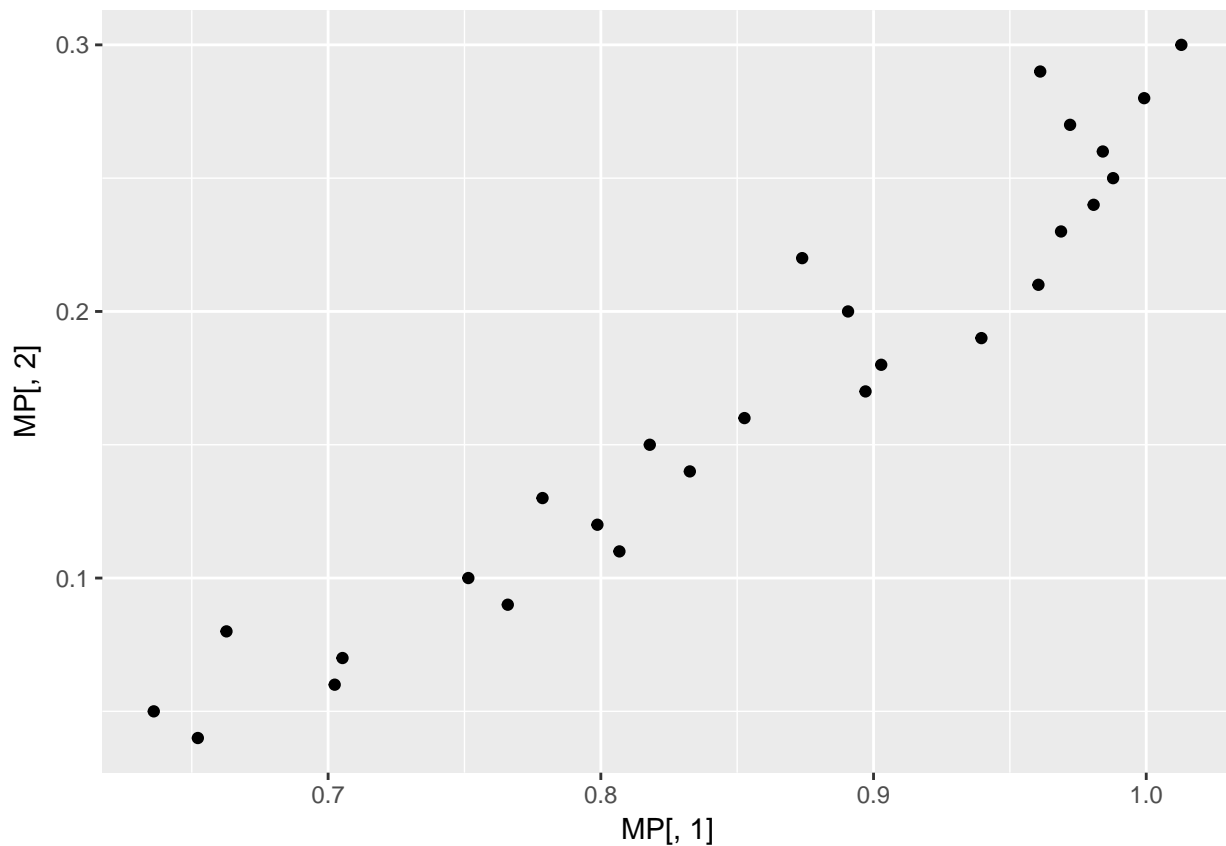
l1 = MP[,1]
m1 = MP[,2]
k1 = MP[,3]
MP1=MP
```

Does it help 100 trees (probably not)

```

mu0 = seq(0.04,0.3,by=0.01)
s = sim_phyl(seed=3)
p <- subplex(par = c(2,0.2,60), fn = llik, n = s$n, E = s$E, t = s$wt)$par
wt = (s$newick.extant.p)$wt
MP = matrix(nrow=length(mu0), ncol=3)
n_trees = 100
for(i in 1:length(mu0)){
  mu = mu0[i]
  trees = sim_srt(wt=wt, pars=c(p[1],mu,p[3]), parallel = F, n_trees = n_trees)
  pars = subplex(par = c(2,60), fn = llik_st, setoftrees = trees, mu = mu, impSam = FALSE)$par
  MP[i,] = c(pars[1],mu,pars[2])
}
qplot(MP[,1],MP[,2])

```



```

lm2 = lm(MP[,2] ~ MP[,1])
summary(lm2)

```

```

##
## Call:
## lm(formula = MP[, 2] ~ MP[, 1])
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -0.034392 -0.014381 -0.003059  0.013309  0.051681
##
## Coefficients:
##              Estimate Std. Error t value Pr(>|t|)

```

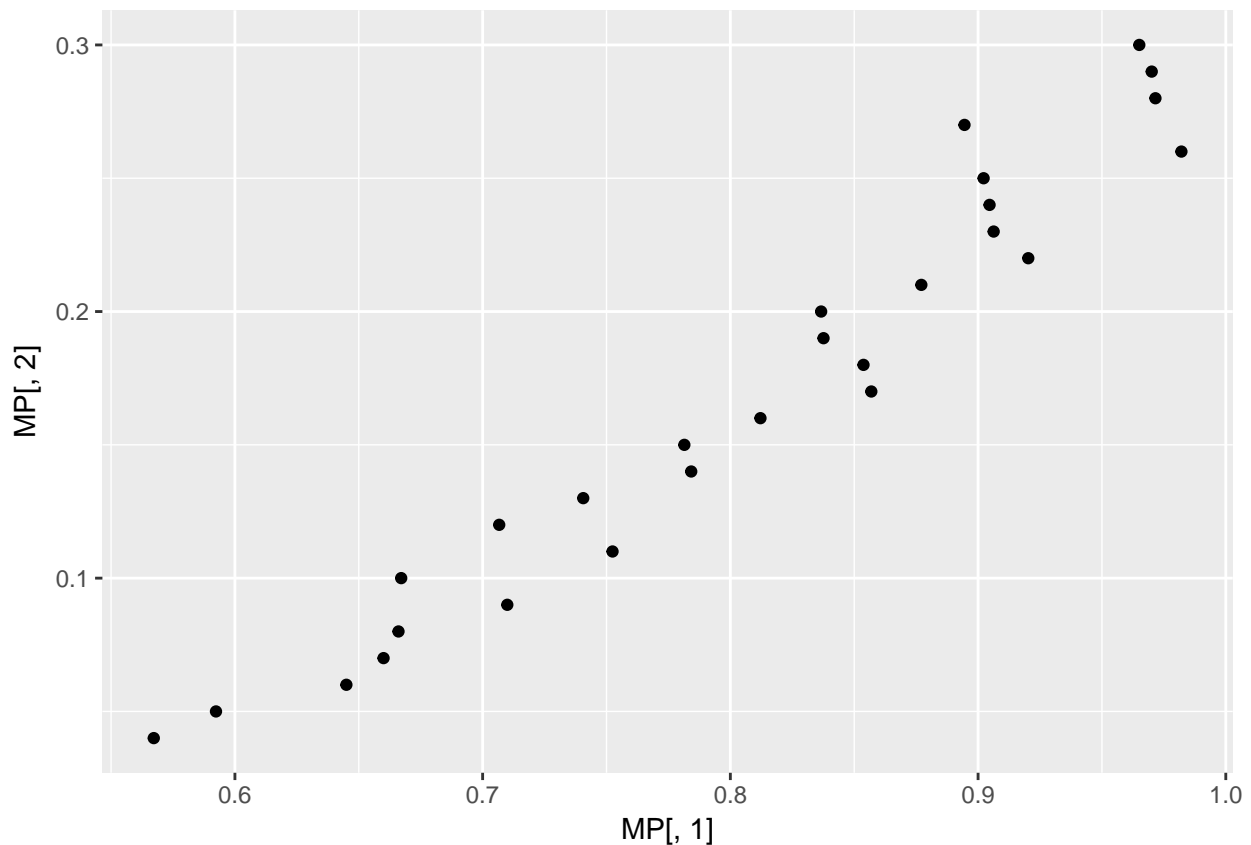
```
## (Intercept) -0.38250    0.03282   -11.65  1.34e-11 ***
## MP[, 1]      0.64593    0.03802    16.99  3.05e-15 ***
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.02285 on 25 degrees of freedom
## Multiple R-squared:  0.9203, Adjusted R-squared:  0.9171
## F-statistic: 288.6 on 1 and 25 DF,  p-value: 3.053e-15

l2 = MP[,1]
m2 = MP[,2]
k2 = MP[,3]
MP2=MP
```

Actually, the variance decreases.

1000 trees?

```
mu0 = seq(0.04,0.3,by=0.01)
s = sim_phyl(seed=3)
p <- subplex(par = c(2,0.2,60), fn = llik, n = s$n, E = s$E, t = s$wt)$par
wt = (s$newick.extant.p)$wt
MP = matrix(nrow=length(mu0), ncol=3)
n_trees = 1000
for(i in 1:length(mu0)){
  mu = mu0[i]
  trees = sim_srt(wt=wt, pars=c(p[1],mu,p[3]), parallel = F, n_trees = n_trees)
  pars = subplex(par = c(2,60), fn = llik_st , setoftrees = trees, mu = mu, impSAM = FALSE)$par
  MP[i,] = c(pars[1],mu,pars[2])
}
qplot(MP[,1],MP[,2])
```



```
lm3 = lm(MP[,2] ~ MP[,1])
summary(lm3)
```

```
##
## Call:
## lm(formula = MP[, 2] ~ MP[, 1])
##
## Residuals:
```

	Min	1Q	Median	3Q	Max
	-0.032094	-0.015010	-0.001572	0.013852	0.044157

```
##
## Coefficients:
```

	Estimate	Std. Error	t value	Pr(> t)
(Intercept)	-0.33899	0.02463	-13.76	3.6e-13 ***
MP[, 1]	0.63144	0.03022	20.89	< 2e-16 ***

```
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.01884 on 25 degrees of freedom
## Multiple R-squared:  0.9458, Adjusted R-squared:  0.9437
## F-statistic: 436.6 on 1 and 25 DF,  p-value: < 2.2e-16
```

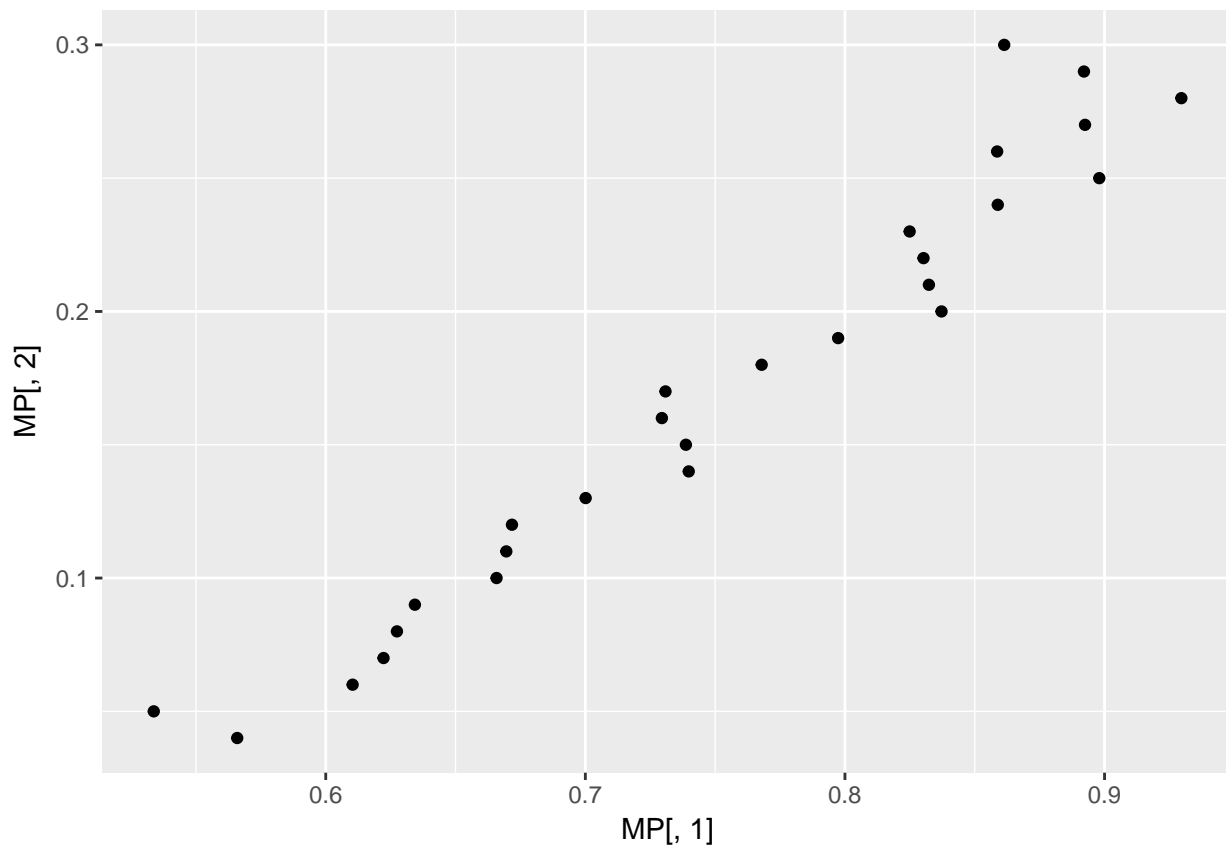
```
l3 = MP[,1]
m3 = MP[,2]
```

10000 trees?

```

mu0 = seq(0.04,0.3,by=0.01)
s = sim_phyl(seed=3)
p <- subplex(par = c(2,0.2,60), fn = llik, n = s$n, E = s$E, t = s$wt)$par
wt = (s$newick.extant.p)$wt
MP = matrix(nrow=length(mu0), ncol=3)
n_trees = 10000
for(i in 1:length(mu0)){
  mu = mu0[i]
  trees = sim_srt(wt=wt, pars=c(p[1],mu,p[3]), parallel = F, n_trees = n_trees)
  pars = subplex(par = c(2,60), fn = llik_st, setoftrees = trees, mu = mu, impSam = FALSE)$par
  MP[i,] = c(pars[1],mu,pars[2])
}
qplot(MP[,1],MP[,2])

```



```

lm4 = lm(MP[,2] ~ MP[,1])
summary(lm4)

```

```

##
## Call:
## lm(formula = MP[, 2] ~ MP[, 1])
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -0.028335 -0.010639 -0.003290  0.005892  0.055001
##
## Coefficients:
##              Estimate Std. Error t value Pr(>|t|)

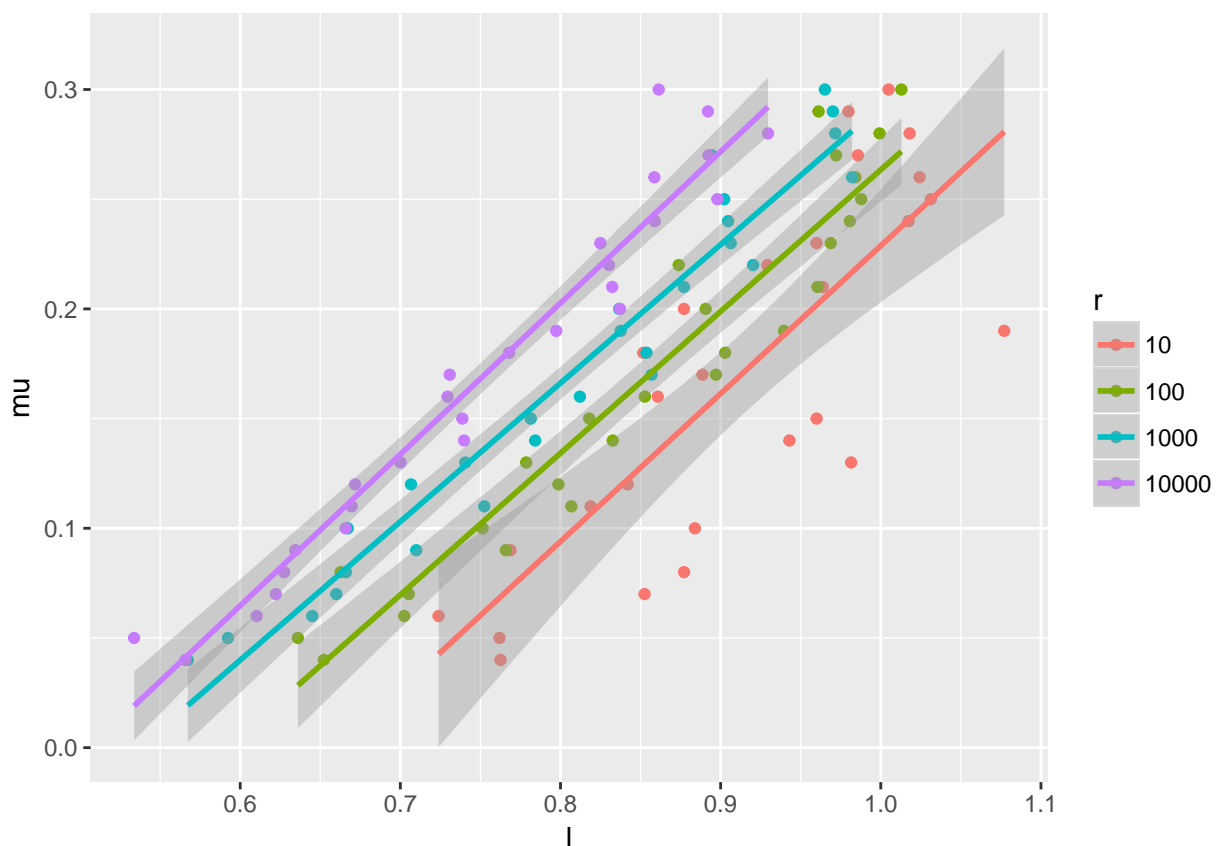
```

```
## (Intercept) -0.34923    0.02364   -14.77  7.43e-14 ***
## MP[, 1]      0.68988    0.03108    22.19  < 2e-16 ***
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.01779 on 25 degrees of freedom
## Multiple R-squared:  0.9517, Adjusted R-squared:  0.9498
## F-statistic: 492.6 on 1 and 25 DF,  p-value: < 2.2e-16
```

```
l4 = MP[,1]
m4 = MP[,2]
```

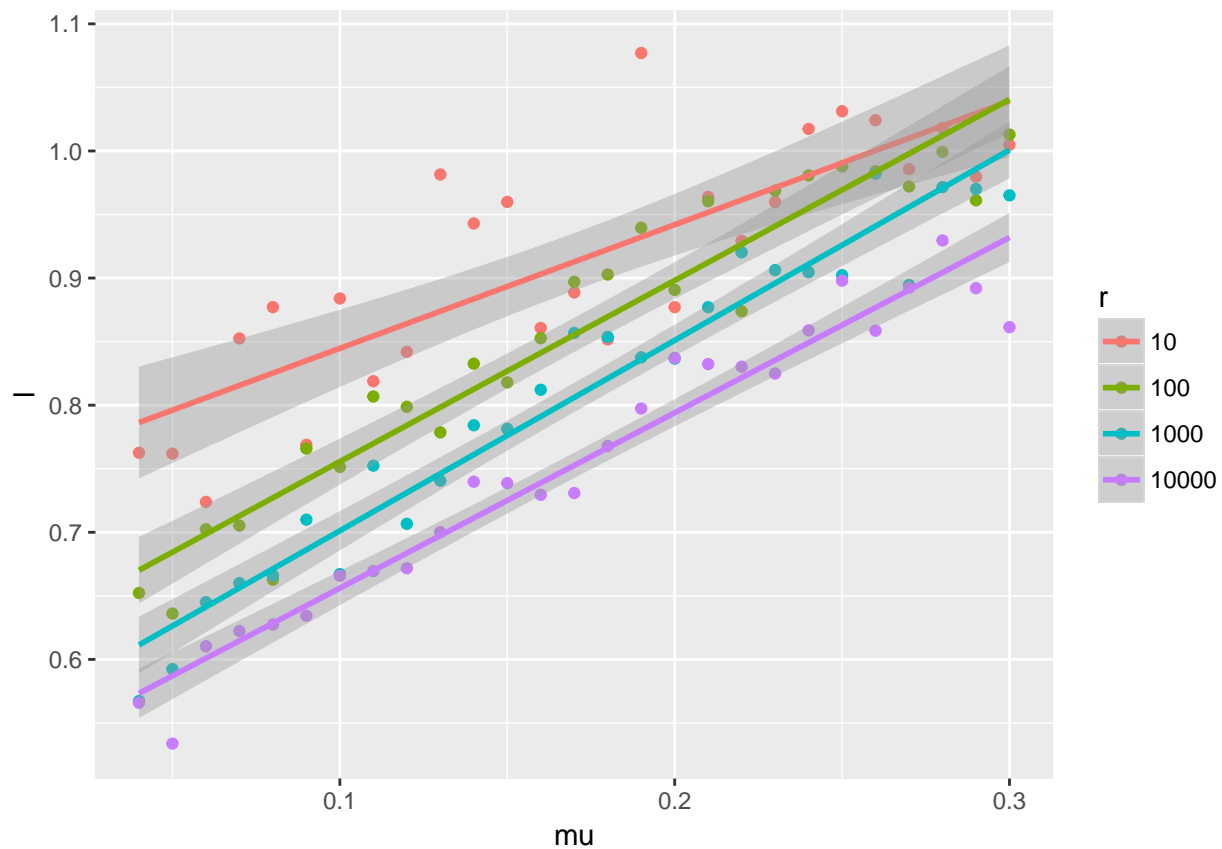
let's look at them all together

```
regressions = data.frame(mu = rep(m1,4), l = c(l1,l2,l3,l4), r = c(rep(x = 10,length(m1)),rep(x = 100,l2),rep(x = 1000,l3),rep(x = 10000,l4)),
regressions$r = as.factor(regressions$r)
ggplot(data=regressions, aes(x=l,y=mu,colour=r)) + geom_point() + geom_smooth(method='lm')
```



Just for convenience: μ vs λ

```
ggplot(data=regressions, aes(x=mu,y=l,colour=r)) + geom_point() + geom_smooth(method='lm')
```



get coefficients

```
lm11 = lm(l1~m1)
lm22 = lm(l2~m2)
lm33 = lm(l3~m3)
lm44 = lm(l4~m4)
lm11$coef
```

```
## (Intercept)      m1
##   0.747476   0.972451
```

```
lm22$coef
```

```
## (Intercept)      m2
##   0.6131469   1.4247474
```

```
lm33$coef
```

```
## (Intercept)      m3
##   0.5514301   1.4979030
```

```
lm44$coef
```

```
## (Intercept)      m4
##   0.5181201   1.3795075
```

```
lm11$coef[1] + lm11$coef[2]*0.1
```

```
## (Intercept)
##   0.8447211
```



```
lm22$coef[1] + lm22$coef[2]*0.1
```

```
## (Intercept)  
## 0.7556216
```

```
lm33$coef[1] + lm33$coef[2]*0.1
```

```
## (Intercept)  
## 0.7012204
```

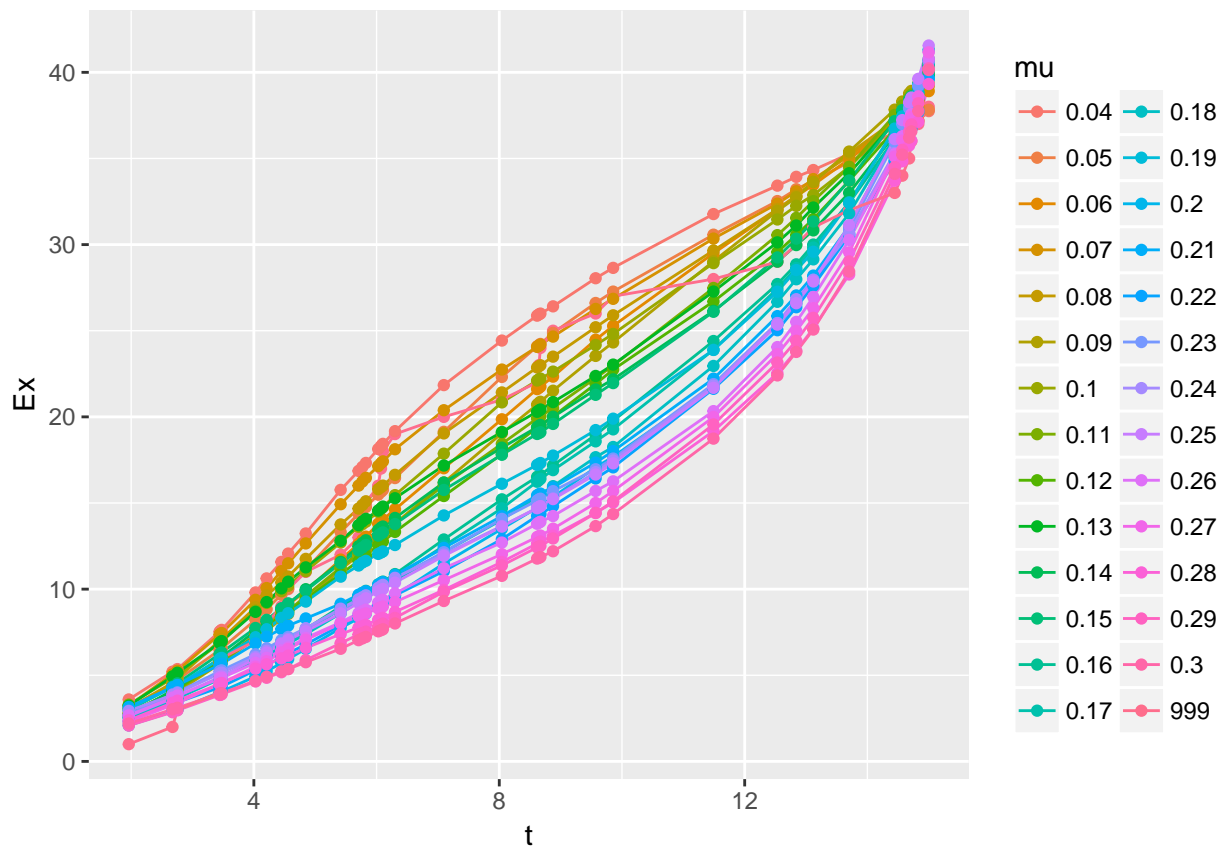
```
lm44$coef[1] + lm44$coef[2]*0.1
```

```
## (Intercept)  
## 0.6560708
```

The Ltt plot

Now let's try to minimize ltt, but first visualize it

```
time = proc.time()
grid = wt
Ltt = data.frame(t=cumsum(wt), Ex = (s$newick.extant.p)$n, mu=999)
for(i in 1:length(mu0)){
  mu = mu0[i]
  pars = c(l1[i],m1[i],k1[i])
  ltt = data.frame(t = cumsum(wt), Ex = expectedLTT(pars,drop.extinct = TRUE, grid=cumsum(wt))$Ex, mu=mu)
  Ltt = rbind(Ltt,ltt)
}
Ltt$mu = as.factor(Ltt$mu)
ggplot(data=Ltt, aes(x=t, y=Ex, colour = mu)) + geom_line() + geom_point()
```



```
print(proc.time()-time)
```

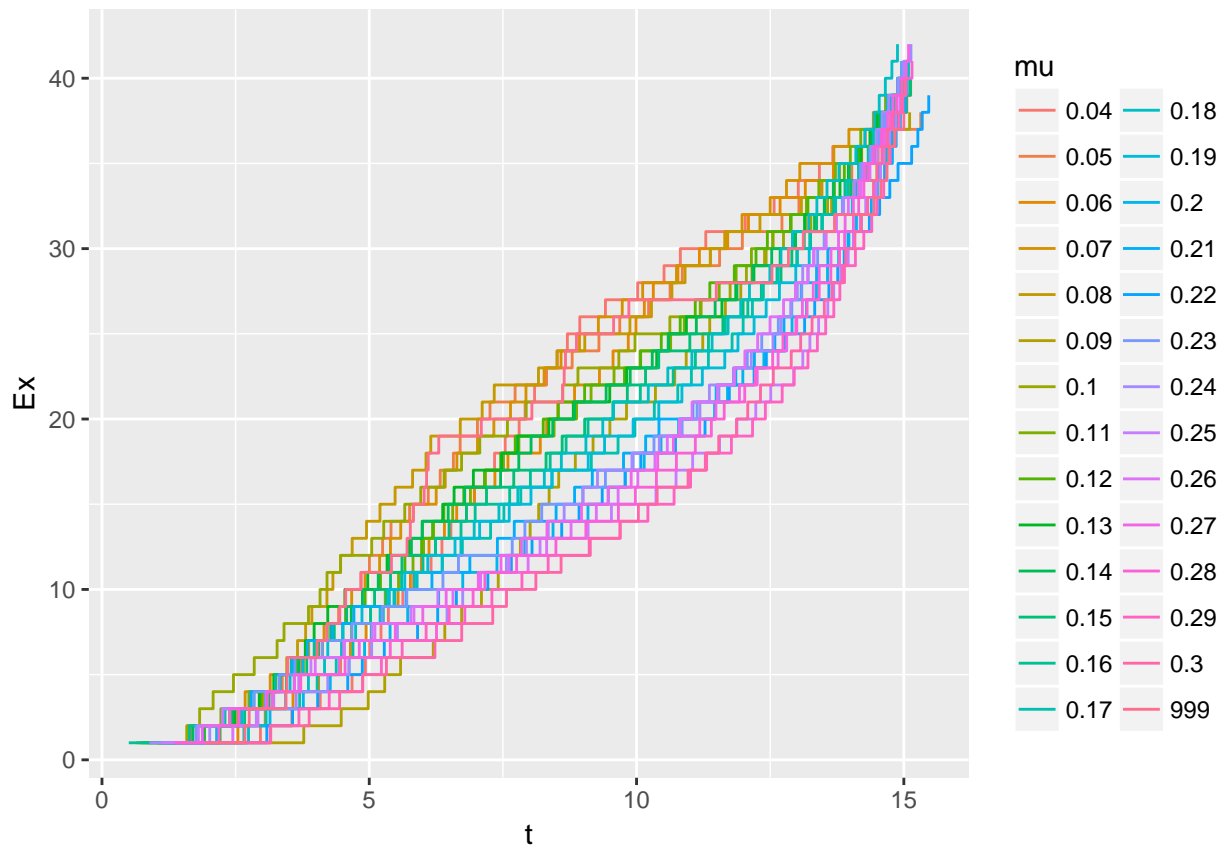
```
##      user  system elapsed
## 800.928   0.028 801.029
```

```
time = proc.time()
Ltt2 = data.frame(t=cumsum(wt), Ex = (s$newick.extant.p)$n, mu=999)
for(i in 1:length(mu0)){
  mu = mu0[i]
  pars = c(l1[i],m1[i],k1[i])
  expe = expectedLTT2(pars)
  ltt2 = data.frame(t = cumsum(expe$t), Ex = expe$Ex, mu=mu)
```

```
Ltt2 = rbind(Ltt2, ltt2)
}
Ltt2$mu = as.factor(Ltt2$mu)
print(proc.time()-time)
```

```
##      user  system elapsed
## 84.236   0.028  84.286
```

```
ggplot(data=Ltt2, aes(x=t, colour = mu)) + geom_step(aes(y=Ex)) #+ scale_color_brewer(name = "Method",
```



```
l1 = Ltt[Ltt$mu == 999,]
diff_ltt = NaN
sq_diff = NaN
diff_mat = matrix(0, ncol=length(mu0), nrow=length(wt))
for(i in 1:length(mu0)){
  mu = mu0[i]
  ltt = Ltt[Ltt$mu == mu,]
  ltt$Ex = abs(l1$Ex-ltt$Ex)
  diff_mat[,i] = ltt$Ex
  diff_ltt[i] = sum(ltt$Ex)
  sq_diff[i] = sum((l1$Ex-ltt$Ex)^2)
}
n_diff_ltt = diff_ltt*wt
```

```
## Warning in diff_ltt * wt: longer object length is not a multiple of shorter
## object length
```

```
n_diff_ltt

## [1] 181.975661 38.790611 5.465826 49.950100 1.084340 1.444825
## [7] 37.970172 16.160830 21.491267 8.775069 22.755875 52.655763
## [13] 39.821809 8.955118 8.924846 23.931219 7.819554 4.824470
## [19] 34.211471 129.433928 152.438611 97.228231 5.400489 4.735825
## [25] 41.375117 146.610782 62.935686 151.773133 56.694765 20.956833
## [31] 20.161568 37.817549 66.199259 8.516344 9.937103 3.252853
## [37] 9.035809 13.466519
```

```
diff_ltt_M = data.frame(mu = mu0, diff_ltt = diff_ltt)
choosed_mu = diff_ltt_M[diff_ltt_M$diff_ltt == min(diff_ltt_M$diff_ltt) ,]
choosed_mu$mu
```

```
## [1] 0.05
```

```
b = matrix(0,ncol=length(mu0),nrow=length(wt))
for(i in 1:length(mu0)){
  b[i,] = diff_mat[i,]==rowMins(diff_mat)[i]
}
colSums(b)
```

```
## [1] 2 2 3 5 3 1 3 0 0 1 0 1 0 0 0 1 2 0 0 0 0 1 0 0 0 2 0
```

```
MP[MP[,2] == choosed_mu$mu]
```

```
## [1] 0.5337676 0.0500000 42.5976211
```

```
MP1[max(colSums(b)) == colSums(b)]
```

```
## [1] 0.8525022 0.0700000 39.4487132
```

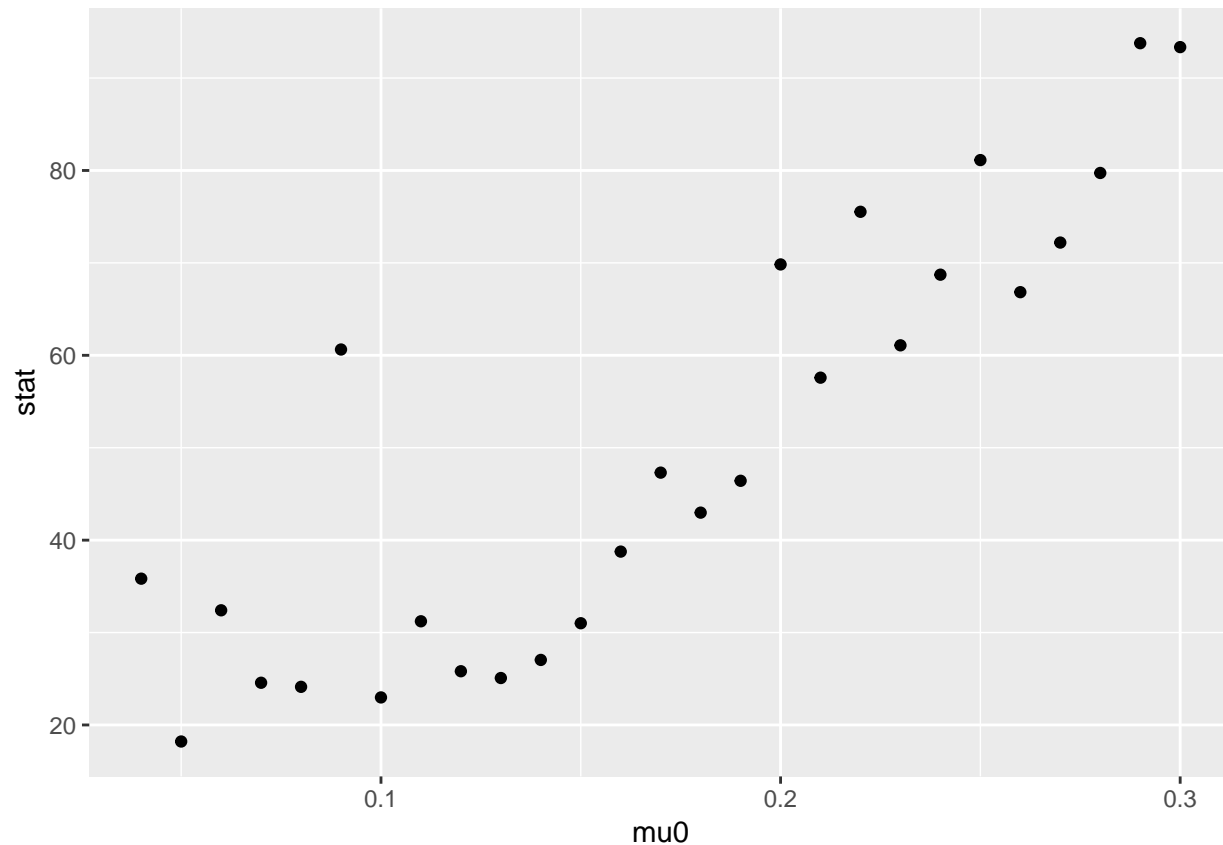
```
b[,max(colSums(b)) == colSums(b)]
```

```
## [1] 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 1 0 0 1 0 0 0 1 0 1 1 0 0 0 0 0 0 0 0
## [36] 0 0 0
```

Let's use the second method

```
stat=NULL
for(i in 1:length(mu0)){
  ltt_temp = Ltt2[Ltt2$mu == mu0[i],]
  ltt_temp = rbind(ltt_temp, data.frame(t=15,Ex=ltt_temp$Ex[length(ltt_temp$Ex)]+1,mu=ltt_temp$mu[1]))
  pp = list(wt=c(ltt_temp$t[1],diff(ltt_temp$t)),E=rep(1,(length(ltt_temp$t)-1)),n=ltt_temp$Ex)
  phy = p2phylo(pp)
  stat[i] = ltt_stat(s$newick.extant,phy)
}

qplot(mu0,stat)
```



that look nice, let's do it a bit more professional

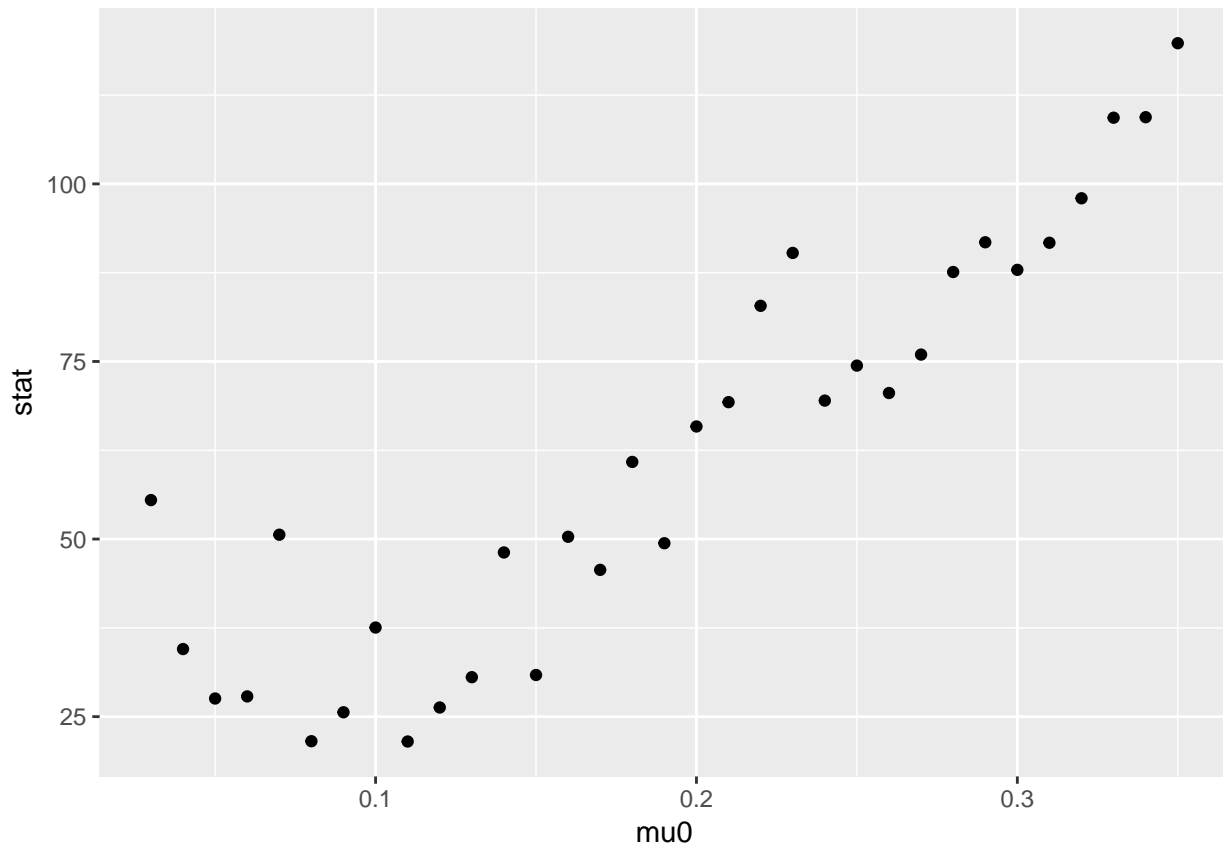
```
time=proc.time()
stat=NULL
mu0 = seq(0.03,0.35,by=0.01)
phylo = s$newick.extant
prior_pars = c(0.8,40)
for(i in 1:length(mu0)){
  mu = mu0[i]
  stat[i] = ltt_mu(mu,phylo,prior_pars = prior_pars)
}
```

```
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
```

```
print(proc.time()-time)
```

```
##      user   system elapsed
## 126.468    0.012 126.506
```

```
qplot(mu0,stat)
```



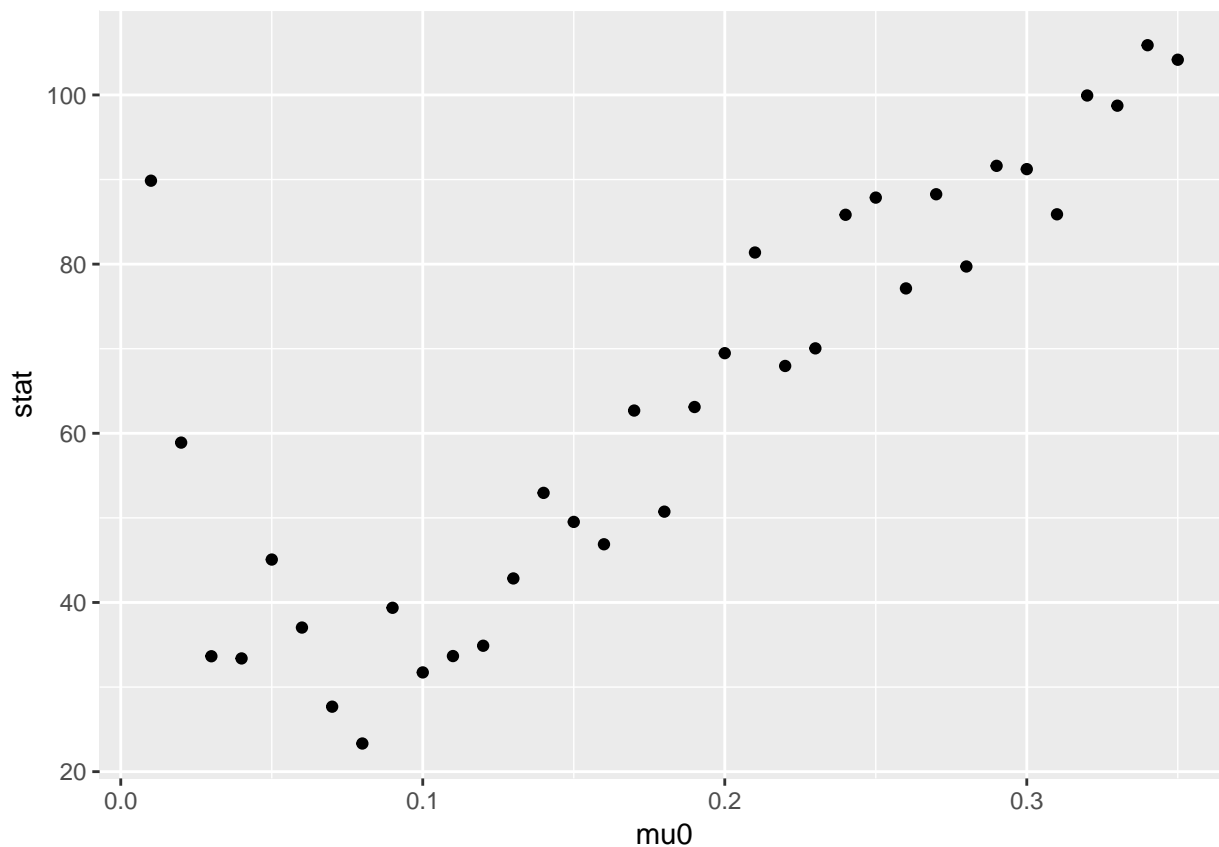
```
time=proc.time()
stat=NULL
mu0 = seq(0.01,0.35,by=0.01)
phylo = s$newick.extant
prior_pars = c(0.8,40)
for(i in 1:length(mu0)){
  mu = mu0[i]
  stat[i] = ltt_mu(mu,phylo,prior_pars = prior_pars,n_trees=50)
}
```

```
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
```

```
print(proc.time()-time)
```

```
##      user   system elapsed
## 609.900    0.056 610.034
```

```
qplot(mu0,stat)
```



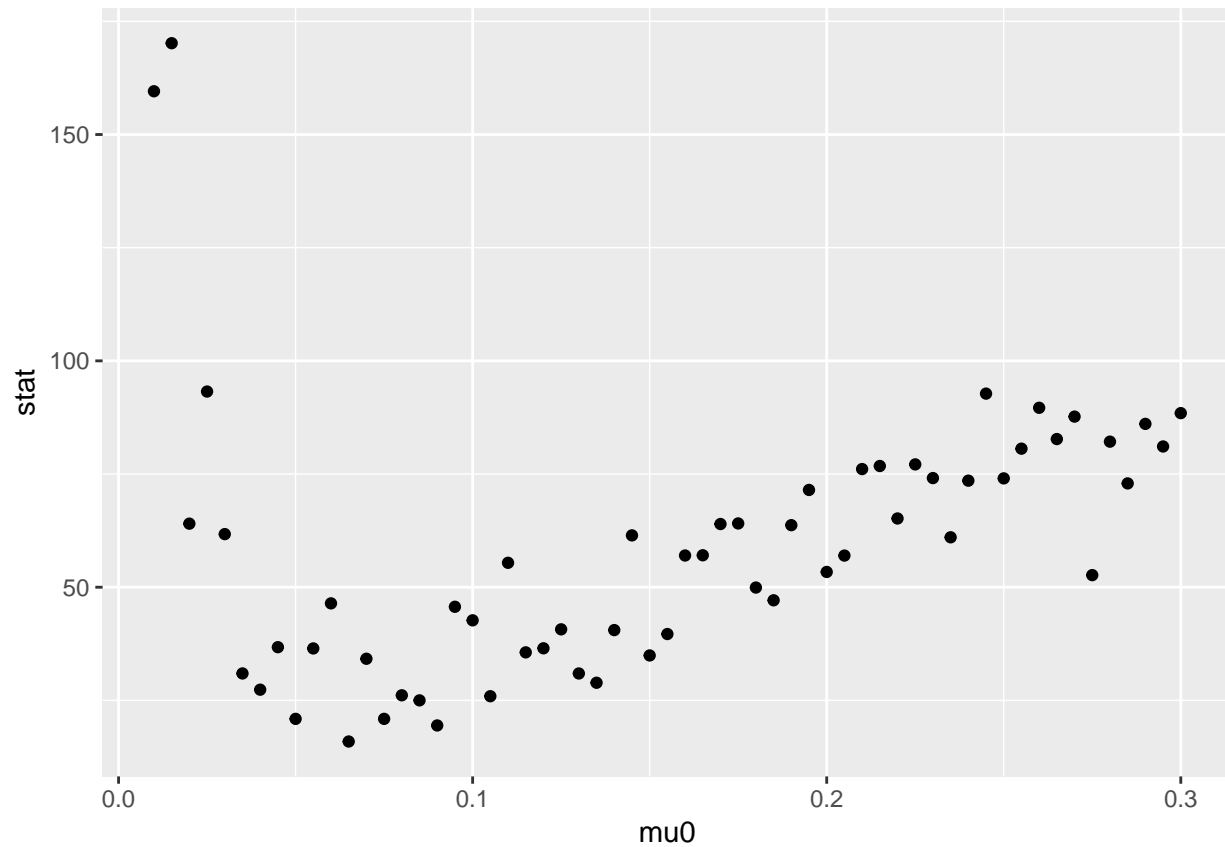
```
time=proc.time()
stat=NULL
mu0 = seq(0.01,0.3,by=0.005)
phylo = s$newick.extant
prior_pars = c(0.8,40)
#n_trees = 10
for(i in 1:length(mu0)){
  mu = mu0[i]
  stat[i] = ltt_mu(mu,phylo,prior_pars = prior_pars)
}
```

```
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
```

```
print(proc.time()-time)
```

```
##      user   system elapsed
## 180.852    0.020 180.868
```

```
qplot(mu0,stat)
```



```
time=proc.time()
stat=NULL
mu0 = seq(0.01,0.3,by=0.001)
phylo = s$newick.extant
prior_pars = c(0.8,40)
#n_trees = 10
for(i in 1:length(mu0)){
  mu = mu0[i]
  stat[i] = ltt_mu(mu,phylo,prior_pars = prior_pars)
}
```

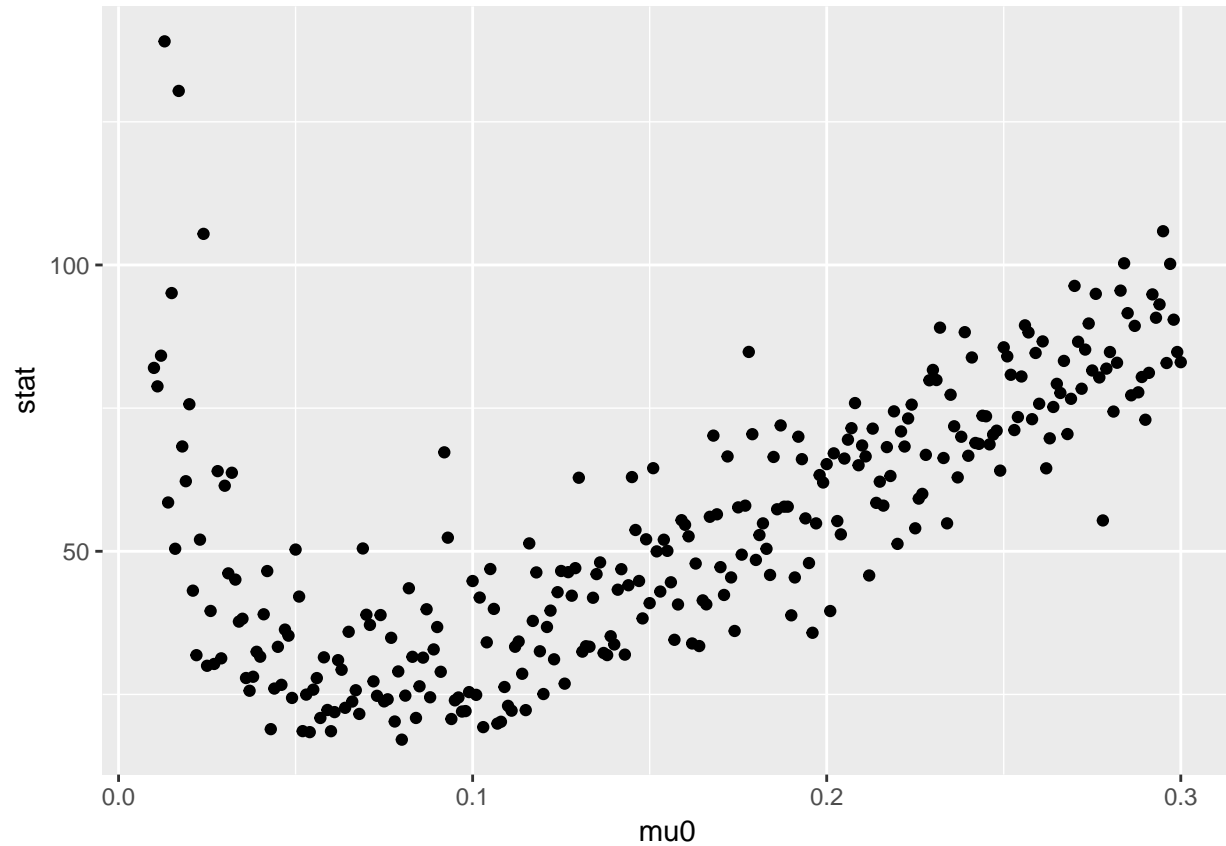
```
## [1] "DIFFERENT LENGHT IN PHYLOGENIES!"
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```



```
print(proc.time()-time)
```

```
##      user  system elapsed  
## 874.912   0.044 874.898
```

```
qplot(mu0,stat)
```



what if we take this min?

```
mu = mean(mu0[stat < quantile(stat,0.1)])  
mu
```

```
## [1] 0.08003448
```

```
trees = sim_srt(wt=wt,pars = prior_pars,n_trees = n_trees,mu = mu)  
lambda_K = subplex(par =c(2,60), fn = llik_st , setoftrees = trees, mu = mu, impSAM = FALSE)$par  
lambda_K
```

```
## [1] 0.6214694 40.9475748
```

```
p
```

```
## [1] 0.88886163 0.08410374 40.74476279
```

now... what about a small EM?

```
pars = c(2,80)  
s = sim_phyl(seed=3)  
stat=NULL  
mu0 = seq(0.01,0.35,by=0.005)  
phylo = s$newick.extant
```

```

wt = (s$newick.extant.p)$wt
for(i in 1:30){
  print(pars)
  stat=NULL
  for(j in 1:length(mu0)){
    mu = mu0[j]
    stat[j] = ltt_mu(mu,phylo,prior_pars = pars)
  }
  qqplot(mu0,stat,main=paste('iteration',i))
  mu = mean(mu0[stat < quantile(stat,0.1)])
  trees = sim_srt(wt=wt,pars = prior_pars,n_trees = n_trees,mu = mu)
  pars = subplex(par =c(2,60), fn = llik_st , setoftrees = trees, mu = mu, impsam = FALSE)$par
}

```

it looks fine, but 10 trees has too much variation, let's repeat it for 100 trees

```

#ct = 15
#dt = 0.1
#grid = wt
Ltt = data.frame(t=cumsum(wt), Ex = (s$newick.extant.p)$n, mu=999)

for(i in 1:length(mu0)){
  mu = mu0[i]
  pars = c(l2[i],m2[i],k2[i])
  ltt = data.frame(t = cumsum(wt), Ex = expectedLTT(pars,drop.extinct = TRUE, grid=cumsum(wt))$Ex, mu=mu)
  Ltt = rbind(Ltt,ltt)
}
Ltt$mu = as.factor(Ltt$mu)
ggplot(data=Ltt, aes(x=t, y=Ex, colour = mu)) + geom_line() + geom_point()

l1 = Ltt[Ltt$mu == 999,]
diff_ltt = NaN
sq_diff = NaN
diff_mat = matrix(0,ncol=length(mu0),nrow=length(wt))
for(i in 1:length(mu0)){
  mu = mu0[i]
  ltt = Ltt[Ltt$mu == mu,]
  ltt$Ex = abs(l1$Ex-ltt$Ex)
  diff_mat[,i] = ltt$Ex
  diff_ltt[i] = sum(ltt$Ex)
  sq_diff[i] = sum((l1$Ex-ltt$Ex)^2)
}
diff_ltt_M = data.frame(mu = mu0, diff_ltt = diff_ltt)
choosed_mu = diff_ltt_M[diff_ltt_M$diff_ltt == min(diff_ltt_M$diff_ltt) ,]
choosed_mu$mu
b = matrix(0,ncol=length(mu0),nrow=length(wt))
for(i in 1:length(mu0)){
  b[i,] = diff_mat[i,]==rowMins(diff_mat)[i]
}
colSums(b)
MP[MP[,2] == choosed_mu$mu]
MP2[max(colSums(b)) == colSums(b)]

```

```
#qqplot()
```

(not working yet)

Meta Analysis

Now we are prepared to estimate parameters of a set of (100) trees and see the distribution.

The algorithm is:

1. simulate tree and save MLE. Drop extinct species and save ltt
2. create a grid μ_g and run monte-carlo for every $\mu \in \mu_g$, then get $(\lambda(\mu), \mu, K(\mu))$, $\forall \mu \in \mu_g$ and the corresponding ltt
3. take the best $(\lambda(\mu), \mu, K(\mu))$ taking min ltt

```
ct = 15
dt = 0.1
#grid = seq(0,ct, by=dt)
n_it = 10
mu0 = seq(0.04,0.35,by=0.03)
n_trees = 10
MMP = matrix(nrow=n_it, ncol=3)
RMP = matrix(nrow=n_it, ncol=3)
for(j in 1:n_it){
  s = sim_phyl()
  p <- subplex(par = c(2,0.2,60), fn = llik, n = s$n, E = s$E, t = s$wt)$par
  MMP[j,] = p
  s2 = s$newick.extant.p
  wt = s2$wt
  grid = cumsum(wt)
  ltt1 = data.frame(t=grid, Ex = s2$n, mu=999)
  Ltt = ltt1
  MP = matrix(nrow=length(mu0), ncol=3)
  for(i in 1:length(mu0)){
    mu = mu0[i]
    trees = sim_srt(wt=wt, pars=c(p[1],mu,p[3]), parallel = F, n_trees = n_trees)
    pars = subplex(par = c(2,60), fn = llik_st, setoftrees = trees, mu = mu, impSAM = FALSE)$par
    pars = c(pars[1],mu,pars[2])
    MP[i,] = pars
    ltt = data.frame(t = grid, Ex = expectedLTT(pars,drop.extinct = TRUE, grid=grid)$Ex, mu=mu)
    Ltt = rbind(Ltt,ltt)
  }
  diff_mat = matrix(0,ncol=length(mu0),nrow=length(wt))
  for(i in 1:length(mu0)){
    mu = mu0[i]
    ltt = Ltt[Ltt$mu == mu,]
    ltt$Ex = abs(ltt1$Ex-ltt$Ex)
    diff_mat[,i] = ltt$Ex
  }
  b = matrix(0,ncol=length(mu0),nrow=length(wt))
  for(i in 1:length(mu0)){
    b[i,] = diff_mat[i,]==rowMins(diff_mat)[i]
  }
}
```

```
RMP[j,] = MP[max(colSums(b)) == colSums(b),]  
}
```

```
RMP  
MMP
```