

# 패널자료분석

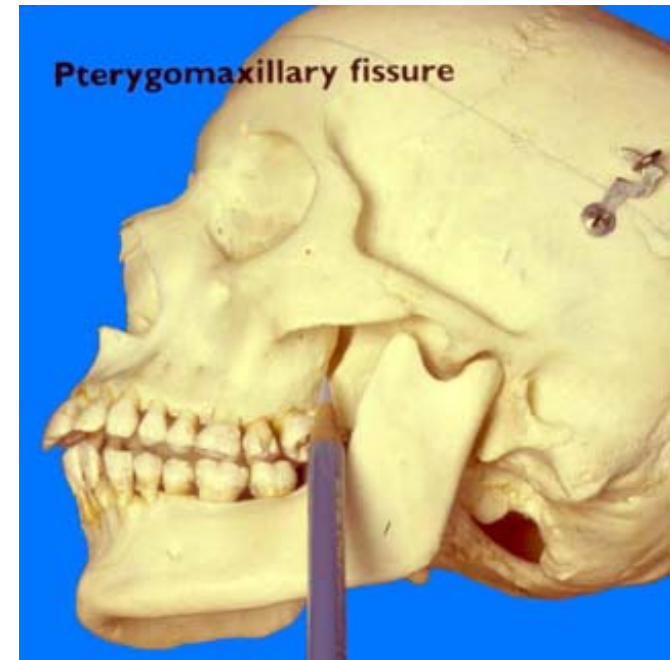
Panel (longitudinal) Data Analysis

## 패널자료란?

- 동일한 대상 (subject, experimental unit)으로부터 여러 시점에 대해 수집한 자료
  - 인간 혹은 동물에 대한 임상실험
  - 농작물의 성장, 부패

# Dental Study

- 27명의 어린이 (16명의 남자, 11명의 여자)
- 각 어린이의 뇌하수체에서 익돌상악열구 (**pterygomaxillary fissure**)까지의 거리(mm)를 8, 10, 12, 14세에 측정
- Questions
  - 시간에 따라 거리가 변화하는가?
  - 변화의 패턴은?
  - 남자와 여자의 패턴 차이는?



# Dental Study: 데이터구조 변형

```
> library(mice)
```

```
> potthoffroy
```

	id	sex	d8	d10	d12	d14
1	1	F	21.0	20.0	21.5	23.0
2	2	F	21.0	21.5	24.0	25.5
3	3	F	20.5	24.0	24.5	26.0
4	4	F	23.5	24.5	25.0	26.5
5	5	F	21.5	23.0	22.5	23.5
6	6	F	20.0	21.0	21.0	22.5
7	7	F	21.5	22.5	23.0	25.0
8	8	F	23.0	23.0	23.5	24.0
9	9	F	20.0	21.0	22.0	21.5
10	10	F	16.5	19.0	19.0	19.5
11	11	F	24.5	25.0	28.0	28.0
12	12	M	26.0	25.0	29.0	31.0
13	13	M	21.5	22.5	23.0	26.5
14	14	M	23.0	22.5	24.0	27.5
15	15	M	25.5	27.5	26.5	27.0
16	16	M	20.0	23.5	22.5	26.0
17	17	M	24.5	25.5	27.0	28.5
18	18	M	22.0	22.0	24.5	26.5
19	19	M	24.0	21.5	24.5	25.5
20	20	M	23.0	20.5	31.0	26.0
21	21	M	27.5	28.0	31.0	31.5
22	22	M	23.0	23.0	23.5	25.0
23	23	M	21.5	23.5	24.0	28.0
24	24	M	17.0	24.5	26.0	29.5
25	25	M	22.5	25.5	25.5	26.0
26	26	M	23.0	24.5	26.0	30.0
27	27	M	22.0	21.5	23.5	25.0



```
> data=reshape(potthoffroy,idvar="id", varying=list(3:6),v.names="dist",direction="long")
```

```
> data$sex.m=1
```

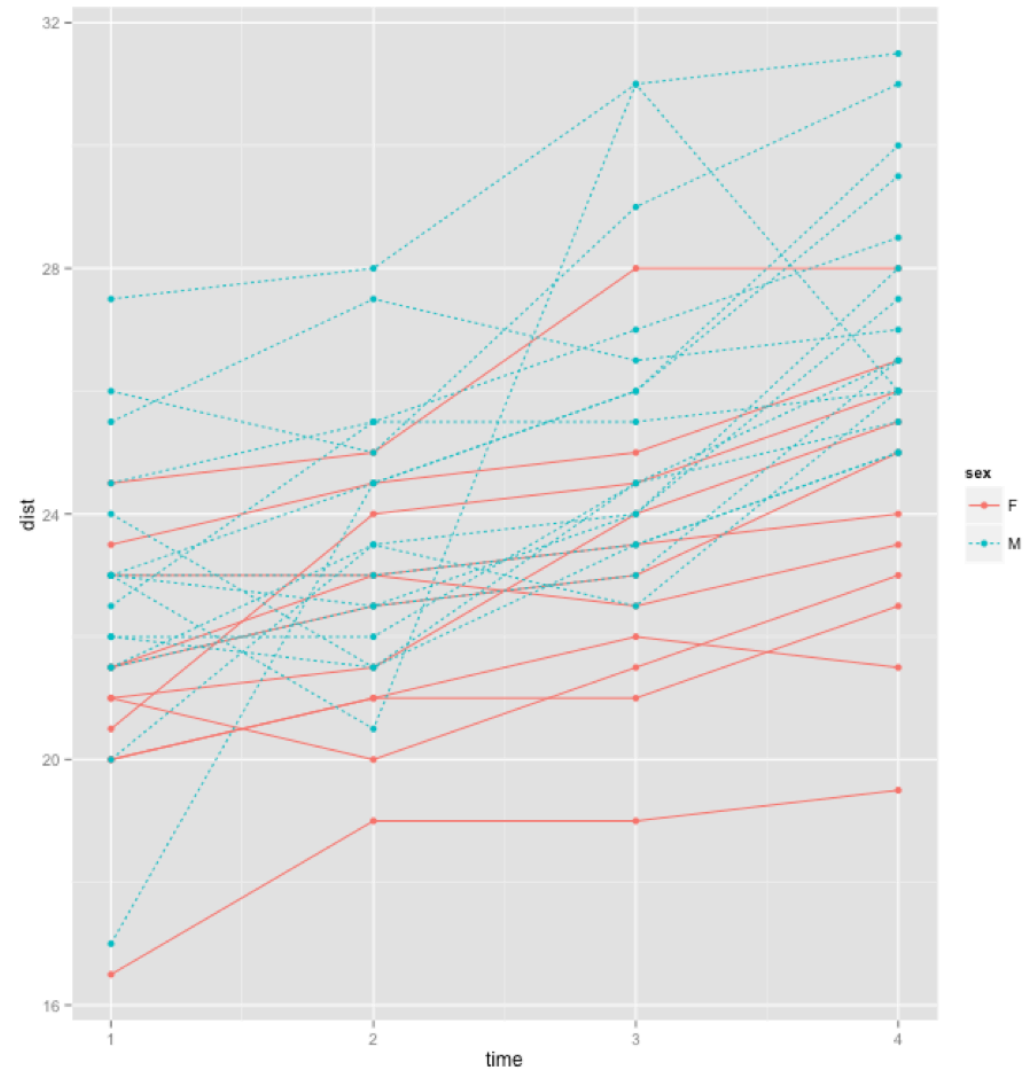
```
> data$sex.m[data$sex=="F"]=0
```

```
> data
```

	id	sex	time	dist	sex.m
1.1	1	F	1	21.0	0
2.1	2	F	1	21.0	0
3.1	3	F	1	20.5	0
4.1	4	F	1	23.5	0
5.1	5	F	1	21.5	0
6.1	6	F	1	20.0	0
7.1	7	F	1	21.5	0
8.1	8	F	1	23.0	0
9.1	9	F	1	20.0	0
10.1	10	F	1	16.5	0
11.1	11	F	1	24.5	0
12.1	12	M	1	26.0	1
13.1	13	M	1	21.5	1
14.1	14	M	1	23.0	1
15.1	15	M	1	25.5	1
16.1	16	M	1	20.0	1
17.1	17	M	1	24.5	1
18.1	18	M	1	22.0	1
19.1	19	M	1	24.0	1
20.1	20	M	1	23.0	1
21.1	21	M	1	27.5	1
22.1	22	M	1	23.0	1
23.1	23	M	1	21.5	1
24.1	24	M	1	17.0	1
25.1	25	M	1	22.5	1
26.1	26	M	1	23.0	1
27.1	27	M	1	22.0	1
1.2	1	F	2	20.0	0
2.2	2	F	2	21.5	0
3.2	3	F	2	24.0	0
4.2	4	F	2	24.5	0
~	~	~	~	~	~

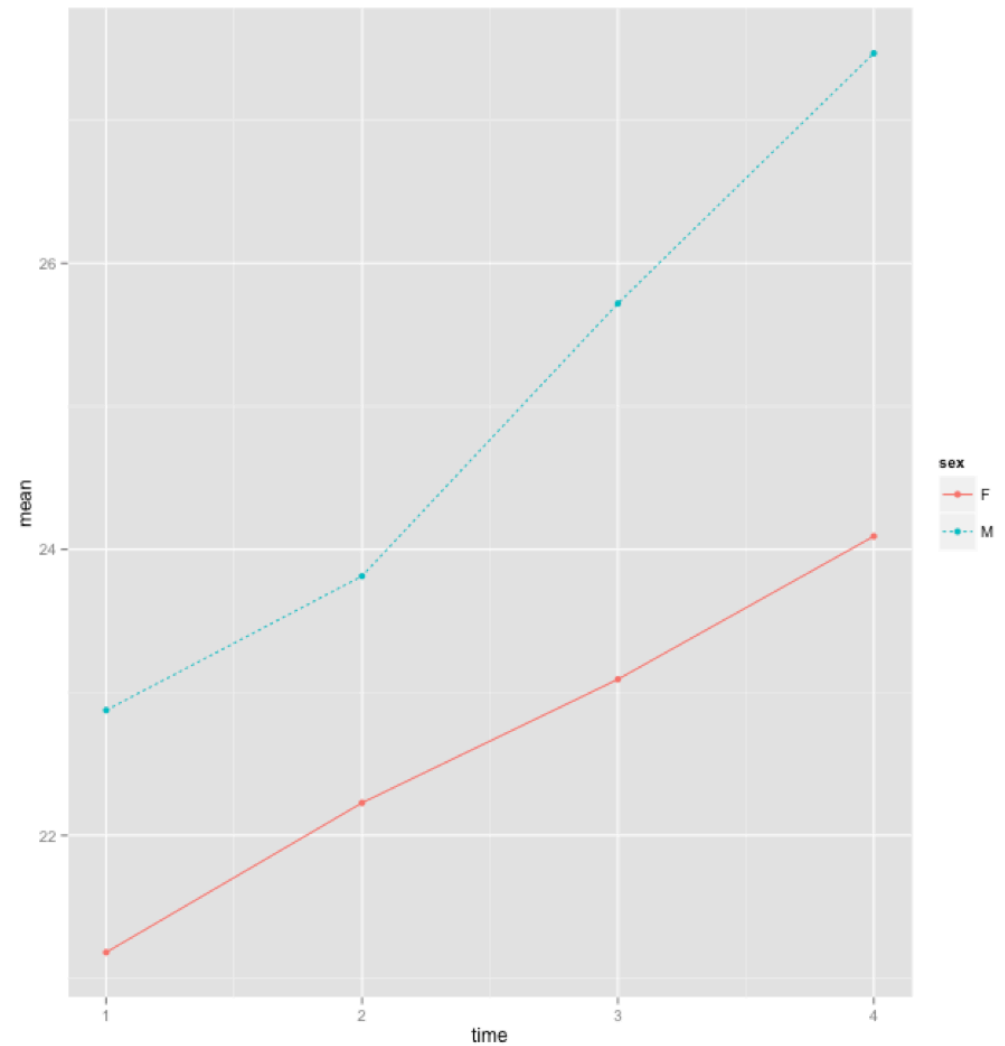
# Dental Study: Interaction Plot

```
library(ggplot2)
ggplot(data, aes(y=dist, x=time, group=id, colour=sex)) +
  geom_line(aes(linetype=sex), stat="identity") +
  geom_point()
```



## Dental Study: 성별 평균

```
library(plyr)
data2=ddply(data,~sex+time,summarize,mean=mean(dist))
ggplot(data2,aes(y=mean,x=time,colour=sex))+
  geom_line(aes(linetype=sex),stat="identity")+
  geom_point()
```



# Repeated Measure ANOVA

- 만일 시간의 흐름에 따른 패턴을 파악하는 것이 목적이 아니라 남자와 여자 사이의 평균적인 차이를 파악하는 것이 목적이라면?
  - 각 시점 간의 관측치가 독립이 아님 → 시점 변수를 factor화 한 ANOVA가 적당하지 않음

$$Y_{ij} = \mu + \alpha_j + \epsilon_{ij}$$

$$\begin{bmatrix} \epsilon_{i1} \\ \epsilon_{i2} \\ \epsilon_{i3} \\ \epsilon_{i4} \end{bmatrix} \sim N(\mathbf{0}, \sigma^2 V)$$

- $Y_{ij}$ : j번째 그룹의 i번째 subject의 관측치
- $\mu$ : 전체 평균
- $\alpha_j$ : 성별 효과 (j번째 그룹 평균과 전체 평균의 차이)

# Repeated Measure ANOVA

- Covariance matrix  $V$ 의 종류

- Autoregressive(AR)

$$\sigma^2 \begin{bmatrix} 1 & \rho & \rho^2 & \rho^3 \\ \rho & 1 & \rho & \rho^2 \\ \rho^2 & \rho & 1 & \rho \\ \rho^3 & \rho^2 & \rho & 1 \end{bmatrix}$$

- Compound symmetry

$$\begin{bmatrix} \sigma^2 + \sigma_1^2 & \sigma_1^2 & \sigma_1^2 & \sigma_1^2 \\ \sigma_1^2 & \sigma^2 + \sigma_1^2 & \sigma_1^2 & \sigma_1^2 \\ \sigma_1^2 & \sigma_1^2 & \sigma^2 + \sigma_1^2 & \sigma_1^2 \\ \sigma_1^2 & \sigma_1^2 & \sigma_1^2 & \sigma^2 + \sigma_1^2 \end{bmatrix}$$

- Unstructured

$$\begin{bmatrix} \sigma_1^2 & \sigma_{12} & \sigma_{13} & \sigma_{14} \\ \sigma_{12} & \sigma_2^2 & \sigma_{23} & \sigma_{24} \\ \sigma_{13} & \sigma_{23} & \sigma_3^2 & \sigma_{34} \\ \sigma_{14} & \sigma_{24} & \sigma_{34} & \sigma_4^2 \end{bmatrix}$$



## library(nlme)사용

### AR(1) correlation matrix

```
> model3=gls(dist~sex,data,correlation=corAR1(form=~1|id)) #AR(1)
> anova(model3)
Denom. DF: 106
```

	numDF	F-value	p-value
(Intercept)	1	3280.246	<.0001
sex	1	8.006	0.0056

```
> summary(model3)
Generalized least squares fit by REML
Model: dist ~ sex
Data: data
```

	AIC	BIC	logLik
	493.0365	503.6903	-242.5183

```
Correlation Structure: AR(1)
Formula: ~1 | id
Parameter estimate(s):
Phi
0.6354623

Coefficients:

```

	Value	Std.Error	t-value	p-value
(Intercept)	22.64243	0.6586210	34.37855	0.0000
sexM	2.42091	0.8555738	2.82957	0.0056

```
Correlation:
(Intr)
sexM -0.77

Standardized residuals:

```

	Min	Q1	Med	Q3	Max
	-2.85756288	-0.73122728	-0.03646313	0.50913515	2.28108149

```
Residual standard error: 2.821756
Degrees of freedom: 108 total; 106 residual
```

### Compound Symmetric correlation matrix

```
> model4=gls(dist~sex,data,correlation=corCompSymm(form=~1|id)) #Compound symmetry
> anova(model4)
Denom. DF: 106
```

	numDF	F-value	p-value
(Intercept)	1	4123.156	<.0001
sex	1	9.292	0.0029

```
> summary(model4)
Generalized least squares fit by REML
Model: dist ~ sex
Data: data
```

	AIC	BIC	logLik
	513.8718	524.5255	-252.9359

```
Correlation Structure: Compound symmetry
Formula: ~1 | id
Parameter estimate(s):
Rho
0.3406282

Coefficients:

```

	Value	Std.Error	t-value	p-value
(Intercept)	22.647727	0.5861390	38.63884	0.0000
sexM	2.321023	0.7614168	3.04829	0.0029

```
Correlation:
(Intr)
sexM -0.77

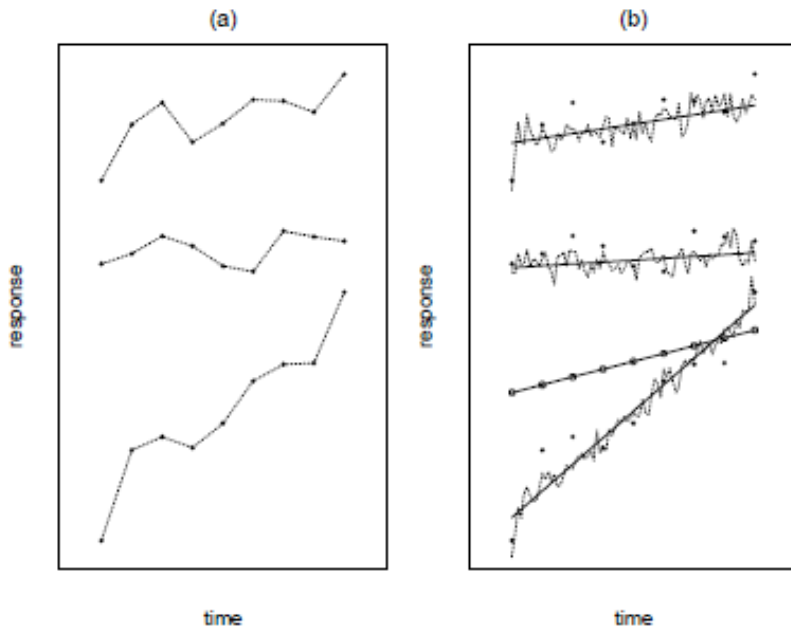
Standardized residuals:

```

	Min	Q1	Med	Q3	Max
	-2.91434827	-0.72001545	-0.02129916	0.56001202	2.38862270

```
Residual standard error: 2.734316
Degrees of freedom: 108 total; 106 residual
```

## 시간의 흐름에 따른 패턴 추정: 선형혼합모형



### IDEA

- 각 subject는 고유한 장기 추세를 가지고 있음
- 실제 관측치는 이 추세를 기반으로 변동성을 가짐
- 측정 오차를 가질 수 있음 (앞의 선형회귀모형은 동일한 성별에 속한 대상들 사이의 측정오차를 고려하지 않음)
- 모든 subject의 추세, 변동성, 측정오차를 각 시점에서 평균을 취하여 전 population에 대한 추세를 얻을 수 있음

### REMARK

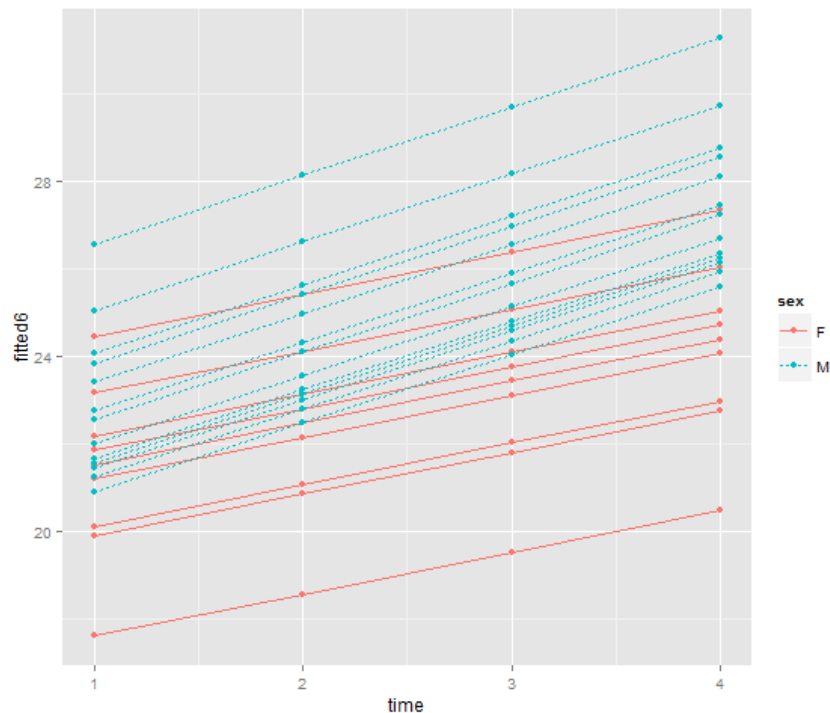
- 추세가 반드시 선형일 필요는 없음

## • Random intercept model

- 추세선의 절편에서 subject 간의 오차를 고려

$$Y_{it} = (\beta_0 + u_i) + (\beta_1 + \beta_3 d_i)t + \beta_2 d_i + \epsilon_{it},$$

$$u_i \sim N(0, \sigma_u^2), \epsilon_{it} \sim N(0, \sigma_\epsilon^2)$$



```
> model7=lmer(dist~sex+time+sex:time+(1+time|id),data=data)
```

```
> summary(model7)
```

Linear mixed model fit by REML ['lmerMod']

Formula: dist ~ sex + time + sex:time + (1 + time | id)

Data: data

REML criterion at convergence: 429.8

Scaled residuals:

Min	1Q	Median	3Q	Max
-3.1681	-0.3859	0.0071	0.4452	3.8495

Random effects:

Groups	Name	Variance	Std.Dev.	Corr
id	(Intercept)	3.4818	1.8660	
	time	0.1301	0.3607	-0.28
Residual		1.7162	1.3100	

Number of obs: 108, groups: id, 27

Fixed effects:

	Estimate	Std. Error	t value
(Intercept)	20.2500	0.7420	27.291
sexM	0.7969	0.9639	0.827
time	0.9591	0.2074	4.623
sexM:time	0.6097	0.2695	2.262

Correlation of Fixed Effects:

	(Intr)	sexM	time
sexM	-0.770		
time	-0.618	0.476	
sexM:time	0.476	-0.618	-0.770

```
data$fitted6=fitted(model6)
```

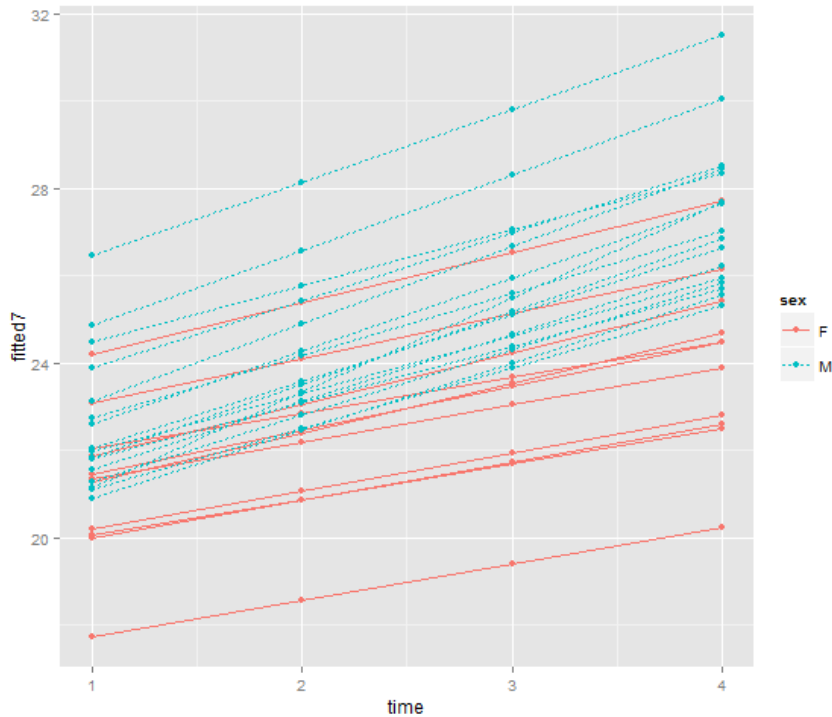
```
ggplot(data,aes(y=fitted6,x=time,group=id,colour=sex))+  
  geom_line(aes(linetype=sex),stat="identity")+  
  geom_point()
```

- Random intercept and slope model

- 추세선의 절편과 기울기에서 subject 간의 오차를 고려

$$Y_{it} = (\beta_0 + u_{1i}) + (\beta_1 + \beta_3 d_i + u_{2i})t + \beta_2 d_i + \epsilon_{it},$$

$$(u_{1i}, u_{2i})^T \sim N(0, \Sigma), \epsilon_{it} \sim N(0, \sigma_\epsilon^2)$$



```
> model6=lmer(dist~sex+time+sex:time+(1|id),data=data)
> summary(model6)
Linear mixed model fit by REML
t-tests use Satterthwaite approximations to degrees of freedom ['merModLmerTest']
Formula: dist ~ sex + time + sex:time + (1 | id)
Data: data
```

REML criterion at convergence: 431

Scaled residuals:

Min	1Q	Median	3Q	Max
-3.5980	-0.4546	0.0158	0.5024	3.6862

Random effects:

Groups	Name	Variance	Std.Dev.
id	(Intercept)	3.299	1.816
Residual		1.922	1.386

Number of obs: 108, groups: id, 27

Fixed effects:

	Estimate	Std. Error	df	t value	Pr(> t )
(Intercept)	20.2500	0.7496	59.3100	27.013	< 2e-16 ***
sexM	0.7969	0.9738	59.3100	0.818	0.4165
time	0.9591	0.1869	79.0000	5.130	2.02e-06 ***
sexM:time	0.6097	0.2428	79.0000	2.511	0.0141 *

---  
Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Correlation of Fixed Effects:

	(Intr)	sexM	time
sexM	-0.770		
time	-0.623	0.480	
sexM:time	0.480	-0.623	-0.770

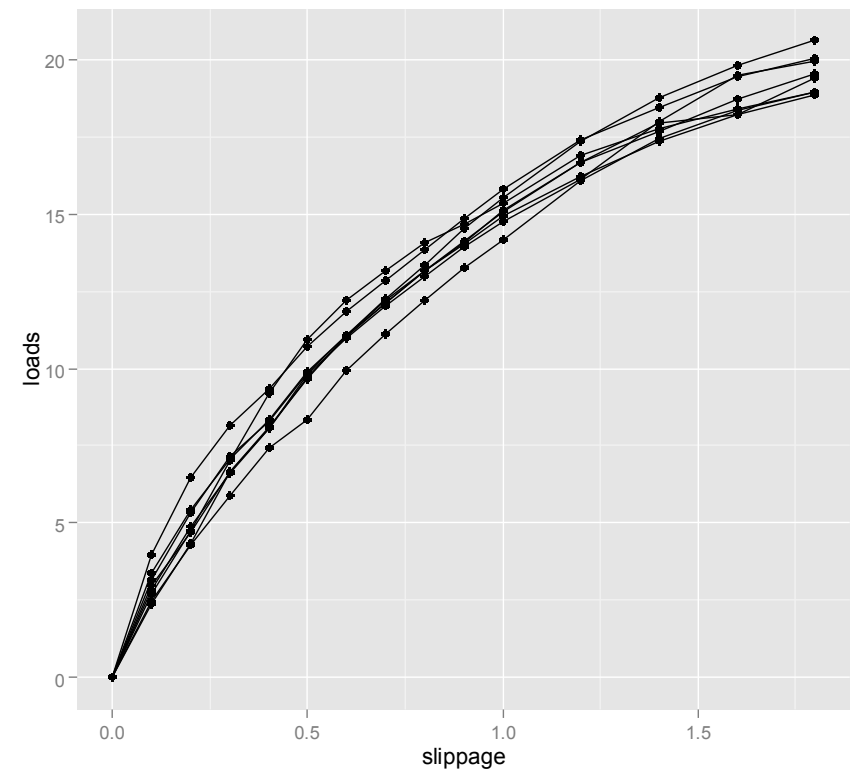
```
data$fitted7=fitted(model7)
ggplot(data,aes(y=fitted7,x=time,group=id,colour=sex))+
  geom_line(aes(linetype=sex),stat="identity")+
  geom_point()
```

## 예: Timber Slippage

- 8개 목재에 대해 클램프에서 미끄러지는데 필요한 무게를 15번씩 반복 측정
- slippage에 따라 load가 어떻게 달라지는가? (각 목재 간의 차이는 큰 관심 없음)

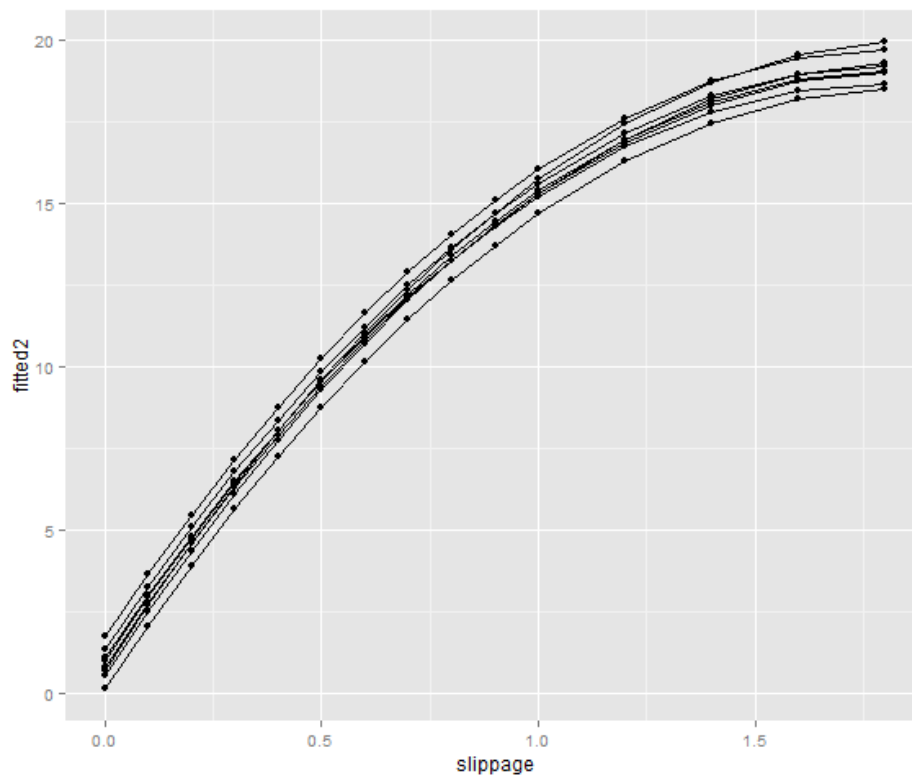
```
> head(timber)
  specimen slippage loads
spec1.1   spec1      0     0
spec2.1   spec2      0     0
spec3.1   spec3      0     0
spec4.1   spec4      0     0
spec5.1   spec5      0     0
spec6.1   spec6      0     0
> tail(timber)
  specimen slippage loads
spec3.15   spec3     1.8 19.40
spec4.15   spec4     1.8 18.93
spec5.15   spec5     1.8 20.62
spec6.15   spec6     1.8 20.05
spec7.15   spec7     1.8 19.54
spec8.15   spec8     1.8 18.87

ggplot(timber, aes(y=loads, x=slippage, group=specimen))+
  geom_line(stat="identity")+
  geom_point()
```



## 예: Timber Slippage

- Timber간 절편과 slippage의 기울기 오차 고려



```
> fit_timber2=lmer(loads~slippage+I(slippage^2)+(1+slippage|specimen),data=timber)
> summary(fit_timber2)
Linear mixed model fit by REML
t-tests use Satterthwaite approximations to degrees of freedom ['merModLmerTest']
Formula: loads ~ slippage + I(slippage^2) + (1 + slippage | specimen)
Data: timber
```

REML criterion at convergence: 205.2

Scaled residuals:

Min	1Q	Median	3Q	Max
-3.4946	-0.5045	-0.0281	0.5953	2.1885

Random effects:

Groups	Name	Variance	Std.Dev.	Corr
specimen	(Intercept)	0.2824	0.5314	
	slippage	0.1270	0.3564	-0.60
Residual		0.2489	0.4988	

Number of obs: 120, groups: specimen, 8

Fixed effects:

	Estimate	Std. Error	df	t value	Pr(> t )	
(Intercept)	0.9434	0.2187	9.1900	4.314	0.00186	**
slippage	19.8891	0.3271	78.8600	60.804	< 2e-16	***
I(slippage^2)	-5.4295	0.1651	103.0000	-32.880	< 2e-16	***

---  
signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

Correlation of Fixed Effects:

	(Intr)	slippg
slippage	-0.594	
I(slippg^2)	0.358	-0.885

```
timber$fitted2=fitted(fit_timber2)
ggplot(timber,aes(y=fitted2,x=slippage,group=specimen))+
  geom_line(stat="identity")+
  geom_point()
```