

# Reference growth charts for Saudi Arabian children and adolescents

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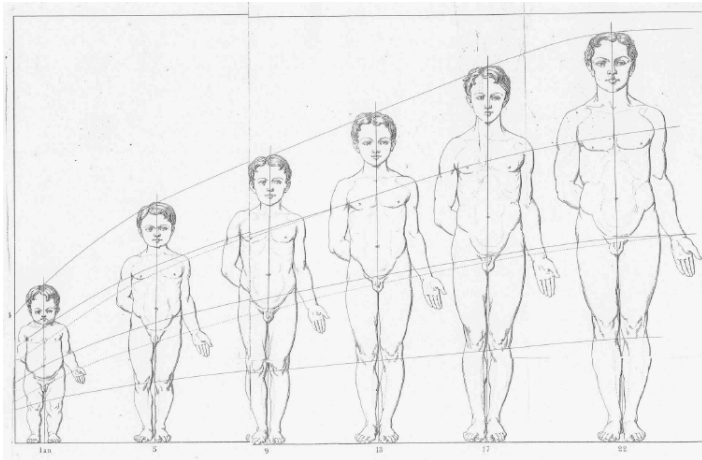
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## Quetelet's (1871) Growth Chart



## Assumption about normality

The reference growth charts are based on normality assumptions for the data.

Age-specific mean  $\mu(t)$  and standard deviation  $\sigma(t)$  curves are estimated and chosen quantile curve for a  $\alpha \in [0, 1]$  can then be constructed as:

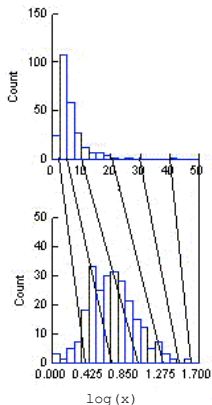
$$\hat{Q}(\alpha | t) = \hat{\mu}(t) + \hat{\sigma}(t)\Phi^{-1}(\alpha)$$

where  $\Phi^{-1}(\alpha)$  denotes the inverse of the standard normal distribution function, in other words normal equivalent deviate of size  $\alpha$  (corresponding to tail area).

## Anthropometric data

Anthropometric data:

- 1 non-normally distributed,
- 2 tends to be right skewed rather than left, which is why a *log* transformation is often suggested to cope with it.



## Penalised Maximum Likelihood Estimation

- Data:  $\{Y(t_{i,j}) : j = 1, \dots, J_i, i = 1, \dots, n\}$

- Model:  $Z(t) = \frac{[Y(t)/\mu(t)]^{\lambda(t)} - 1}{\lambda(t)\sigma(t)} \sim \mathcal{N}(0, 1)$

- Estimation:

$$\ell(\lambda, \mu, \sigma) = \sum_{i=0}^n \left[ \lambda(t_i) \log \frac{Y(t_i)}{\mu(t_i)} - \log \sigma(t_i) - \frac{1}{2} Z^2(t_i) \right]$$

$$\max[\ell(\lambda, \mu, \sigma) - \nu_\lambda \int (\lambda''(t))^2 dt - \nu_\mu \int (\mu''(t))^2 dt - \nu_\sigma \int (\sigma''(t))^2 dt]$$

- Quantile:  $Q(\alpha | t) = \mu(t)[1 + \lambda(t)\sigma(t)\Phi^{-1}(\alpha)]^{1/\lambda(t)}$

## Data

	A	B	C	D	E	F	G	H	I
1	REGION	IDNUMBER	SEX	WEIGHTOF	HEIGHTOF	HEADCIRC	AGEYEARS	AGEMON	
2		1	1	.NA	.NA	.NA	.NA	.NA	
3		1	1	2	.NA	.NA	.NA	.NA	
4		1	1	2	64.5	148	59	17.9688	215.63
5		1	2	1	.NA	.NA	.NA	.NA	
6		1	2	2	.NA	.NA	.NA	.NA	
7		1	2	2	14.6	99	50	4.6525	55.83
8		1	3	1	.NA	.NA	.NA	.NA	
9		1	3	2	.NA	.NA	.NA	.NA	
10		1	3	1	8.5	74.5	49	0.914	10.97
11		1	3	2	9.5	82	48	1.3855	16.63
12		1	3	1	15.6	113	51	4.6197	55.44
13		1	3	2	29.5	129.5	55	7.6469	91.76
14		1	6	1	.NA	.NA	.NA	.NA	
15		1	6	2	.NA	.NA	.NA	.NA	
16		1	6	.NA	.NA	.NA	.NA	.NA	
17		1	6	2	15.8	104	50	5.8009	69.61
18		1	6	1	35.2	130	51	9.6737	116.08
19		1	6	2	40	148	56	16.1846	194.21

## Codes and description of variables (health profile of Saudi children)

- Region** The ID number of the region. There are 13 regions in the Kingdom. All are covered in this survey.
- Id number** This is the id number of the household (family).
- Sex** 1=male, 2= female.
- Measure** The variables **weightof** (in Kg), **heightof** (in cm), **headcirc** (in cm), refer to the corresponding body measurements.
- Age** **Ageyears** and **agemon** refer to the date of measurement, recorded in Hijri calendar but subsequently converted to Gregorian.

## Issues related to constructing the reference growth charts

- Detecting the **outliers**
- **Smoothing** the curves
- **Averaging** the overlapping period 2 to 3 years of age
- **Goodness-of-fit** of the centile curves
- **Comparison** between different geographical regions and between genders

## Robust regression

R - a public domain language for data analysis

MASS package (contributed by W.N. Venables and B.D. Ripley)

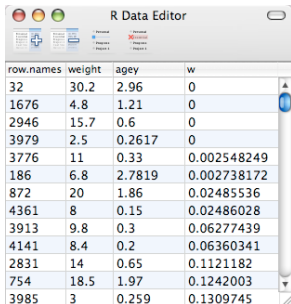
```
> library(MASS)
> mp<-rlm(log(weight)~1+agey+I(agey^2)+I(agey^3), method="MM")
```

An object of class `rlm` inherited from `lm` is used to fit linear models and it can be used to carry out regression.

Using `rlm` fitting is done by iterated re-weighted least squares (IWLS).

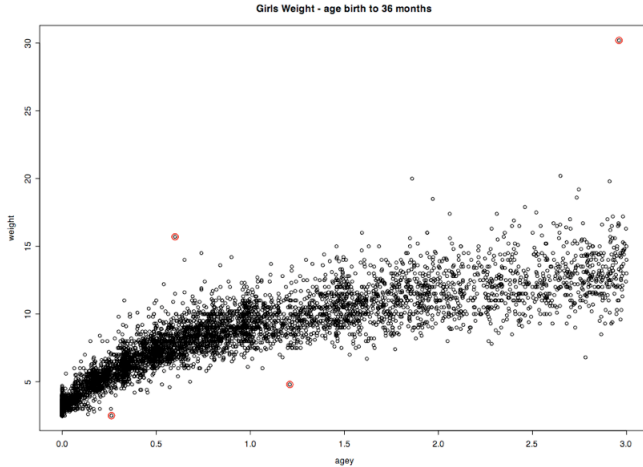
An additional component in `rlm` that is not in an `lm` object is:

`w` – the weights used in the IWLS process.



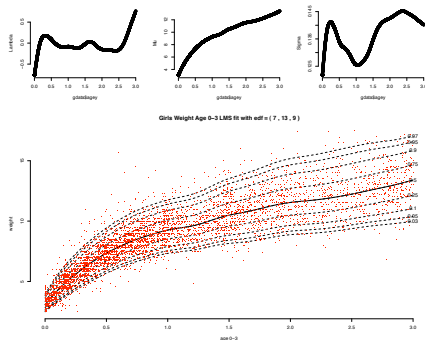
row.names	weight	agey	w
32	30.2	2.96	0
1676	4.8	1.21	0
2946	15.7	0.6	0
3979	2.5	0.2617	0
3776	11	0.33	0.002548249
186	6.8	2.7819	0.002738172
872	20	1.86	0.02485536
4361	8	0.15	0.02486028
3913	9.8	0.3	0.06277439
4141	8.4	0.2	0.06360341
2831	14	0.65	0.1121182
754	18.5	1.97	0.1242003
3985	3	0.259	0.1309745

## Detecting the outliers

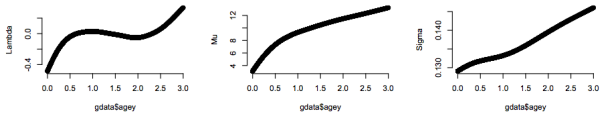


R package `lmsqreg` (contributed by V. J. Carry)

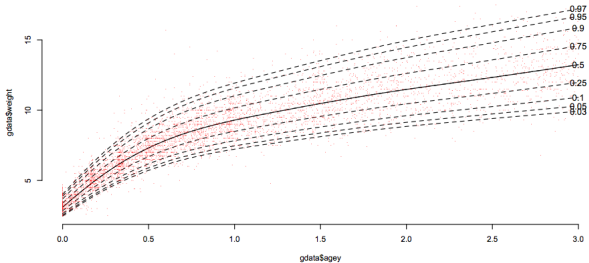
```
> library(lmsqreg)  
> mw3<-lmsqreg.fit(gdata$weight, gdata$agey, edf=c(7, 13, 9),  
pvec = c(0.03, 0.05, 0.1, 0.25, 0.5, 0.75, 0.9, 0.95, 0.97))
```



## Smoothing with edf(4, 6, 3)

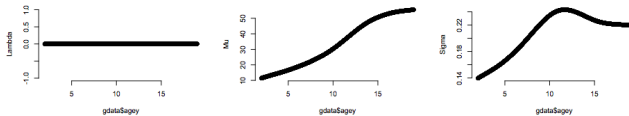


LMS fit with edf = (4,6,3), PL=9198.316

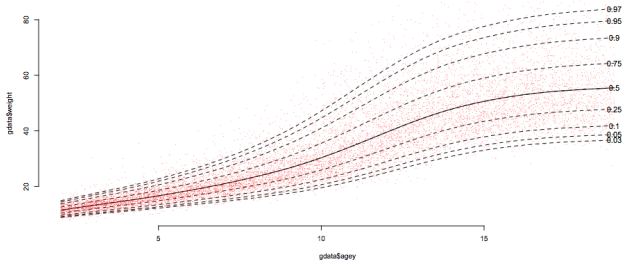


Construction of the growth charts  
Comparisons of the growth charts

$$\lambda = 0$$



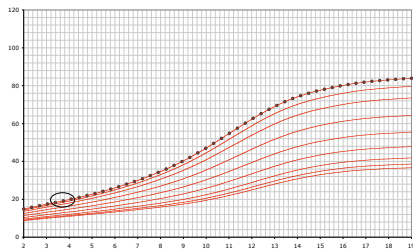
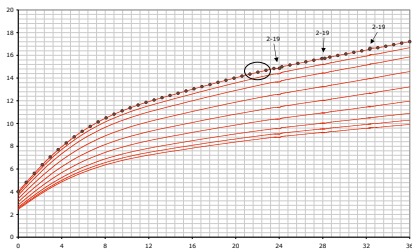
LMS fit with edf = (0,9,5), PL=12912.915



## Overlap for age 2 to 3 years

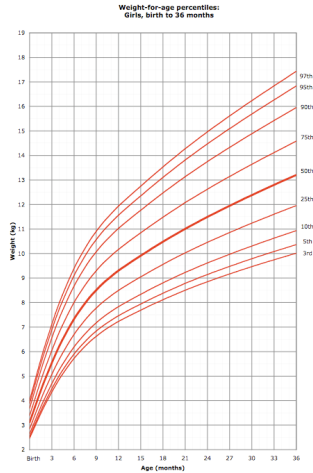
```
lmsqreg.fit(YY, TT, edf = c(3, 5, 3), targlen = 50, pvec = c(0.05, 0.1, 0.25, 0.5,  
0.75, 0.9, 0.95))
```

**targlen** - Number of points at which smooth estimates of L, M, S should be extracted for quantile plotting.



$$\hat{Y} = X [X'X]^{-1} X'Y$$

## Averaged Chart



R package `lmsqreg` (mw3)

```
> mw3
```

```
lms quantile regression, version , fit date Thu Jun  4 14:37:52 2009
```

```
Dependent variable: gdata$weight , independent variable: gdata$agey
```

```
The fit converged with EDF=( 4,6,3 ), PL= 9198.316
```

```
nominal percentile    0.030 0.050 0.10 0.25 0.500 0.750 0.900 0.95 0.970
```

```
estimated percentile 0.025 0.052 0.09 0.24 0.506 0.755 0.905 0.95 0.972
```

```
KS tests: (intervals in gdata$agey //p-values)
```

(-0.001,0]	(0,0.348]	(0.348,0.802]	(0.802,1.54]	(1.54,3]	Overall
0.000	0.000	0.271	0.324	0.676	0.001

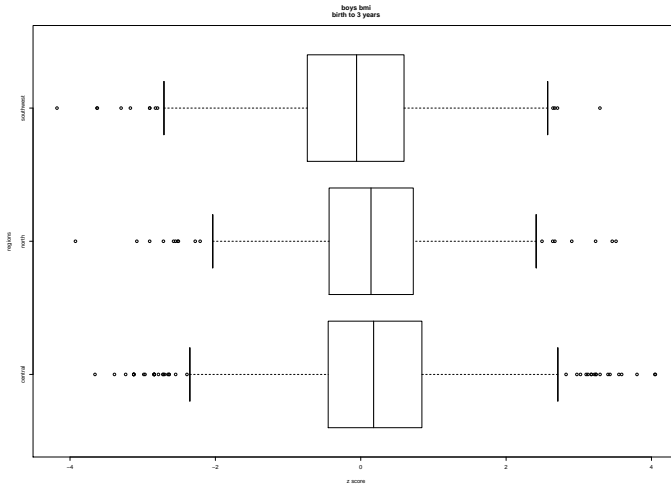
```
t tests: (intervals in gdata$agey //p-values)
```

(-0.001,0]	(0,0.348]	(0.348,0.802]	(0.802,1.54]	(1.54,3]	Overall
0.006	0.000	0.562	0.369	0.568	0.810

```
X2 tests (unit variance): (intervals in gdata$agey //p-values)
```

(-0.001,0]	(0,0.348]	(0.348,0.802]	(0.802,1.54]	(1.54,3]	Overall
0.000	0.000	0.717	0.050	0.462	0.979

## BoxPlots of mean SD scores of the three geographical regressions



## R analysis of variance (ANOVA)

```
> summary(fm<-aov(z~group))
              Df Sum Sq Mean Sq F value    Pr(>F)
group          2   69.4    34.7  37.589 < 2.2e-16 ***
Residuals    3941 3640.0     0.9
---
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

> TukeyHSD(fm)
  Tukey multiple comparisons of means
    95% family-wise confidence level

Fit: aov(formula = z ~ group)

$group
              diff             lwr             upr             p adj
north-central  -0.01495461 -0.1000653  0.07015605  0.9107089
southwest-central -0.33239976 -0.4255862 -0.23921328  0.0000000
southwest-north  -0.31744515 -0.4224195 -0.21247080  0.0000000
```

## Procedure for using ANCOVA to compare the growth standards between the regions

- 1 STEP 1: Find the best fitting polynomials having the lowest possible common degree for each of the three regions.
- 2 STEP 2: We want to answer the question "Is a common polynomial of the same degree as found in STEP 1 appropriate for all three regions or do the polynomials vary with region?" ie. for a particular measurement, sex and age group we want to test:

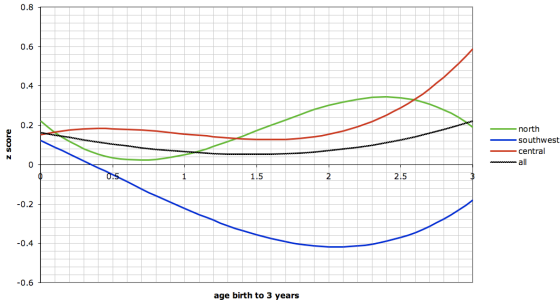
$H_0 : E[z | age] = \beta_0 + \dots + \beta_q age^q$  for each region, where  $q \leq 3$  is the degree of the common best fitting polynomial.

vs.  $H_1$  : The polynomial for at least two regions differ.

- 3 STEP 3: After finding a significant result in STEP 2 carry out pairwise comparisons between the regions.

## SD score regression models

**Boys BMI 0-3**

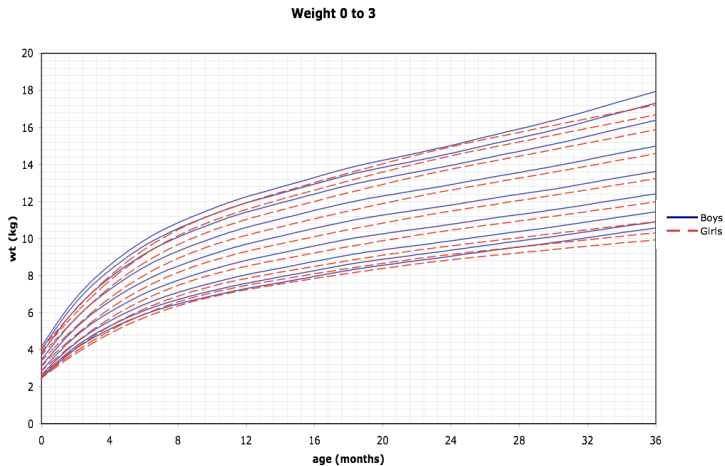


sex: male, age: birth to 36 months

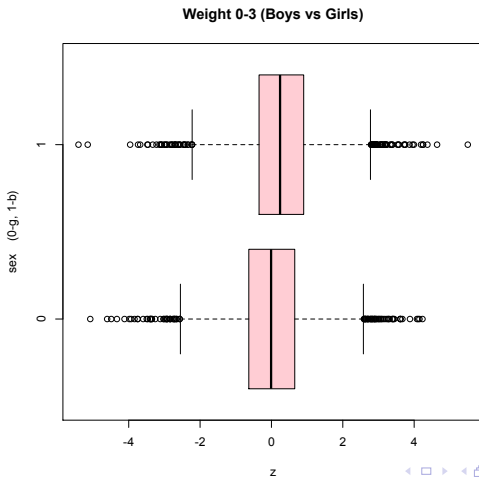
body mass index

	<i>p</i>
north-central	0.061050
southwest-central	0.000000
southwest-north	0.000000

## Weight, age birth to 36 months



## Box Plots



## t-test

```
> t.test(z~sex)
```

```
Welch Two Sample t-test
```

```
data: z by sex
```

```
t = -14.4148, df = 12473.99, p-value < 2.2e-16
```

```
alternative hypothesis: true difference in means is not equal to 0
```

```
95 percent confidence interval:
```

```
-0.2958223 -0.2249998
```

```
sample estimates:
```

```
mean in group 0 mean in group 1  
0.003078598      0.263489639
```

## lm function used to fit linear models

```
> m<-lm(z~sex+sex*x+sex*I(x^2)+sex*I(x^3))
> summary(m)
```

Call:

```
lm(formula = z ~ sex + sex * x + sex * I(x^2) + sex * I(x^3))
```

Residuals:

	Min	1Q	Median	3Q	Max
	-5.727730	-0.642032	-0.002139	0.649591	5.140159

Coefficients:

	Estimate	Std. Error	t value	Pr(> t )
(Intercept)	-0.0003369	0.0234113	-0.014	0.988518
sex1	0.1680739	0.0326333	5.150	2.64e-07 ***
x	0.1004580	0.1011812	0.993	0.320801
I(x^2)	-0.1245990	0.0993347	-1.254	0.209744
I(x^3)	0.0325240	0.0248738	1.308	0.191047
sex1:x	0.6006027	0.1418117	4.235	2.30e-05 ***
sex1:I(x^2)	-0.5300762	0.1395383	-3.799	0.000146 ***
sex1:I(x^3)	0.1160172	0.0350394	3.311	0.000932 ***

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

Residual standard error: 1.007 on 12473 degrees of freedom

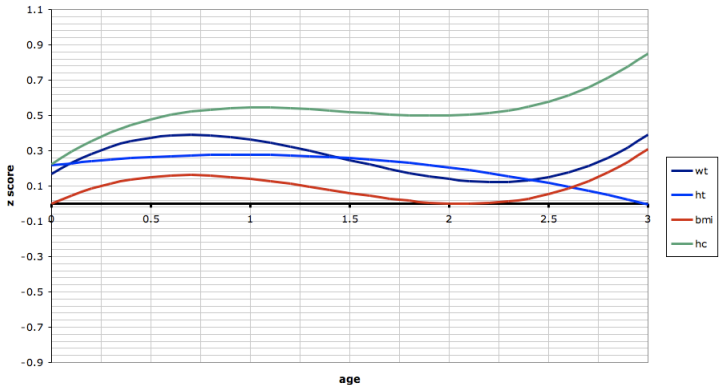
Multiple R-Squared: 0.02064, Adjusted R-squared: 0.02009

F-statistic: 37.56 on 7 and 12473 DF, p-value: < 2.2e-16



## Boys vs Girls, age birth to 3 years

### Z Scores Boys vs Girls



## Things to do

- 1 Assessing the difference in fits of quantiles fitted by a parametric function and by a smooth non-parametric curve.
- 2 Test for a significant difference between the curves for the overlapping period with the original estimations.

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