Chapter 1 Basic Biostatistics

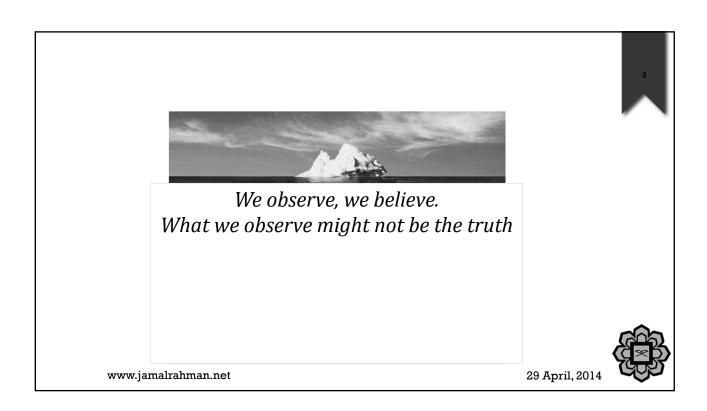
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Department of Community Medicine
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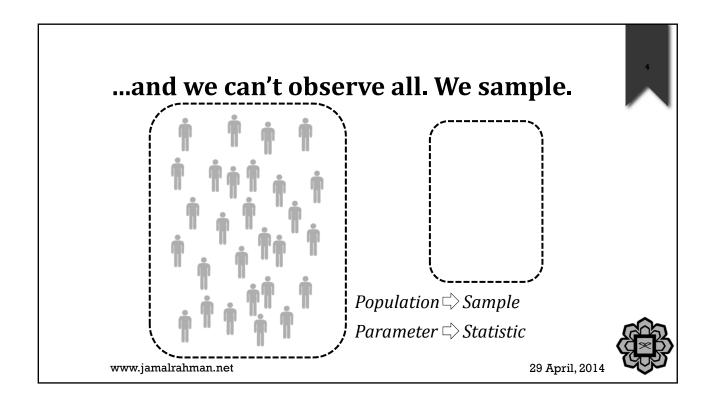


Content

- Basic premises variables, level of measurements, probability distribution
- Descriptive statistics
- Inferential statistics, hypothesis testing

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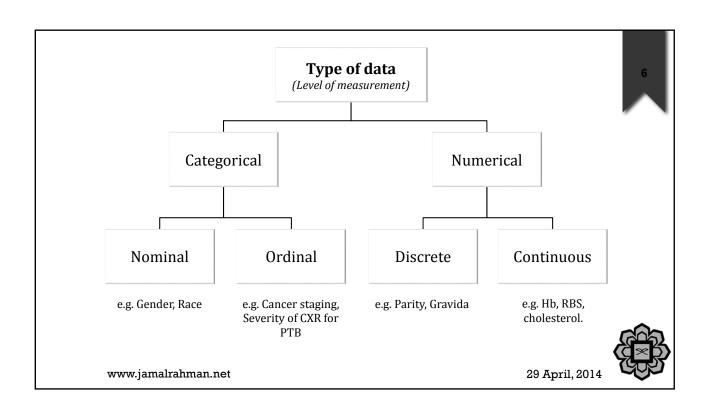


Variable

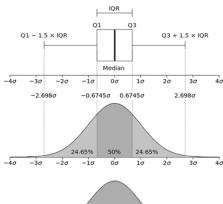
- Characteristic of a population
- Can take different values
- Data = measurements collected/observed

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$$f(x; \mu, \sigma^2) = \frac{1}{\sqrt[\sigma]{2\pi}} e^{-\frac{1}{2}(\frac{x-\mu}{\sigma})^2})$$

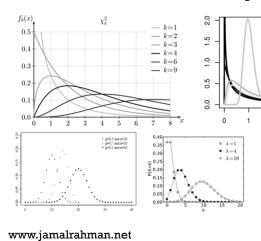


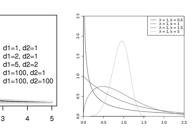
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Other distributions

Discrete vs. continuous probability distribution





χ², F, Weibull, Binomial & Poisson



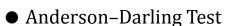
Characteristics of Normal distribution

- Smooth, symmetrical (around the mean), uni-modal, bell shaped curve
- Mean = Median = Mode
- \bullet Skewness = 0
- Kurtosis = 0

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Test of Normality



- Corrected Kolmogorov-Smirnov Test (Lilliefors Test)
- Cramér-von-mises Criterion
- D'agostino's K-squared Test
- Jarque-Bera Test
- Pearson's Chi-square Test
- Shapiro-Francia
- Shapiro-Wilk Test

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Use Normality test with caution



- Small samples almost always pass a normality **test.** Normality tests have little power to tell whether or not a small sample of data comes from a Gaussian distribution.
- With large samples, minor deviations from normality may be flagged as statistically significant, even though small deviations from a normal distribution won't affect the results of a t test or ANOVA.



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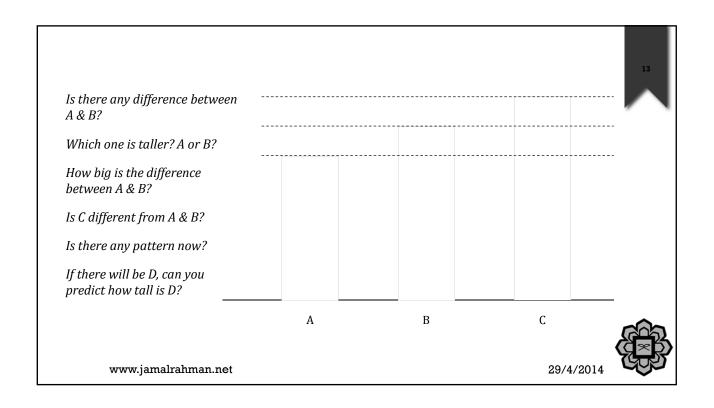
Statistical objectives

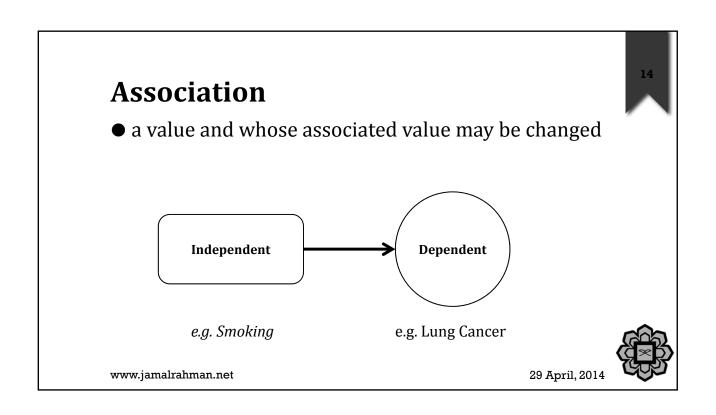


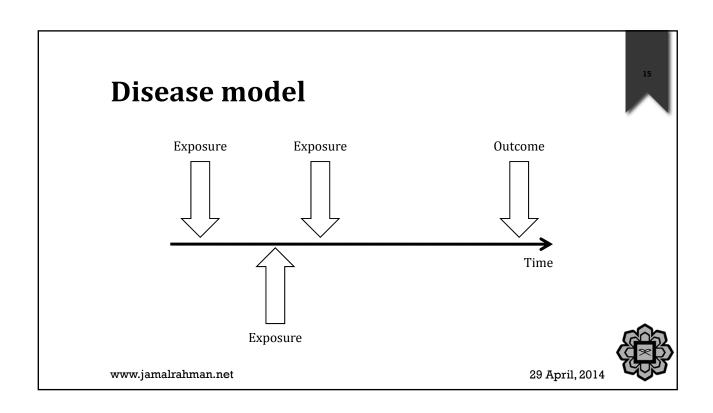
- 1. Determine presence of difference (or similarity)
- 2. Determine degree of difference
- 3. Determine the direction of changes (outcome)
- 4. Predict changes (outcomes)

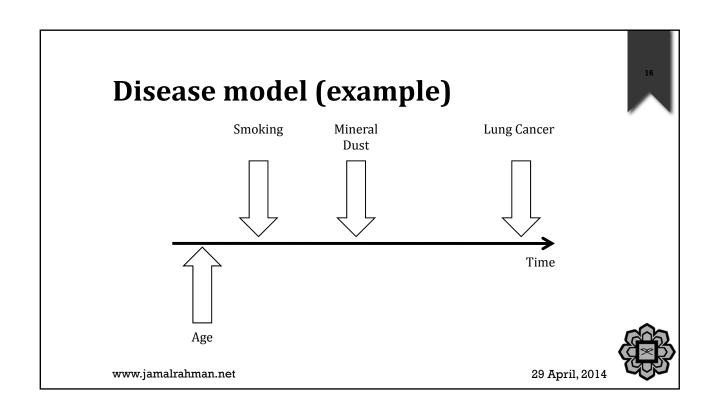


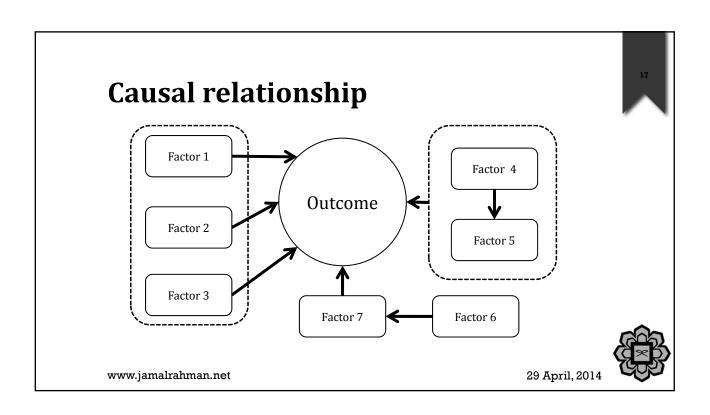
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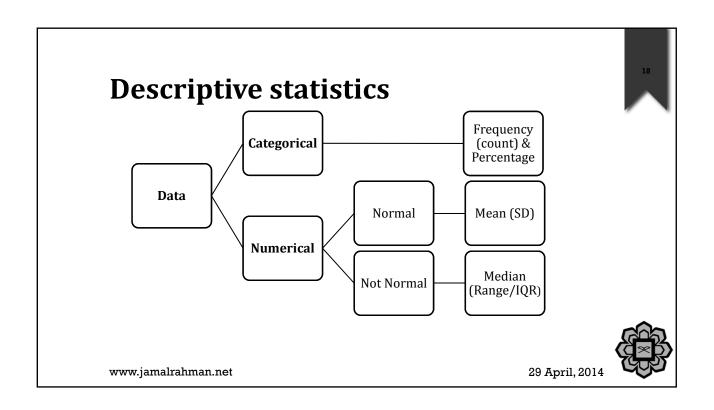












Hypothesis Testing

- Involve more than one variables
 - exposure & outcome,
 - predictor & criterion,
 - risk & disease
- Try to prove that Exposure causes the Disease e.g. Smoking causing Lung Cancer
- Example ~ Ho: No difference of risk to get Lung Cancer between smoker and non-smoker



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	Lung Cancer	No Lung Cancer
Smoking	20 (18.2%)	90 (81.8%)
Not Smoking	5 (4.5%)	105 (95.5%)

 χ^2 (df=1)= 10.150, p =0.001, OR = 4.7 (CI95% 1.7 – 13.0)

Because p < 0.05, we reject H_0 . Therefore there is a different between smoker & non smoker



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Statistical Test

● Univariate ~ One dependent & one independent

 Multivariate ~ Multiple dependent & multiple independent variable 41

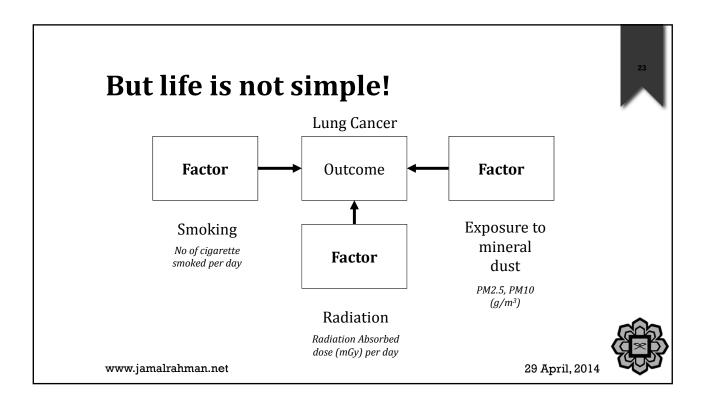
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What test to use?

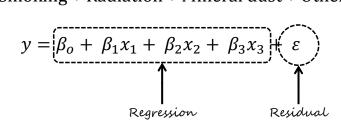
Variable 1	Variable 2	Test
Categorical	Categorical	Chi-square
Categorical (2 pop)	Numerical (Normal)	Independent sample t-test
Categorical (2 pop)	Numerical (Not Normal)	Mann-Whitney U test
Categorical (> 2 pop)	Numerical (Normal)	One-way ANOVA
Categorical (> 2 pop)	Numerical (Not Normal)	Kruskal-Wallis test
Numerical (Normal)	Numerical (Normal)	Pearson Correlation Coefficient Test
Numerical (Normal/ Not Normal)	Numerical (Not Normal)	Spearman Correlation Coefficient Test
Numerical (Normal)	Numerical (Normal) – Paired	Paired t-test
Numerical (Not Normal)	Numerical (Not Normal) – Paired	Friedman test

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The multivariate model

Lung CA = Smoking + Radiation + Mineral dust + Others



A good model is when Regression > Residual

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Multivariate Analysis

- Hypothesis testing & control for confounders
 - e.g. General Linear Model, Logistic Regression
- Modeling
 - e.g. Linear Regression
- Data reduction
 - e.g. Factor Analysis, Cluster Analysis



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Writing plan for statistical analysis #1

Data were analyzed using the complex sample function of SPSS (version 13.0). Sampling errors were estimated using the primary sampling units and strata provided in the data set. Sampling weights were used to adjust for nonresponse bias and the oversampling of blacks, Mexican Americans, and the elderly in NHANES. The prevalence of hypertension, as well as the awareness, treatment, and control rates, were age adjusted by direct standardization to the US 2000 standard population. 10 To analyze differences over time, the 2003-2004 data were compared with the 1999–2000 data. Estimates with a coefficient of variation >0.3 were considered unreliable. A 2-tailed P value < 0.05 was considered statistically significant.

(Ong et al. 2009)

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Writing plan for statistical analysis #2

To assess the effect of the selection process on the characteristics of the cases, we compared cases included in the final analysis to the rest of the cases. Since controls included in the present analysis were different from the rest of the diabetes free participants by design, no similar comparisons were performed for that group. To compare baseline characteristics of cases and controls appropriate univariate statistics were used. Similar binary logistic and multiple linear regression models were built with incident diabetes or HbA1c as respective outcomes and additive block entry of adiponectin and potential confounders. For linear regression CRP and triglycerides were log transformed. Since HbA1c could be modified by drug treatment, we ran a sensitivity analysis excluding all participants on antidiabetic medication. A p-value of <0.05 was considered significant. Analyses were performed with SPSS 14.0 for Windows.



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Reporting analysis (example)

TABLE 1. Characteristics of the cohort No known diabetes Known diabetes Total P value 37.28 (422) 35.22 (342) 62.72 (710) 64.78 (629) Admissions 1132 971 6.96 ± 2.08 31.80 (360) 156.52 ± 125.01 218.32 ± 150.13 37.90 (429) Patients HbA1c (%)
HbA1c ≥7.0 (yes)
Admission glucose (mg/dl)
Maximum glucose (mg/dl)
Glucose ≥200 mg/dl (yes) 64.78 (629) 6.05 ± 0.87 9.44 (67) 118.39 ± 52.65 158.48 ± 87.85 17.61 (125) 35.22 (342) 8.49 ± 2.56 69.43 (293) 220.68 ± 175.32 318.98 ± 177.09 72.04 (304) < 0.001 <0.001 <0.001 <0.001 <0.001 56.62 ± 18.16 50.70 (360) 61.17 ± 14.70 47.63 (201) 58.32 ± 17.09 49.56 (561) Age (yr) 0.001 Race/ethnicity 30.57 (129) 9.95 (42) 41.23 (174) 18.25 (77) 76.25 (321) 87.68 (370) 144.50 ± 25.93 76.90 ± 14.68 28.88 ± 7.46 98.04 ± 43.52 45.45 ± 16.94 161.19 ± 187.01 0.04 27.32 (194) 15.35 (109) 41.97 (298) Black 28.53 (323) White Hispanic Other 13.34 (151) 41.70 (474) 16.43 (186) Prior medication/hospital/clinic (ves) < 0.001 63.66 (452) 68.35 (773) 65.99 (747) Prior medication/nosp HTN (yes) Systolic BP (mm/Hg) Diastolic BP (mm/Hg) BMI (kg/m²) LDL (mg/dl) HDL (mg/dl) HDL (mg/dl) 53.66 (452) 53.10 (377) 136.51 ± 23.75 76.43 ± 15.12 27.55 ± 7.62 103.81 ± 42.81 50.70 ± 20.88 118.60 ± 89.33 <0.001 <0.001 5.99 (747) 139.50 ± 24.88 76.61 ± 14.95 28.04 ± 7.58 101.48 ± 43.16 48.54 ± 19.52 136.09 ± 139.56 Triglycerides (mg/dl) < 0.01

Data are presented as mean \pm so for continuous variables and percentage (number) for categorical variables based on total number of admissions Totals may not equal 100% due to rounding. P values were calculated by generalized estimating equations. HTN, Hypertension; BP, blood pressure; LDL, low-density lipoprotein; HDL, high-density lipoprotein.

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Reporting analysis (example)



Table 1. Sociodemographic Characteristics of the Participants

		Women (n=18	300)	Men (n=1281)		
Sociodemographic Characteristics	n	Unweighted, %*	Weighted, %*	n	Unweighted, %*	Weighted, %*
Place of residence						
Urban	893	49.6	30.9	652	50.9	33.0
Rural	907	50.4	69.1	629	49.1	67.0
Age, y						
25 to 34	725	40.3	42.6	470	36.7	35.8
35 to 44	500	27.8	27.9	350	27.3	27.3
45 to 54	367	20.4	18.8	276	21.6	21.1
55 to 64	208	11.6	10.7	185	14.4	15.8
Education, y†						
None	685	38.1	44.7	196	15.3	21.2
1 to 4	642	35.7	37.3	423	33.1	36.1
5	191	10.6	7.5	248	19.4	19.4
6 to 7	164	9.1	6.1	199	15.6	12.2
≥8	115	6.4	4.4	213	16.6	11.2



Reporting analysis (example)

	Hypertension										
		Women				Men					
		Urban	Rural		Urban		Rural				
Participant Characteristics	%	∯5% CI	%	95% CI	%	95% CI	%	95% CI			
All participants	41.0	36.4 to 45.7	26.8	19.5 to 34.1	40.0	34.0 to 46.1	33.5	27.8 to 39.2			
Age, y											
25 to 34	17.6	12.7 to 22.5	11.1	5.9 to 16.2	31.8	25.0 to 38.6	32.8	26.4 to 39.2			
35 to 44	43.2	31.8 to 54.5	27.1	18.8 to 35.5	35.3	25.7 to 44.9	27.7	18.8 to 36.6			
45 to 54	69.5	57.6 to 81.3	45.5	35.6 to 55.4	49.8	39.6 to 60.1	32.0	20.3 to 43.7			
55 to 64	73.0	64.0 to 81.9	57.9	44.9 to 70.9	59.4	38.3 to 80.5	46.0	46.0 to 60.8			
Education, y											
0 to 4	45.8	38.0 to 53.6	27.0	19.7 to 34.3	42.7	26.3 to 59.1	33.7	26.1 to 41.3			
5 to 7	37.6	32.4 to 42.9	18.5	7.6 to 29.4	39.0	28.3 to 49.7	32.8	21.8 to 43.8			
≥8	27.2	20.8 to 33.6	46.8	4.6 to 89.0	38.3	29.5 to 47.2	35.0	4.0 to 66.0			
Body mass index, kg/m ²											
<25.0	33.0	29.4 to 36.7	25.6	18.7 to 32.4	34.4	29.3 to 39.6	30.5	23.8 to 37.2			
25.0 to 29.9	54.1	45.2 to 63.1	42.2	26.1 to 58.2	53.9	39.6 to 68.2	62.5	48.3 to 76.7			
≥30	54.9	44.5 to 65.3	31.8	8.4 to 55.1	78.6	67.0 to 90.2	89.7	66.6 to 100.			
Waist circumference, cm											
Women <8 and men <102	38.0	33.2 to 42.8	27.4	19.9 to 34.9	38.3	32.5 to 44.1	32.5	27.0 to 38.0			
Women ≥8 and men ≥102	60.0	50.4 to 69.6	50.9	26.1 to 75.7	79.3	65.3 to 93.2	100				
Current drinking											
No	40.4	33.8 to 47.0	24.4	16.2 to 32.6	37.8	28.9 to 46.8	28.5	23.1 to 33.9			
<1 d/wk	40.3	33.6 to 47.0	33.0	23.3 to 42.8	43.4	33.2 to 53.6	35.4	20.7 to 50.2			
≥1 d/wk	48.2	33.1 to 63.3	34.6	20.1 to 49.1	38.6	27.7 to 49.5	40.5	31.4 to 49.6			

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^{*}Within each variable, the sum of the proportions may not be 100% because of rounding. †The sum of the number of participants in each category is <1800 for women and 1281 for men because of missing data.

Summary

- 1. Identify & define variables
- 2. Type independent vs. dependent
- 3. Level of measurements nominal, ordinal or continuous
- 4. Check distribution Normal vs. Not Normal
- 5. Decide what to do descriptive vs. analytical

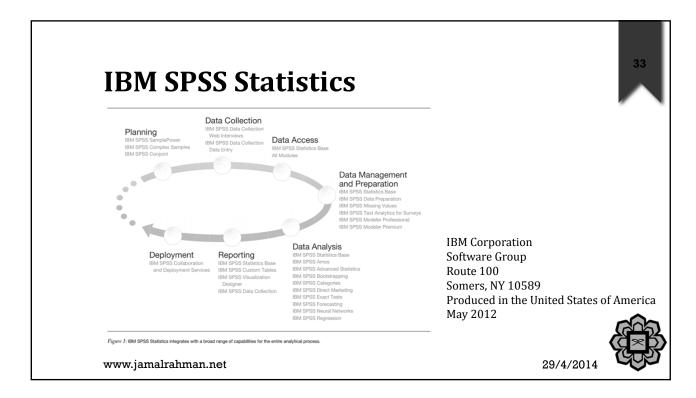


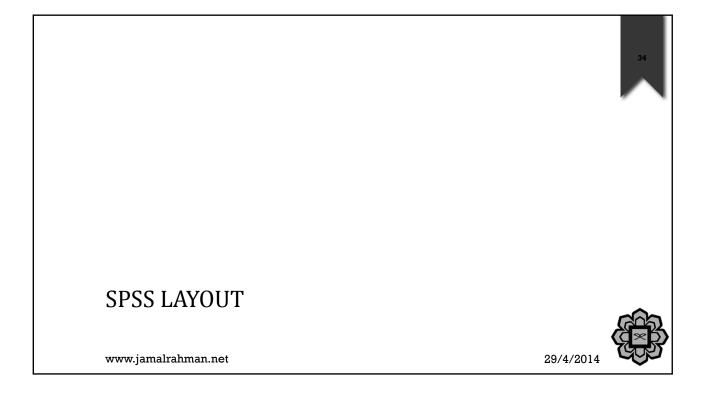
Chapter 2 **Introduction to SPSS**

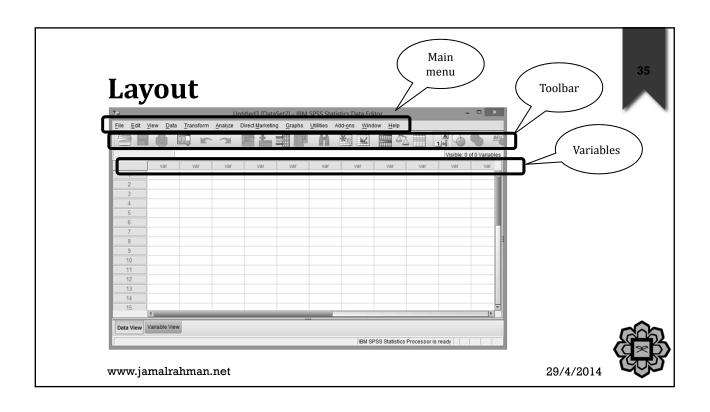
IBM SPSS Statistics v21 for Windows

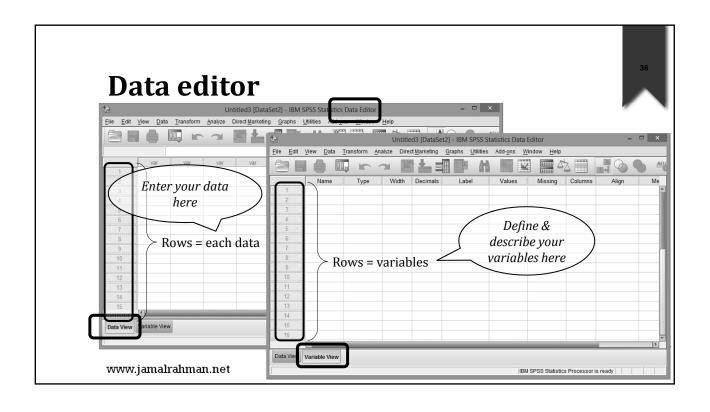
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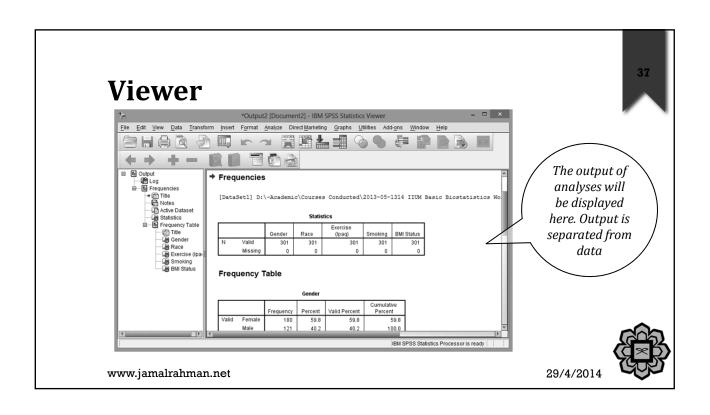


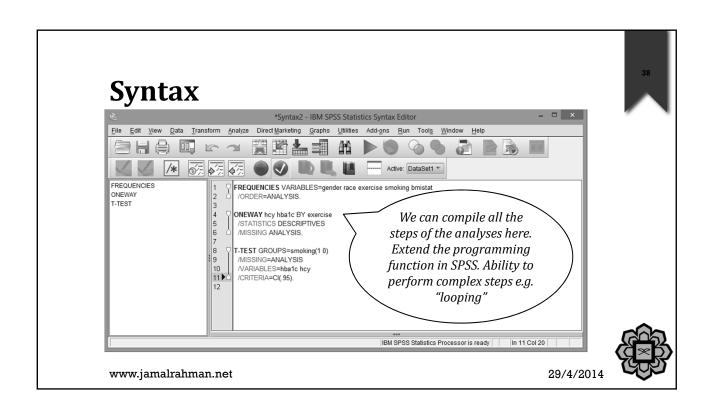












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CREATING DATASET

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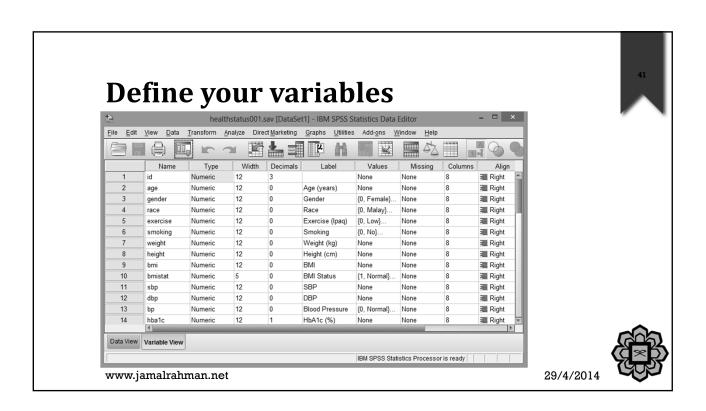
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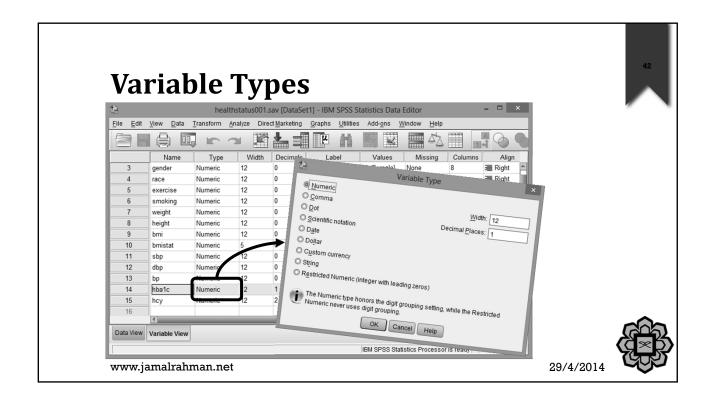
Before even you start SPSS!

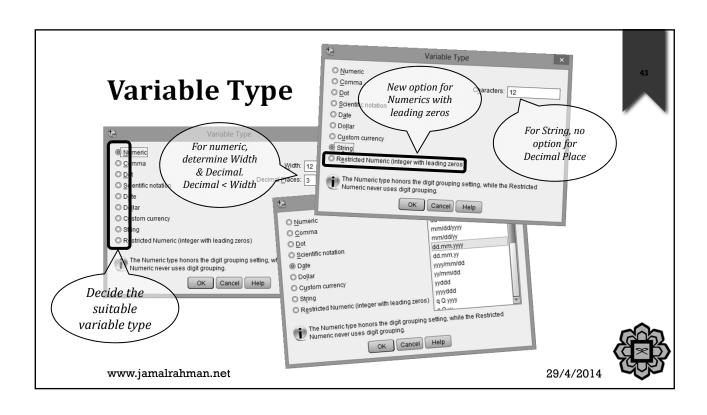
- You must identify & define relevant variables
- Define means
 - 1. Name preferably short single name, begins with alphabet, no special character, no space
 - 2. Type of data e.g. Numeric, Date, String
 - 3. Width & Decimal Places (if numeric)
 - 4. Label description for the Name (will be displayed in Viewer)
 - 5. Values labels for value e.g. 1=Male, 2=Female
 - 6. Missing define missing value e.g. 999 for N/A

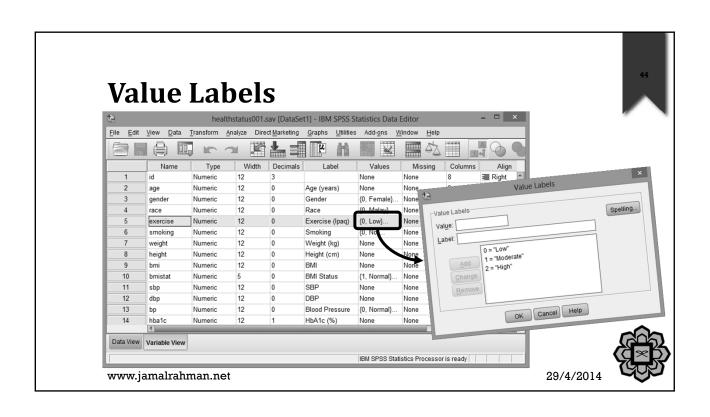


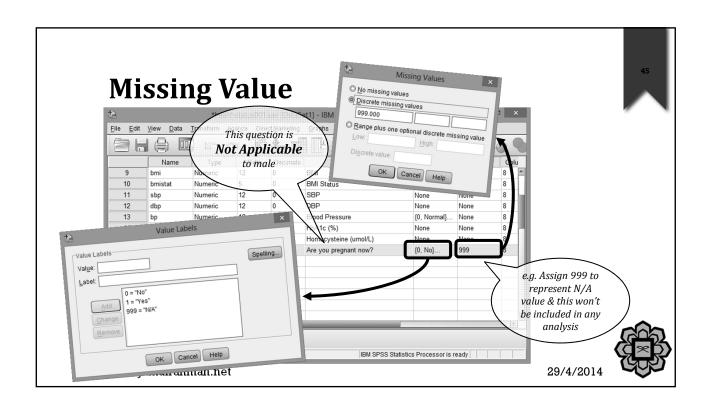
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Chapter 3 **Descriptive Statistics**

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Exercise data

- Hypothetical
- Study to describe factors related to HbA1c & Homocystein (HCY)
- N=301
- 13 variables

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Retrieve file information healthstatus001.sav [DataSet1] - IBM SPSS Statistics Data Editor <u>File Edit View Data Transform Analyze Direct Marketing Graphs Utilities Add-ons Window Help</u> m <u>O</u>pen Visible: 15 of 15 Variables Open Data<u>b</u>ase Read Text Data... Chinese @ Read Cognos Data... Low 31 OI OI No Close emale Indian Moderate No 154 32 Save Male Chinese Low Yes 78 154 33 Save As... 19 Male Malay High Yes 40 Save All Data Moderate OI OI Overw 🗟 Export to Database... Male Malay Moderate 33 Mark File Read Only emale Malay Moderate No 69 154 29 Male 26 Malay High No 65 159 mi Rename Dataset... Moderate Display Data File Information Cache Data... Moderate No 86 158 35 01 😱 Switch Server.. Repository IBM SPSS Statistics Processor is ready

healthstatus001

Variable	Position	Label	Measurement Level	Role	Column Width	Alignment	Print Format	Write Format
id	1	<none></none>	Scale	Input	8	Right	F12.3	F12.3
age	2	Age (years)	Scale	Input	8	Right	F12	F12
gender	3	Gender	Nominal	Input	8	Right	F12	F12
race	4	Race	Nominal	Input	8	Right	F12	F12
exercise	5	Exercise (lpaq)	Nominal	Input	8	Right	F12	F12
smoking	6	Smoking	Nominal	Input	8	Right	F12	F12
weight	7	Weight (kg)	Scale	Input	8	Right	F12	F12
height	8	Height (cm)	Scale	Input	8	Right	F12	F12
sbp	9	SBP	Scale	Input	8	Right	F12	F12
dbp	10	DBP	Scale	Input	8	Right	F12	F12
hba1c	11	HbA1c (%)	Scale	Input	8	Right	F12.1	F12.1
hcy	12	Homocystein e (umol/L)	Scale	Input	8	Right	F12.2	F12.2

Value		Label		
gender	0	Female		
	1	Male		
race	0	Malay 🖟		
	1	Chinese		
	2	Indian		
exercise	0	Low		
	1	Moderate		
	2	High		
smoking	0	No		
	1	Yes		

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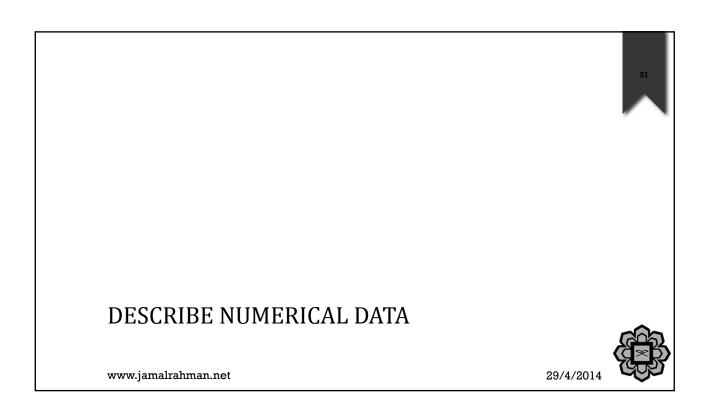
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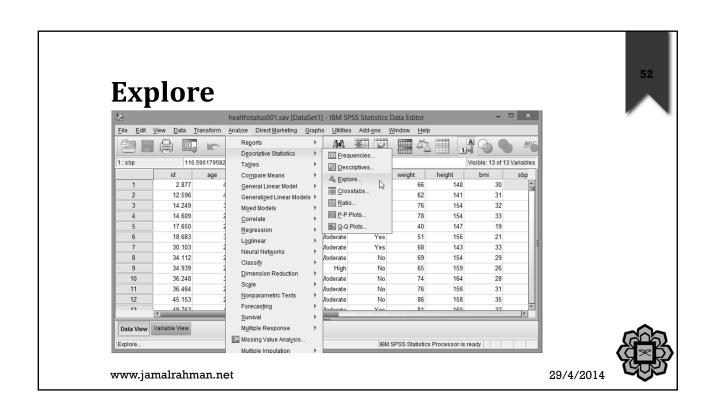


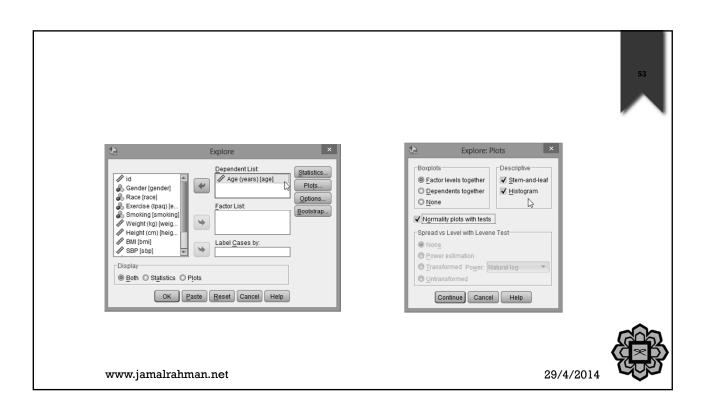
Task 1

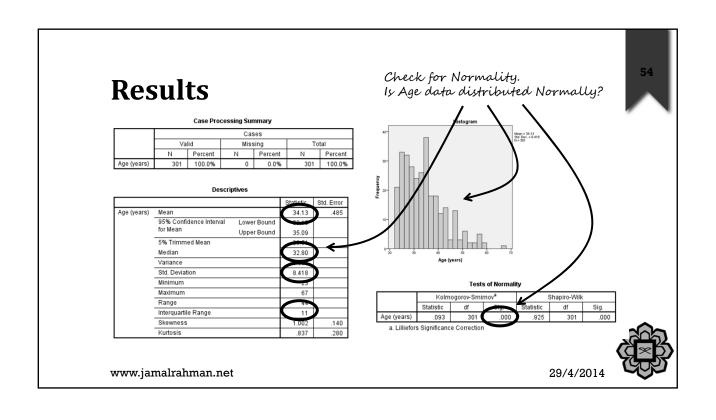
- 1. Describe socio-demographic characteristics of the respondent (age, gender & race)
- 2. Describe the explanatory variables
 - 1. Exercise
 - 2. smoking status
 - 3. BMI status &
 - 4. BP status
- 3. Describe HbA1c (taking cut-off for Poor HbA1c ≥ 6.5%) & HCY

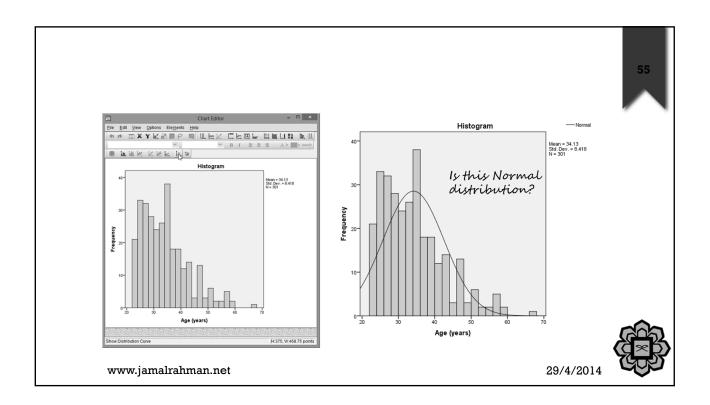
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Describe age

Normal

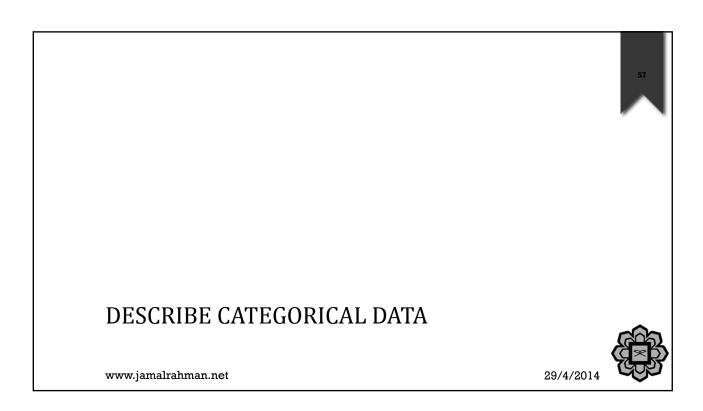
• The subjects distributed between 23-67 years old with the average of 34 (SD=8) years.

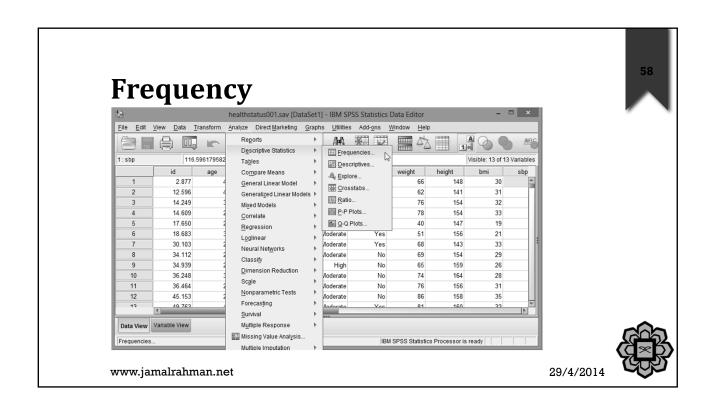
If not Normal

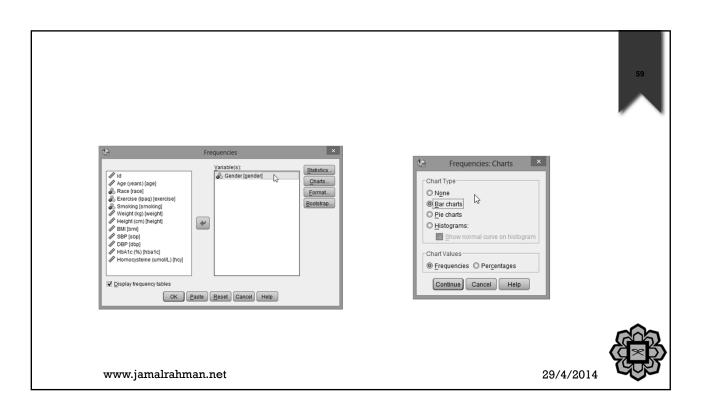
 The subjects distributed between 23-67 years old with the median of 33 (IQR=11) years

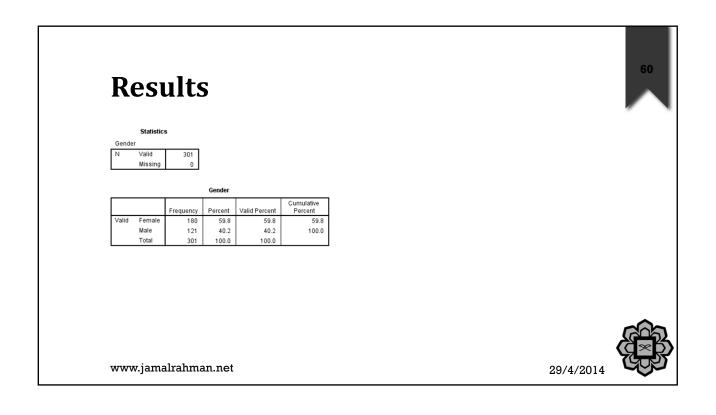
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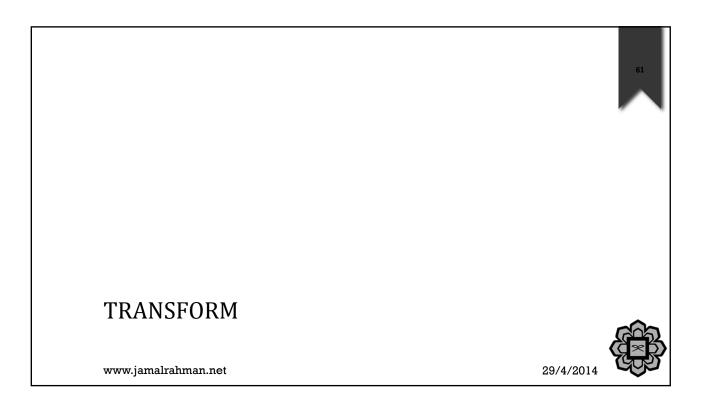
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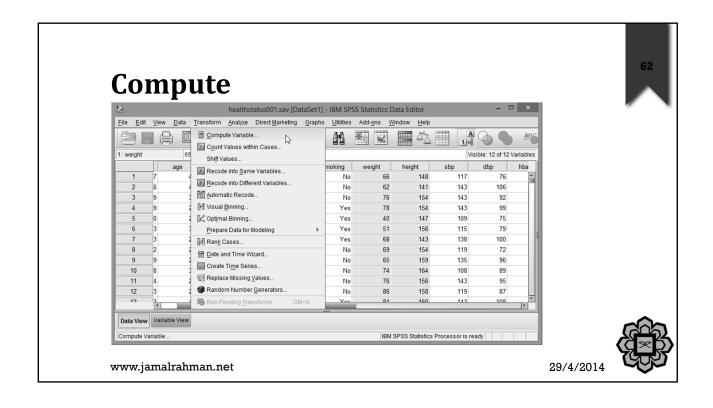


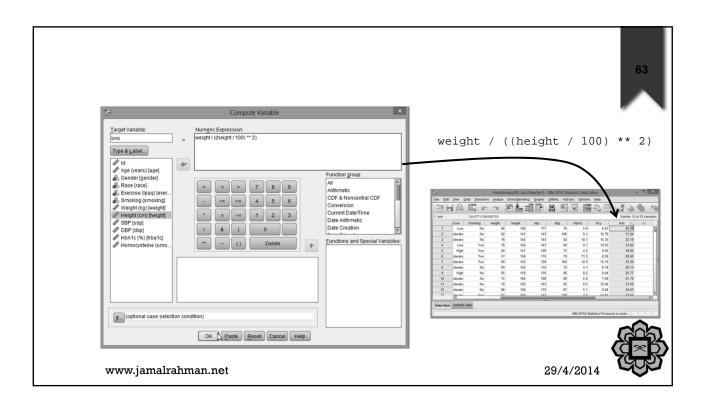


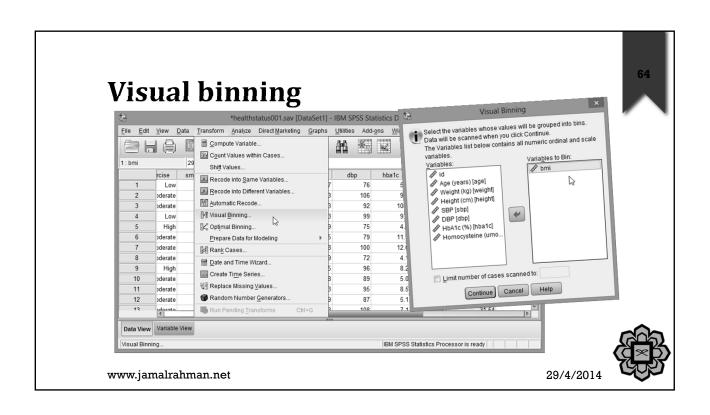


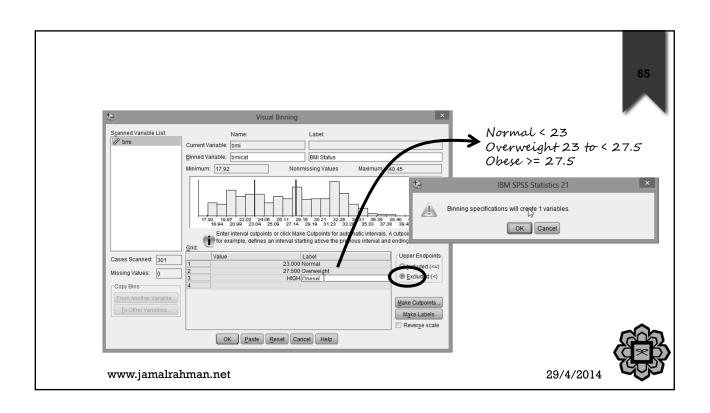


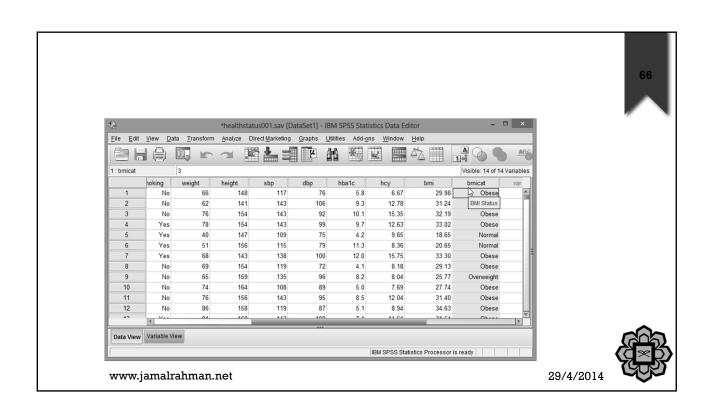












Chapter 4 Bivariable analyses IBM SPSS Statistics v21 for Windows

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To check association of two variables? HbA1c Age 29/4/2014 www.jamalrahman.net

The steps

- 1. Determine which is dependant & which is independent
- 2. Determine level of measurements
- 3. Determine Normality of the numerical measurement
- 4. Determine the suitable statistical test



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What are the tests?

Variable 1	Variable 2	Test	
Categorical	Categorical	Chi-square	
Categorical (2 pop)	Numerical (Normal)	Independent sample t-test	
Categorical (2 pop)	Numerical (Not Normal)	Mann-Whitney U test	
Categorical (> 2 pop)	Numerical (Normal)	One-way ANOVA	
Categorical (> 2 pop)	Numerical (Not Normal)	Kruskal-Wallis test	
Numerical (Normal)	Numerical (Normal)	Pearson Correlation Coefficient Test	
Numerical (Normal/ Not Normal)	Numerical (Not Normal)	Spearman Correlation Coefficient Test	
Numerical (Normal)	Numerical (Normal) – Paired	Paired t-test	
Numerical (Not Normal)	Numerical (Not Normal) – Paired	Friedman test	
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Tasks 2



- 1. Determine association between socio-demographic characteristics & all the risk factors with HbA1c
- 2. Determine association between socio-demographic characteristics & all the risk factors with HCY

Note: It would be good if you could construct **dummy table** for the answers even before the analyses started



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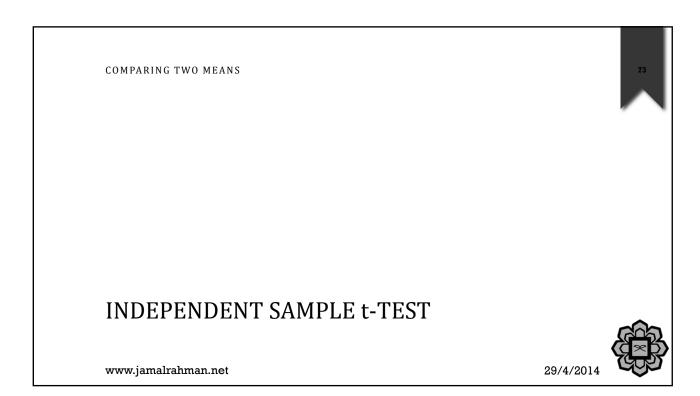
HCY normal range

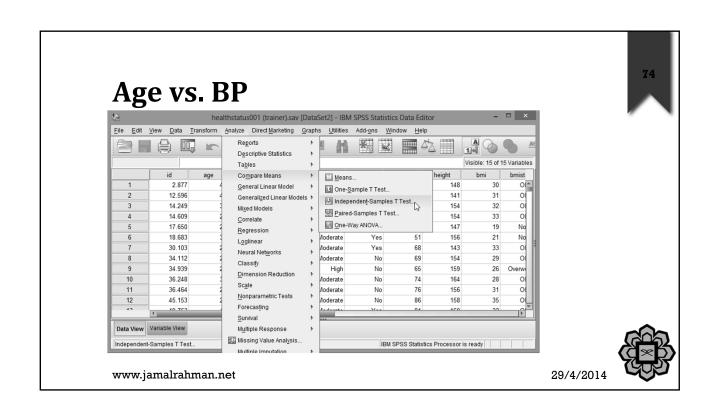


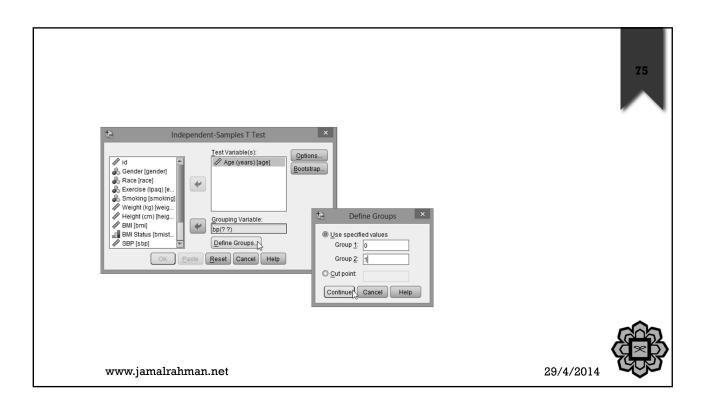
Blood reference ranges for homocysteine:

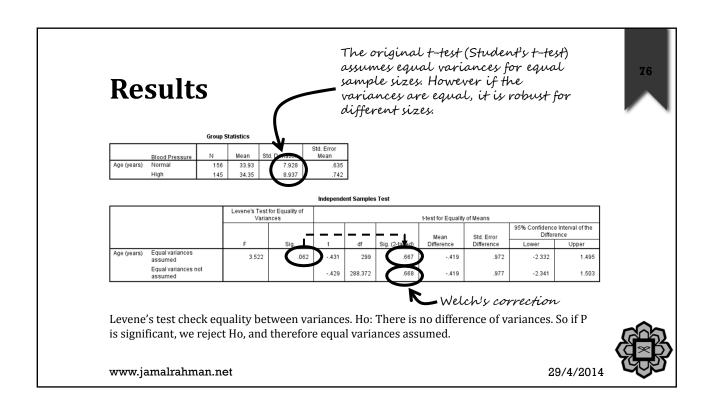
	blood reference ranges for nomocysteine:							
Sex	Age	Lower limit	Upper limit	Unit	Elevated	Therapeutic target		
	40.40	3.3 ^[12]	7.2 ^[12]	µmol/L				
CI-	12–19 years	45 ^[14]	100 ^[14]	μg/dL	> 10.4 µmol/L			
	>60 years	4.9 ^[12]	11.6 ^[12]	µmol/L	or > 140 μg/dl	< 6.3 µmol/L ^[13] or < 85 µg/dL ^[13]		
		66 ^[14]	160 ^[14]	μg/dL	r r v pgran			
	40.40	4.3 ^[12]	9.9 ^[12]	µmol/L	> 11.4 µmol/L			
M-I-	12-19 years	60 ^[14]	130 ^[14]	μg/dL				
Male		5.9 ^[12]	15.3 ^[12]	µmol/L	or > 150 μg/dL			
>60 ye	>60 years	>60 years 80 ^[14]	210 ^[14]	μg/dL	, or pgrac			

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Table - Distribution of age by blood pressure status

	N	Mean	SD	Statistics	df	P
Normal BP	156	33.9	7.9	t=0.431	299	0.667
High BP	145	34.4	8.9			



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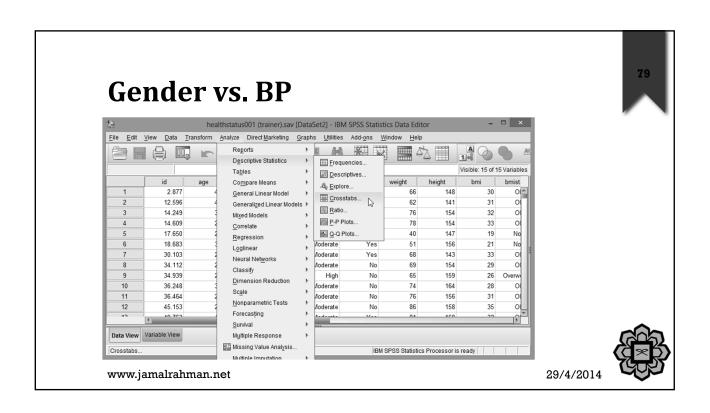
DIFFERENCE OF TWO PROPORTIONS

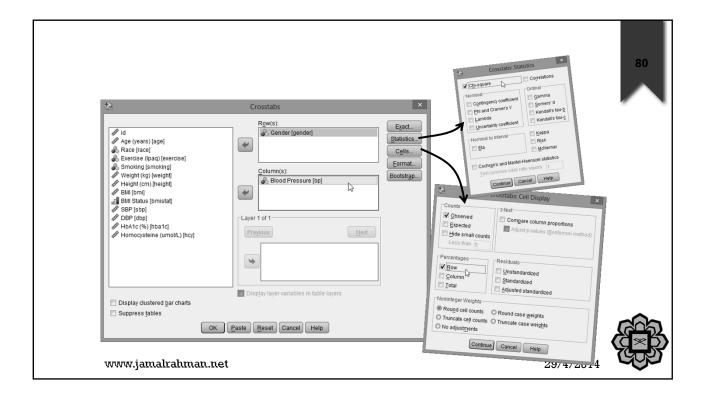
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CHI-SQUARED TEST

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Results

Some books may suggest the use of Continuity Correction at ALL time, but recent simulations showed that CC (or Yate's correction) is OVERCONSERVATIVE. Hence, use Pearson χ^2 when < 20% of cells

Case Processing Summary

Cases						
Valid		Missing		Total		
N	Percent	N	Percent	N	Percent	
301	100.0%	0	0.0%	301	100.0%	
	N	N Percent	Valid Miss	Valid Missing N Percent N Percent	Valid Missing To N Percent N Percent N	

Gender*	Blood Pres	sure	301	100.0%	0	0.0%	301	100.0
	Gen	der * B	lood Pressu	re Crosstab	ulation			
				Blood P	ressure		1.	
				Normal	High	Total		
Gender	Female	Coun	ıt	92	88	180	1	
		% wit	hin Gender	51.1%	48.9%	100.0%		Pears
	Male	Coun	ıt	64	57	121	1	Conti
		% wit	hin Gender	52.9%	47.1%	100.0%		Likelih
Total		Coun	ıt	156	145	301	1	Fishe
		% wit	hin Gender	51.8%	48.2%	100.0%] [Linea

Chi-Square Tests

have expected count < 5

	Value	df	Asymp. Sig. (2-sided)	Exact Sig. (2- sided)	Exact Sig. (1- sided)
Pearson Chi-Square	.092ª	1	.762		
Continuity Correction ^b	.034	1	959		
Likelihood Ratio	.092	1	.762		
Fisher's Exact Test				.814	.426
Linear-by-Linear Association	.092	1	.762		
N of Vallet Cases	301				

e expected count less than 5. The minimum expected count is 58.29.

Describe this table first. What is your impression? 49% women vs. 47% men with high BP

When \geq 20% of cells have EC < 5, use Fisher's Exact Test



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This is given because we code the variables using numbers. Can be used to measure P-trend

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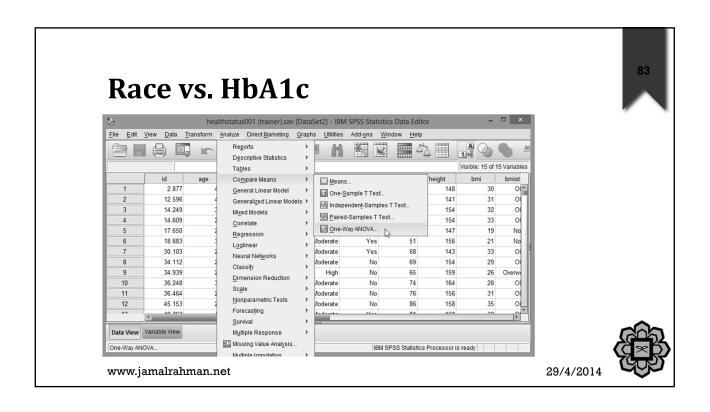
COMPARING MORE THAN TWO MEANS

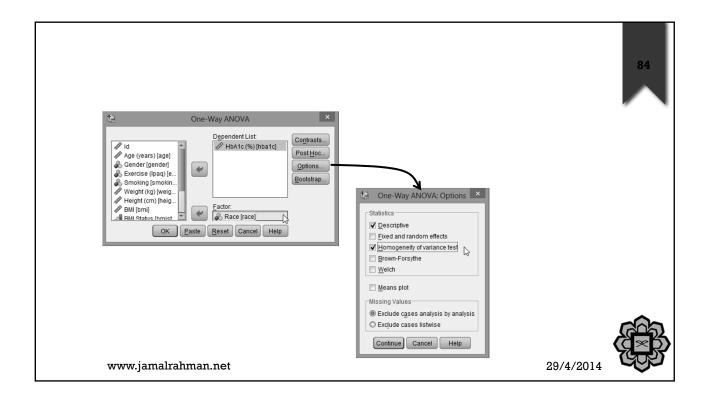
ONE-WAY ANOVA

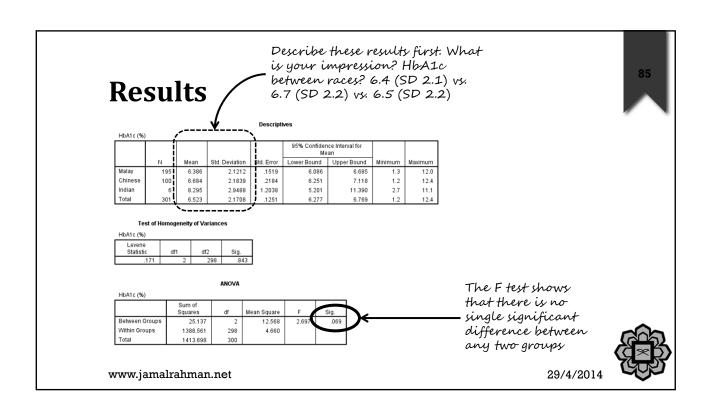
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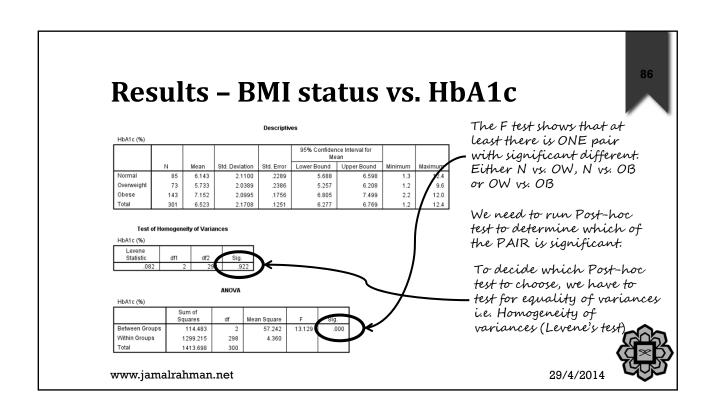


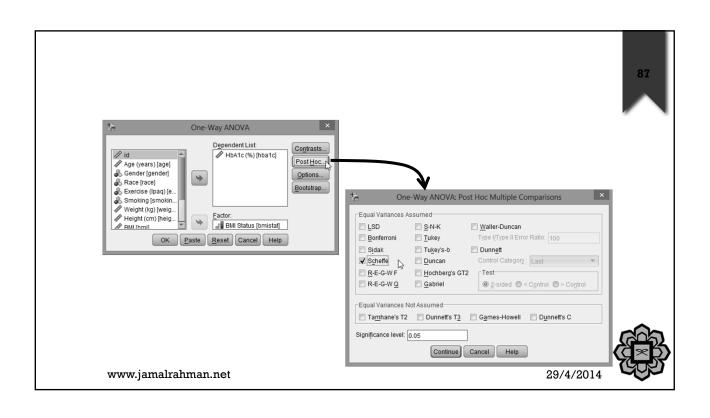
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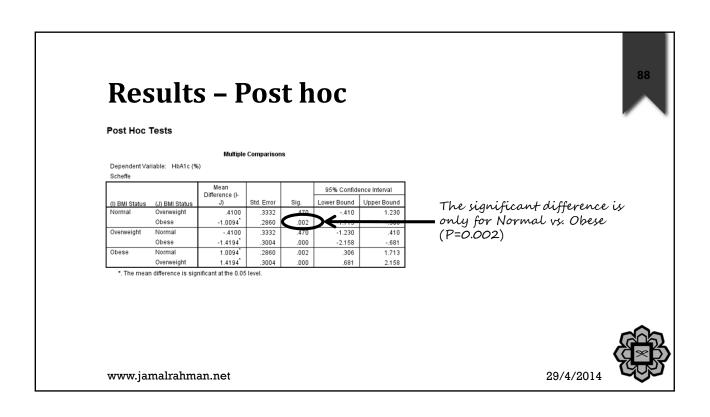












Report

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There is a significant association between BMI Status and HbA1c (F(2,298)=13.129, P<0.001). Post-hoc test showed that Obese subjects have significantly higher HbA1c compared to Normal and Overweight subjects (P=0.001 and P<0.001 respectively).



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NON PARAMETRIC TESTS

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