\mathbf{r}		-		
ĸ	03	<u>a</u>	เท	0
1/	ca	u.	ш	ے

Nature Biotechnology **26**, 303 - 304 (2008) doi:10.1038/nbt0308-303

What is principal component analysis?

Markus Ringnér1

Principal component analysis is often incorporated into genome-wide expression studies, but what is it and how can it be used to explore high-dimensional data?

PCA as a tool to Quantify and Visualise

1

Multivariate Analysis

Multiple Regression/two way-ANOVA/mixed models /machine learning algorithms

Ordination methods

2

What is the difference between these two pairwise correlation matrices?

	X_1	X ₂	X ₃	X_4	X ₅
X1	1.00	0.80	0.90	0.78	0.87
X 2	0.80	1.00	0.76	0.87	0.78
Хз	0.90	0.76	1.00	0.78	0.89
X4	0.78	0.87	0.78	1.00	0.95
X 5	0.87	0.78	0.89	0.95	1.00

- 1			-	-	
	X_1	X ₂	X ₃	X ₄	X ₅
X 1	1.00	0.87	0.96	0.04	0.05
X ₂	0.87	1.00	0.95	0.03	0.07
Хз	0.96	0.95	1.00	0.04	0.05
	0.04	0.03	0.04	1.00	0.84
X 5	0.05	0.07	0.05	0.84	1.00

What is the difference between these two pairwise
correlation matrices?

	X_1	X_2	X ₃	X_4	X ₅
X1	1.00	0.80	0.90	0.78	0.87
X2	0.80	1.00	0.76	0.87	0.78
Хз	0.90	0.76	1.00	0.78	0.89
X4	0.78	0.87	0.78	1.00	0.95
X 5	0.87	0.78	0.89	0.95	1.00

One dimension

	X_1	X_2	. X ₃	X_4	. X ₅
X ₁	1.00 0.87	0.87 1.00	0.96 0.95	0.04 0.03	0.05 0.07
Хз	0.96	0.95	1.00	0.04	0.05
	0.04	0.03	0.04	1.00	0.84
X5	0.05	0.07	0.05	0.84	1.00

Two dimensions

4

Ordination analyses

- Uncover, organize and summarize the main patterns of variation in a set of variables measured over multiple observations.
- Patterns of variation are structured in a reduced space with smaller number number of dimensions.
- Reduction is possible because often variables are associated (e.g., correlated). Dimensions represent combinations (e.g., linear combinations of variables).

5

Ordination analyses

A procedure for adapting a multidimensional swarm of data points in such a way that when it is projected onto a reduced number of dimensions any intrinsic pattern will become apparent.

Adapted from Connie Clark

Ordination analyses - uncover and organiz
data; a quick example:

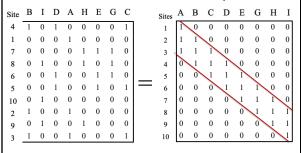
Species

Site	В	I	D	A	Н	Е	G	C
4	1	0	1	0	0	0	0	1
1	0	0	0	1	0	0	0	0
7	0	0	0	0	1	1	1	0
8	0	1	0	0	1	0	1	0
6	0	0	1	0	0	1	1	0
5	0	0	1	0	0	1	0	1
10	0	1	0	0	0	0	0	0
2	1	0	0	1	0	0	0	0
9	0	1	0	0	1	0	0	0
3	1	0	0	1	0	0	0	1

7

Ordination analyses – uncover and organize data; a quick example:

Species Species



8

Ordination methods

- Principal Component Analysis (PCA)
- Correspondence Analysis (CA)
- Principal Coordinate Analysis (PCoA)
- Discriminant Function Analysis (DFA)
- Principal Curve Analysis
- Etc, etc, etc...

Principal components analysis (PCA) is perhaps the most common technique used to summarize patterns among variables in multivariate datasets.



Some treat Principal Component Analysis (PCA) as an unsupervised learning method (an exploratory technique such as k-means) 10 Unsupervised Learning 373 10.1 The Challenge of Unsupervised Learning 373 10.2 Principal Components Analysis 374 10.1 What Are Principal Components 379 10.2 Another Interpretation of Principal Components 379 10.3 Chartering Methods 385 10.3 Chartering Methods 385 10.3 Practical Issues in Clustering 386 10.3 Practical Issues in Clustering 386 10.3 Practical Issues in Clustering 386 10.3 Practical Issues in Clustering 389 10.3 Practical

11

Supervised versus unsupervised learning techniques

- Techniques for unsupervised learning are fast growing in a number of fields, particularly biology.
- A cancer researcher might assay gene expression levels in 100 patients with breast cancer. They might then look for subgroups among the breast cancer samples, or among the genes, in order to obtain a better understanding of the disease.
- A search engine might choose what search results to display to a particular individual based on the click histories of other individuals with similar search patterns. These statistical learning tasks, and many more, can be performed via unsupervised learning techniques.

Adapted from James et al. 2013

Supervised versus unsupervised learning techniques

In contrast, unsupervised learning is often much more challenging. The exercise tends to be more subjective, and there is no simple goal for the analysis, such as prediction of a response.

Unsupervised learning is often performed as part of an exploratory data analysis.

 $\mbox{\sc Hard}$ to assess the results obtained given that there is no universally accepted mechanism for performing cross-validation or validating results on an independent data set; there is no way to check how the models does because we don't know the true answer—the problem is unsupervised.

Adapted from James et al. 2013

13

Examples of Principal Component Analysis



14

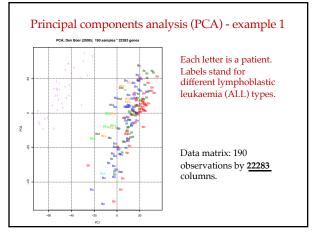
Principal components analysis (PCA) - example 1

A subtype of childhood acute lymphoblastic leukaemia with $\Re W$ poor treatment outcome: a genome-wide classification study



Quantification and Visualisation

	lhood acute lymphoblastic leukaemia with $ ightarrow rac{1}{2} ightarrow rac{1}{2}$ utcome: a genome-wide classification study					
Laura J CM Van Zutven, H Berna Beverlos	Maniquet Den Boer", Marjon van Septenberst", Renie X De Menzee, Moyling H Ored, Jenisco G C AM Beije - Gladdines, Souan T C J M Peter, Laury L Miron Johnen H Bernst Genedien, Pater J Van der Spek, Gally Eschwicht, Martin A Henchmannt, Gettas E Jordo Schwalet, Walters A Kazapu, Malinne E Gene, Bah Feller, 1991.					
children. 25% of precursor B-Al	Summary Badgoond Genetic toletypes of acute lymphoblastic leukemin [AL1] are used to determine risk and treatment in children 25% of personne FALL cases are genetically unclassified and have intermediate prognosis. We aimed to accommodate and accommodate and accommodate to the contractions are a generous-when only in improve prognosis classification of this findings.					
Data matri	x: 190 observations by 22283 columns					
	Gene expression (22283 genes)					
Gene expression (190 patients)						



17

Principal components analysis (PCA) - example 2



PCA – A Powerful Method for Analyze Ecological Niches

Franc Janžekovič and Tone Novak University of Maribor, Faculty of Natural Sciences and Mathematics, Department of Biology, Maribor Silocenia

Principal components analysis (PCA) - example 2

2.1 Environmental niche of three hymenopteran and two spider species

Between 1977 and 2004, 63 caves and artificial tunnels were ecologically investigated in Slovenia; the three most abundant Hymenoptera species found in these studies have been ecologically evaluated (details in Novak et al. 2010a). In the caves, many environmental data were collected, as follows. The following abbreviations of the environmental variables are used: Dist-E = distance from entrance; Dist-S = distance from surface; Illum = illumination; PCS = passage cross-section; Tair =air temperature; RH = relative air humidity; Tgr = ground temperature; HY = substrate moisture. The hymenopteran spatial niche breadth was originally represented by nine variables.

Data matrix: 63 observations (caves) by 9 columns

Environmental variables (9)

63 caves

Franc Jandokovič and Tone Novak aculty of Natural Sciences and Mathematics, Department of Ricings, Maribor

19

Principal components analysis (PCA) - example 2

(pairwise correlation among environmental variables)

	1	2	3	4	5	6	7	8	9
1 Air temperature	1.00								
2 arc-sin relative air humidity	0.15 0.133	1.00							
3 Ground temperature	0.94 <0.001	0.18 0.079	1.00						
4 arc-sin substrate moisture	0.388 <0.001	0.59 <0.001	0.37 <0.001	1.00					
5 Airflow	-0.48 <0.001	-0.36 <0.001	-0.43 <0.001	-0.55 <0.001	1.00				
6 Distance from entrance	-0.34 <0.001	0.14 0.153	-0.41 <0.001	0.10 0.312	0.04 0.712	1.00			
7 Distance from surface	-0.02 0.837	0.24 0.017	-0.04 0.683	0.46 <0.001	-0.11 0.275	0.67 <0.001	1.00		
8 Passage cross-section	0.35 <0.001	0.17 0.089	0.23 0.025	0.39 <0.001	-0.40 <0.001	-0.11 0.274	0.05 0.656	1.00	
9 log illumination	0.45 <0.001	-0.18 0.077	0.46 <0.001	-0.04 0.690	-0.07 0.494	-0.821 <0.001	-0.679 <0.001	0.37 <0.001	1.00

Table 1. Pearson correlations coefficient among re correlations in bold. (Upper row r, lower row p).

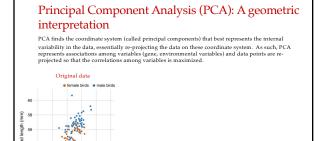
3

20

Principal components analysis (PCA) - example 2 (niche differences – dots represent different caves ellipsoids are confidence intervals for where species is found)

Each dot represents E. longicornis a cave PC 2 (29.5%)

PC 1 (37.6%) Fig. 5. Ordination of the nine environmental variables in $1^{\rm st}$ and $2^{\rm nd}$ PC axes. Ellipses (95% confidence) represent spatial niches in the three hymenopteran species.



Source https://wilkelab.org/SDS375/slides/dimension-reduction-1.html#9

22

Principal Component Analysis (PCA): A geometric interpretation PCA finds the coordinate system (called principal components) that best represents the internal variability in the data, essentially re-projecting the data on these coordinate system. As such, PCA represents associations among variables (gene, environmental variables) and data points are re-projected so that the correlations among variables in saximized. Original data Standardization and PCA fitting

23

Principal Component Analysis (PCA): A geometric interpretation PCA finds the coordinate system (called principal components) that best represents the internal variability in the data, essentially re-projecting the data on these coordinate system. As such, PCA represents associations among variables (gene, environmental variables) and data points are re-projected so that the correlations among variables is maximized. Original data Standardization and PCA fitting Rotation PCA aligns their axes with directions of maximum variation in the data Source https://willelabo.org/SOS375/slides/dimension-reduction-1.html#9

Principal Component Analysis (PCA): A geometric interpretation

- PCA constructs a new coordinate system (new variables, PCs) which are linear combinations of the original data and which are defined to align the samples along their major axes of variation (assuming linearity).
- Thus, PCA determines the coordinate system that best represents the internal variability in the data, essentially re-projecting the data.

25

The association among variables need to be measured by either (in most cases):

Correlation Matrix (for variables that have different units or scales, e.g., ph, temperature).

Covariance Matrix (variables have the same units, e.g., body length & body width in cm).

Raw data when variables are in the same units (more difficult to interpret) and calculations differ (very rare to find applications in the literature); rarely used.

26

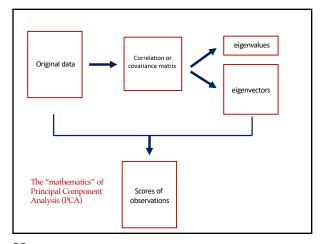
Correlation versus covariance

$$COV_{xy} = \frac{\sum_{i=1}^{n} (X_i - \bar{X})(Y_i - \bar{Y})}{n-1}$$

$$X = 0 \& Y = 0 : s_x = s_x \& s_y = s_y$$

$$COR_{xy} = \frac{COV_{xy}}{s_x s_y}$$

$$X = 0 \& Y = 0 : s_x = 1 \& s_y = 1$$



The mathematics of Principal Component Analysis (PCA):

Eigen-analysis is a mathematical operation on a *square symmetric* matrix (e.g., pairwise correlation matrix, pairwise covariance matrix).

A *square* matrix has the same number of rows as columns.

A *symmetric* matrix is the same if you switch rows and columns.

29

square and symmetric matrix

(e.g., pairwise correlation matrix)

	X ₁	X ₂	X ₃	X_4	X ₅
X ₁	1.00	0.80	0.90	0.78	0.87
X ₂	0.80	1.00	0.76	0.87	0.78
Хз	0.90	0.76	1.00	0.78	0.89
X4	0.78	0.87	0.78	1.00	0.95
X 5	0.87	0.78	0.89	0.95	1.00

The important components of Principal Component Analysis (pun intended)



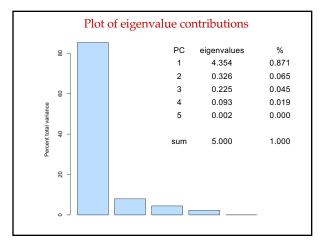
31

Principal component analysis presents three important structures:

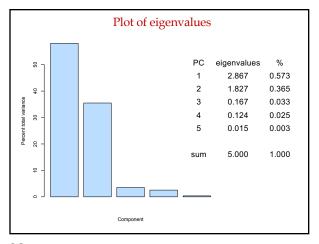
1 – **Eigenvalues:** represent the amount of variation in the original data summarized by each principal component. The first principal component (PC-1) presents the largest amount, PC-2 presents the second largest amount, and so on.

32

			Eigenv	zalue	es
\mathbf{X}_{1}	X_2	X_3	X_4	X_5	
1.00	0.80	0.90	0.78	0.87	
0.80	1.00	0.76	0.87	0.78	"one
5					
0.07	0.78	0.89	0.95	1.00	dimension"
РС	eigenvalu	ies	%	_	
PC		ies		7	
1	4.354		0.871		"Lower"
2	0.326		0.065		dimensionality
3	0.225		0.045		dimensionality
4	0.093		0.019		because it kept a
5	0.002		0.000		large proportion of
sum	5.000		1.000		the variation in the data in the first PC.
	1.00 0.80 0.90 0.78 0.87 Eigenva PC 1 2 3 4 5	1.00	X ₁ X ₂ X ₃ 1.00 0.80 0.90 0.80 1.00 0.76 1.00 0.78 0.87 0.78 0.87 0.78 0.87 0.78 0.87 0.78 0.87 0.78 0.87 0.78 0.87 0.78 0.89 0.87 0.25 0.326 3 0.225 4 0.093 5 0.002	X1	1.00



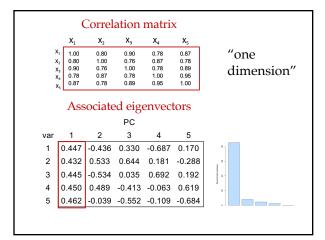
		Eig	genval	ues	
1.00		0.96	0.04	0.05	
0.87		0.95 1.00	0.03	0.07 0.05	"two
0.04		0.04	1.00	0.84	dimensions"
0.05	0.07	0.05	0.84	1.00	ullilelisions
2	2.867 1.827	0.57		Higher"	dimensionality
1	2.867	0.57		T: 1 //	
					,
3	0.167			wo components	
4	0.124	0.025 are ne		e neede	ed to summarize
_	0.015	0.003 variation.			
5					



Principal component analysis presents three important structures:

- 2 **Eigenvectors**: Each principal component is a linear function with coefficients for each variable.
- Eigenvectors contain these coefficients. High values, positive or negative, represents high association with the component.

37



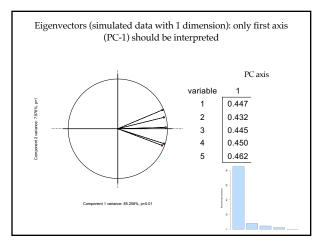
38

Eigenvectors can be seen as regression coefficients, where the component is the dependent variable. A "one dimension" matrix has only one interpretable principal component.

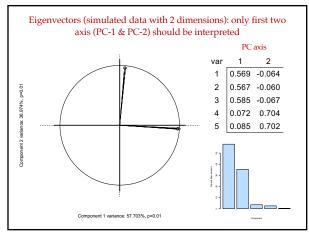
$PC\text{-}1\text{=}0.447X_1\text{+}0.432X_2\text{+}0.445X_3\text{+}0.450X_4\text{+}0.462X_5$

Unlike the numbers after =, this is not a subtraction but a hyphen stating that this is the first and second Principal Components (PC).

			PC		
var	1	2	3	4	5
1	0.447	0.436	0.330	-0.687	0.170
2	0.432	-0.533	0.644	0.181	-0.288
		0.534			
4	0.450	-0.489	0.413	-0.063	0.619
5	0.462	0.039	-0.552	-0.109	-0.684



	X ₁	X_2	X_3	X ₄	X ₅	
	1.00	0.87	0.96	0.04	0.05	"two
	0.87	1.00	0.95	0.03	0.07	1:
	√₃ 0.96	0.95	1.00	0.04	0.05	dimensions
	< ₄ 0.04 _{ζ5} 0.05	0.03	0.04 0.05	1.00 0.84	0.84 1.00	
			PC			
		_	_		_	
var	1	2	3	4	5	
var 1	1 0.569			-0.642	5 0.445	
				-0.642		:]
1	0.569	-0.064 -0.060	0.249	-0.642	0.445	* * * * * * * * * * * * * * * * * * *
1	0.569 0.567	-0.064 -0.060	0.249	-0.642 0.661 -0.810	0.445	months and a second a second and a second and a second and a second and a second an



Principal component analysis presents three important structures:

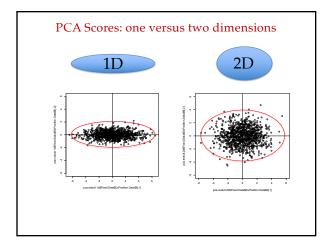
3 – Multivariate scores: Since each component is a linear function of the variables, when multiplying the standardized variables (in the case of correlation matrices) by the eigenvector structure, a matrix containing the position of each observation in each principal component is produced.

The plot of these scores in the first few dimensions, represents the main patterns of variation among the original observations (more in the empirical example).

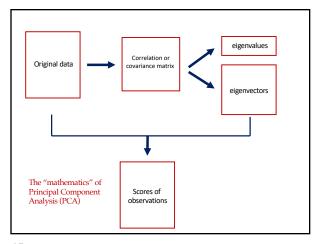
 $\begin{aligned} & \text{PC-1=0.569X}_1 + 0.567X_2 + 0.585X_3 + 0.072X_4 + 0.085X_5 \\ & \text{PC-2=-0.064X}_1 - 0.060X_2 - 0.067X_3 + 0.704X_4 + 0.702X_5 \end{aligned}$



43



44



Next lecture: How many PCA dimensions? Inferential frameworks for determining number of axes to interpret and the significance of each variable on each axis (lots of work on this area).

1st) determine how many axes to interpret (i.e., how many PCs capture correlated variation in the data?).



ecience @ direct.



How many principal components? stopping rules for determining the number of non-trivial axes revisited

Pedro R. Peres-Neto*, Donald A. Jackson, Keith M. Somers

46

Inferential frameworks for determining number of axes to interpret and the significance of each variable on each axis are usually nor performed.

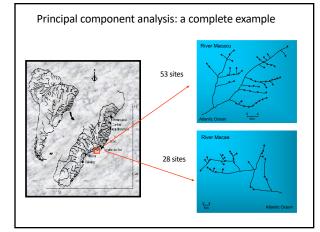
2nd) for each significant axis, determine which variable is significant on each of them.

Ecology, 84(9), 2003, pp. 2347-2363 © 2003 by the Ecological Society of America

> GIVING MEANINGFUL INTERPRETATION TO ORDINATION AXES: ASSESSING LOADING SIGNIFICANCE IN PRINCIPAL COMPONENT ANALYSIS

> > Pedro R. Peres-Neto, $^{\rm l}$ Donald A. Jackson, and Keith M. Somers

47



What is the correlation structure and differences among streams in terms of their environmental features?

Depth

Depth variation

Current velocity

Current variation

Substrate composition: Boulder, rubble, gravel and sand Substrate variation (variance in composition) Stream width variation (irregularity)

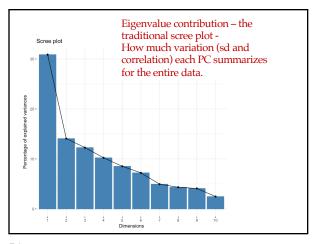
Area

Altitude

49

Correlation matrix 1.00 0.04 0.28 -0.07 0.06 -0.33 -0.02 0.12 -0.02 0.05 0.01 -0.11 0.04 1.00 0.12 0.85 0.31 0.07 0.07 -0.35 -0.42 0.84 0.86 -0.66 0.28 -0.12 1.00 -0.07 -0.17 -0.08 0.02 0.19 -0.13 -0.13 -0.17 -0.03 -0.07 0.85 -0.07 1.00 0.36 0.17 0.06 -0.44 -0.33 0.71 0.71 -0.57 0.06 0.31 -0.17 0.36 1.00 0.08 -0.33 -0.81 0.33 0.36 0.20 -0.17 -0.33 0.07 -0.08 0.17 0.08 1.00 0.11 -0.52 0.09 0.00 0.11 0.23 -0.02 0.07 0.02 0.06 -0.33 -0.11 1.00 0.04 -0.22 0.14 0.05 0.00 0.12 -0.35 0.19 -0.44 -0.81 -0.52 -0.04 1.00 -0.25 -0.39 -0.25 0.04 -0.02 -0.42 -0.13 -0.33 0.33 0.09 -0.22 -0.25 1.00 -0.38 -0.38 0.33 0.05 0.84 -0.13 0.71 0.36 0.00 0.14 -0.39 -0.38 1.00 0.66 -0.58 0.01 0.86 -0.17 0.71 0.20 0.11 0.05 -0.25 -0.38 0.66 1.00 -0.60 -0.11 -0.66 -0.03 -0.57 -0.17 0.23 0.00 0.04 0.33 -0.58 -0.60 1.00

50

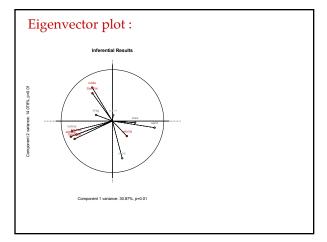


Eigenvector structure (2 first dimensions)

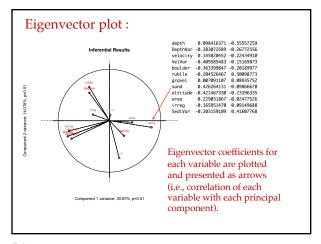
PC-1 PC-2

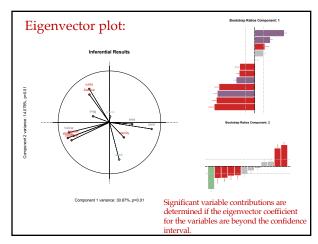
depth	0.098416371	-0.55557259
DepthVar	-0.383072589	-0.26772556
velocity	0.145820452	-0.22434910
VelVar	-0.409585483	-0.15169873
boulder	-0.363399847	-0.20189977
rublle	-0.204526467	0.50098773
gravel	0.007091107	0.08935752
sand	0.426264131	-0.09866678
altitude	-0.421467330	-0.23396335
area	0.229031867	-0.02477526
irreg	-0.165951470	0.09149688
SediVar	-0.203159109	0.41607768

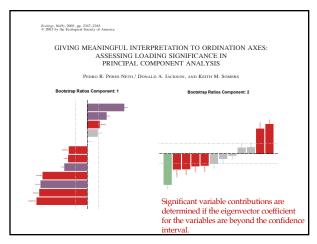
52

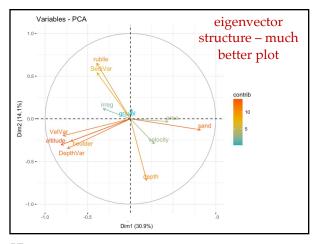


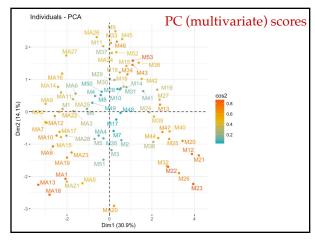
53

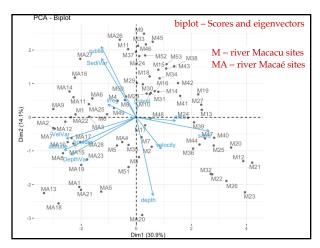


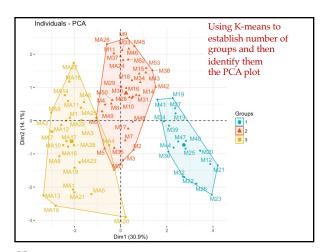












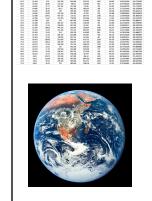
Mapping the environment of our planet - a very "small" example



61

Long 7-70.5 69.5 69.5 69.5 69.5 69.5 69.5 7-70.5 69.5 7-71.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70.5 69.5 7-70 avg_pre 89.49 50.23 37.9 47.71 36.37 29.06 28.69 149.71 108.16 33.43 31.43 13.61 13.63 13. way, VI 421,958 442,354 452,354 452,354 452,354 452,354 452,354 452,354 452,354 571,81 778,905 562,878 761,104 565,229 562,878 5761,504 567,405 562,878 5761,504 567,405 562,878 5761,504 567,405 562,878 562,878 562, arage, Age
2160
2160
2170
1258
1047
763
763
763
342
224
1590
1190
355
263
2785
263
2785
2600
861
418
3405
1756
491
1756
491
1756
491
1756
491
1756
491
1756
493
1567
494
1590
1756
493
1567
494
1590
1756
493
1567
1447 55.344 55.344 55.344 55.344 55.345 55.345 55.345 55.345 54.046 54.046 52.788 52.788 52.788 51.564 51 14909 geographic cells (110Km by 110Km)

62



- 1) Latitude (Lat) and Longitude (Long) at the centre of geographic cell.
- centre or geographic cell.

 2) Average precipitation (last 40 years; avg_prec)

 3) Average actual evapotranspiration (avg_ET, a proxy of productivity)

 4) Average vegetation index (avg_VI)

- Mean altitude (avg_Alt)
 Maximum altitude minus minimum altitude (altitudinal range; range_Alt)
- 7) Average temperature (avg_temp)
 8) Seasonal temperature (annual range in
- temperature; seas_temp)
 9) Seasonal precipitation (annual range in precipitation; seas_prec)

