Introduction to SaTScan, seasonality and time series analysis

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GIS training course for animal health workers Jakarta, Indonesia 28 November 2016





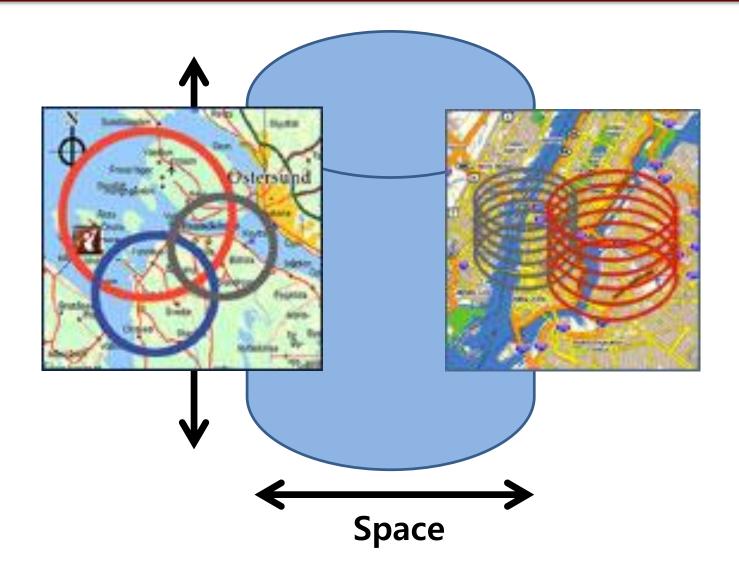


Outline

- Introduction of SaTScan
- Examples
 - Leptospirosis in the USA
 - FMD in Korea
- Introduction of seasonality analysis
- Example
 - Viral Encephalitis in Vietnam
- Introduction of time series analysis
- Example
 - Brucellosis in South Korea

- SaTScan is a freely available software that uses the scan statistic to detect clusters (www.satscan.org)
- To test whether a disease is randomly distributed over space, over time or over space and time.
- To perform geographical surveillance of disease, to detect areas of significantly high or low rates.

What is SaTscan?



For each distinct window, calculate the likelihood, proportional to:

$$\left(\frac{n}{\mu}\right)^n \left(\frac{N-n}{N-\mu}\right)^{N-n}$$

- *n* = number of cases inside circle
- N = total number of cases
- μ = expected number of cases inside circle

Scan Statistics

 Circles of different sizes (from zero up to 50 % of the population size)

 For each circle a likelihood ratio statistic is computed based on the number of observed and expected cases within and outside the circle and compared with the likelihood *LO* under the null hypothesis.

- The **scan statistic** is the maximum likelihood over all possible circles
 - Identifies the most unusual clusters
- To find p-value, use Monte Carlo hypothesis testing
 - Redistribute cases randomly and recalculate the scan statistic many times
 - Proportion of scan statistics from the Monte Carlo replicates which are greater than or equal to the scan statistic for the true cluster is the p-value

What SaTScan can/can't do?

• CAN

- Identify spatial, temporal, spatial-temporal clusters
- Provide flexible geographic units

• CANNOT

- Display maps of events and clusters locations
- Need GIS or mapping software (such as ArcGIS)
- Create other statistical and regression models

Introduction of Statistical models in SaTScan

Bernoulli Model

- There are animals with or without a disease (represented by a 0/1 variable)
 - A set of cases and controls

• Purely temporal/spatial or the space-time scan statistics

Discrete Poisson Model

- The number of cases in each location is Poissondistributed.
- Under the null hypothesis, and when there are no covariates, the expected number of cases in each area is proportional to its population size
- Purely temporal, purely spatial and space-time
- This model a very good approximation to the **Bernoulli model** if few cases VS controls (less than 10%)

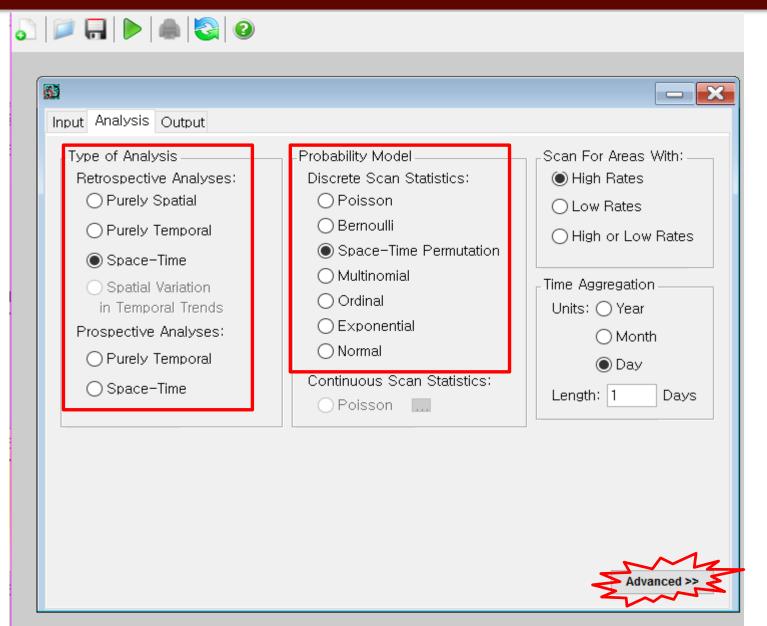
Space-Time Permutation Model

- Requires only cases data with information about the spatial location and time for each case (No information needed for population at risk)
- If the population increase (or decrease) is the same across the study region, that is okay, and will not lead to biased results
- The user is advised to be very careful when using this method for data spanning several years
 - population in some areas grows faster than in others

Screenshot of SaTScan

File	Atum	n_cases - 메모장		
	파일(F)	"편집(E) 서식(O) 보기(
	1001 1002 1007 1008 1020 1028 1033 1035 1035 1036 1039 1040 1057 1060 1069 1077 1085	42.062 -72.6255 42.3818 -72.4542 42.2788 -72.4028 42.1881 -72.9663 42.1759 -72.5676 42.0593 -72.4973 42.2579 -72.5066 42.3547 -72.5724 42.0669 -72.4127 42.4043 -72.6883 42.2133 -72.6408 42.0895 -72.3216 42.3161 -72.6276 42.1849 -72.3098 42.0508 -72.7769 42.1511 -72.7653	Input Analysis Output Case File: Control File: (Bernoulli Model) Study Period Study Period Start Date: 2000 1 1 End Date: 2010 12 3	Month O Day Generic
	1095 1096 1104	42.1297 -72.4284 42.4204 -72.776 42.1351 -72.5702	Population File: (Poisson Model)	
	1118 1119 1201 1230 1238 1247	42.0951 -72.5259 42.1239 -72.511 42.4531 -73.2856 42.1906 -73.406 42.2951 -73.2268 42.703 -73.0702	Grid File: (optional)	Coordinates Cordinates Cartesian Lat/Long
	1301 1351 1370	42.5383 -72.5154 42.5933 -72.727		Advanced >>

Input



Input – Advanced Analysis

(Advanced Analysis Features					
	Spatial Window Temporal Window Space and Time Adjustments Inference Border Analysis Power Evaluation					
Input Analysis Output Type of Analysis Retrospective Analyses: Purely Spatial Space-Time Spatial Variation in Temporal Trends Prospective Analyses: Purely Temporal Space-Time Space-Time	Maximum Spatial Cluster Size 50.0 percent of the population at risk (<= 50%, default = 50%) 					

Output

File Session Windows Help

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	8	— ×
	Input Analysis Output	
	Text Output Format Main Results File:	
	Geographical Output Format	
	Shapefile for GIS software	
	Column Output Format	
	Cluster Information Stratified Cluster Information Location Information Risk Estimates for Each Location Simulated Log Likelihood Ratios/Test Statistics	dBase
		Advanced >>

Executing

saiscan - software for the spatial and space-lime scan statistic

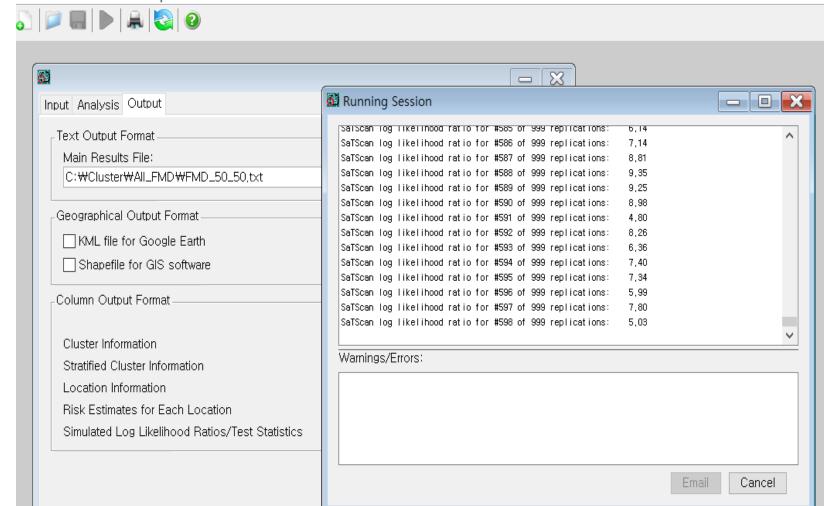
File Session Windows Help



Input Analysis Output
Case File: Time Precision C:\U00cmCluster\U00cmAll_FMD\U00cmCase_FMD.txt Control File: (Bernoulli Model)
Study Period Year Month Day Start Date: 2010 11 1 End Date: 2015 4 30
Population File: (Poisson Model) C:\Cluster\All_FMD\Pop_FMD.txt
Coordinates File:
C:\Uster\All_FMD\Geo_FMD.txt OCartesian
Grid File: (optional) Cat/Long
Advanced >>

Processing

File Session Windows Help

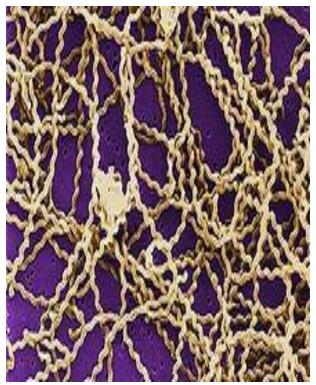


Advanced >>

Example: Lepto

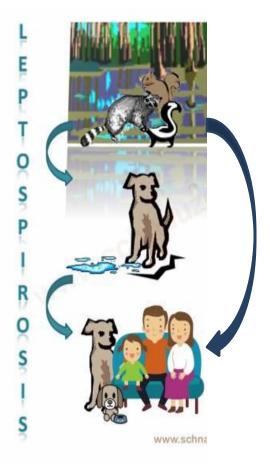
Leptospirosis

- A bacterial zoonotic disease caused by spirochetes of the genus *Leptospira*
- Pathogenic*:*
 - L. interrogans and L. kirschneri
 - More than 200 serovars



Epidemiology

- Zoonotic disease
 - Transmitted to humans from a variety of wild and domesticated animal hosts
 - Most common reservoirs: rodents (rats),
 wild animals and farm animals in the US
 - Occupations that involve animal handling are more likely to contract disease
- Transmitted through damaged skin or mucus membranes of exposed humans and animals
- Indirect contact (water, soil and feed) with infected urine from an animal with leptospiruria



Host animals

Species	Common infections	Possible others	
Dogs	Canicola, Icterohemorrhagiae, Gri ppotyphosa, Pomona	Bratislava, Autumnalis	
Cats	rarely identified		
Cattle (and deer)	Hardjobovis, Pomona, Grippotyph osa, Icterohemorrhagiae	Australis, Autumnalis, Canicola, Bataviae, Hebdomadis, Kremat osis, Tarassovi, Sejroe, Bratislav a	
Pigs	Pomona, Bratislava, Canicola, Tar assovi, Icterohemorrhagiae	Grippotyphosa, Sejroe	
Sheep	Pomona, Grippotyphosa, Bratislav a, Hardjo		
Horses	Pomona, Bratislava, Canicola, Icter ohemorrhagiae, Sejroe		

Reference: Bharti, A.R., Nally, J.E., Ricaldi, J.N., Matthias, M.A., Diaz, M.M., Lovett, M.A., Levett, P.N., Gilman, R.H., Willig, M.R., Gotuzzo, E., 2003. Leptospirosis: a zoonotic disease of global importance. Lance Infect. Dis. 3, 757-771.

Objective of study

• To evaluate spatial clustering by each serovar in the United States between 2008 and 2010 using data from IDEXX Laboratories Inc.

Materials and methods

- Procedures
 - Spatial clustering analysis (2008-2010)
 - Overall (any titers ≥ 1600) / each serovar
 - Spatial window was set at maximum 20%
 - *P*-value < 0.05
 - Geocoding was conducted from hospital zipcodes into geographical coordinates (latitude and longitude)
 - Sample locations with clusters were visualized on the map

Materials and methods

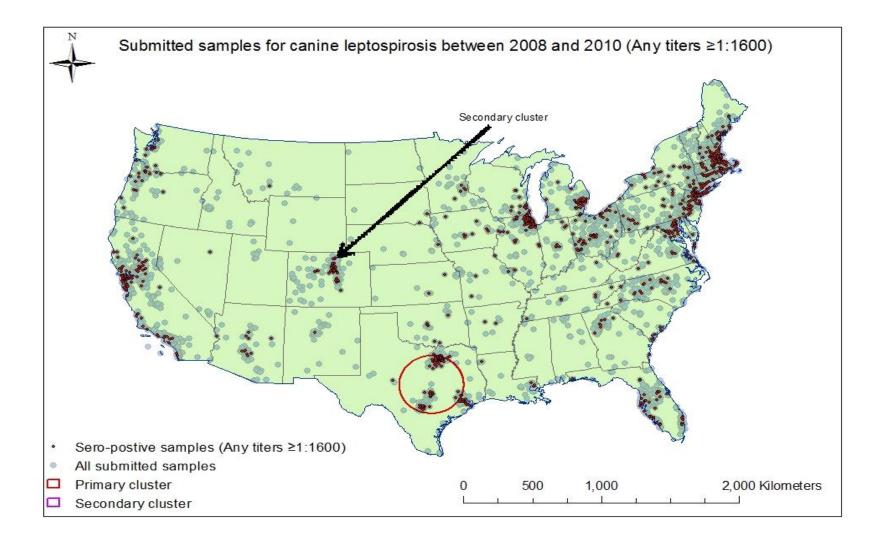
- Which model is appropriate?
 - Discrete Poisson Model
 - May consider... Bernoulli Model
- Software programs
 - Microsoft Excel & Note
 - STATA version
 - ArcGIS (ESRI, CA, USA)
 - SaTScan

Results from Poisson and Bernoulli Models

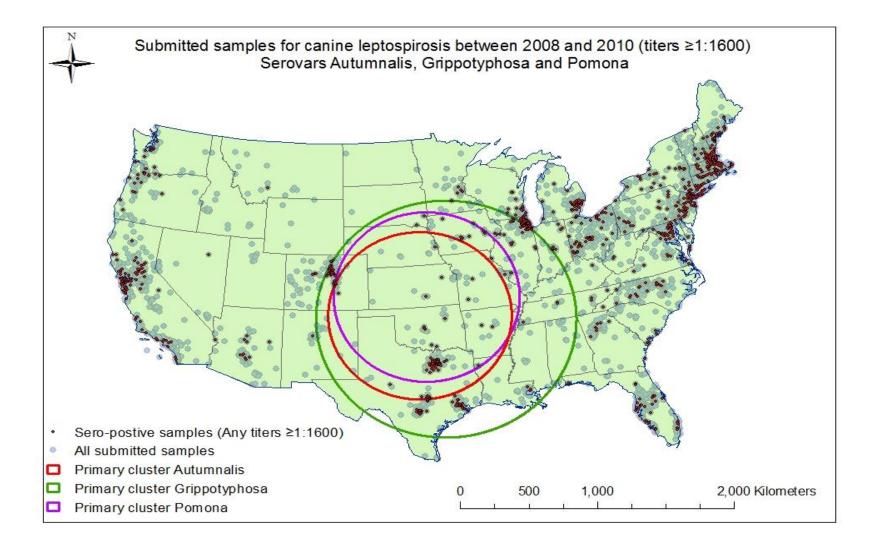
- 50% of the population at risk area (serovar Autumnalis)
- Only primary cluster is significant in both models

	77043, 45828, 75071, 75092, 43207, 75042, 75013, 75070, 74070, 771 75228, 77035, 43204, 74006, 77042, 77053, 75081, 43026, 75218, 321 75238, 77079, 34293, 43206, 27560, 75023, 43318, 75009, 26150, 751 27513, 43212, 75025, 75243, 75080, 77077, 27501, 43064, 77084, 431 77072, 43221, 43232, 77433, 75214, 75035, 75231, 75223, 43227, 341 45822, 77082, 77477, 61614, 73446	77043, 45828, 75071, 75092, 43207, 75042, 75013, 750 75228, 77035, 43204, 74006, 77042, 77053, 75081, 430 75238, 77079, 34293, 43206, 27560, 75023, 43318, 750 27513, 43212, 75025, 75243, 75080, 77077, 27501, 430 77072, 43221, 43232, 77433, 75214, 75035, 75231, 752 45822, 77082, 77477, 61614, 73446
Coordinates / radius Gini Cluster Population Number of cases Expected cases Annual cases / 100000. Observed / expected Relative risk Log likelihood ratio	(33.227300 N, 87.578900 W) / 856.51 km Yes 519 107 54.61 20573.9 1.96 2.08 2	e with clusters. No Overlap mates / radius (33.227300 N, 87.578900 W) / 856.51 km uster Yes ion
	1742, 1827, 1887, 1803, 1450, 1810, 1719, 1432, 2420, 1844, 2421, 1843, 3060, 3051	n IDs included.: 59068, 59044, 82414, 59102, 59101, 59718, 59714, 597 59754, 59601, 59405, 59725, 82520, 82604, 83201, 832 82636, 83204, 59802, 59840, 82901, 59875, 59808, 833
Poisso	on Model	Bernoulli Model Model

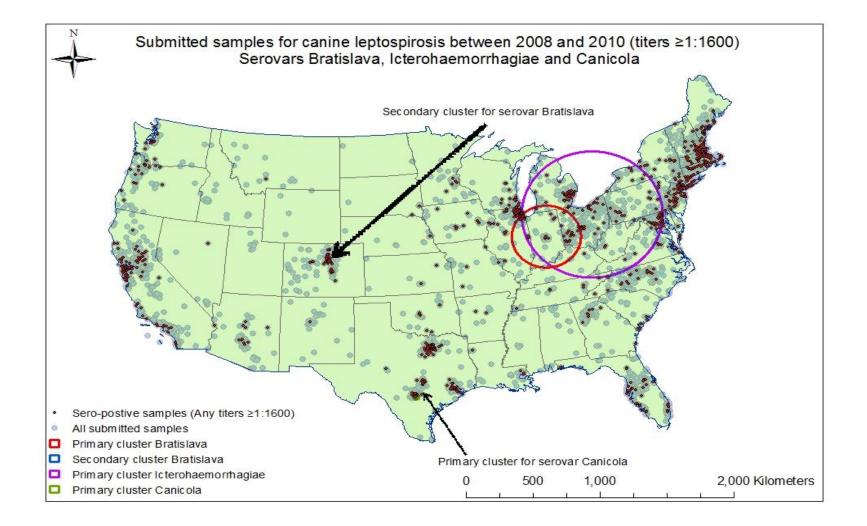
Cluster map from Poisson Model



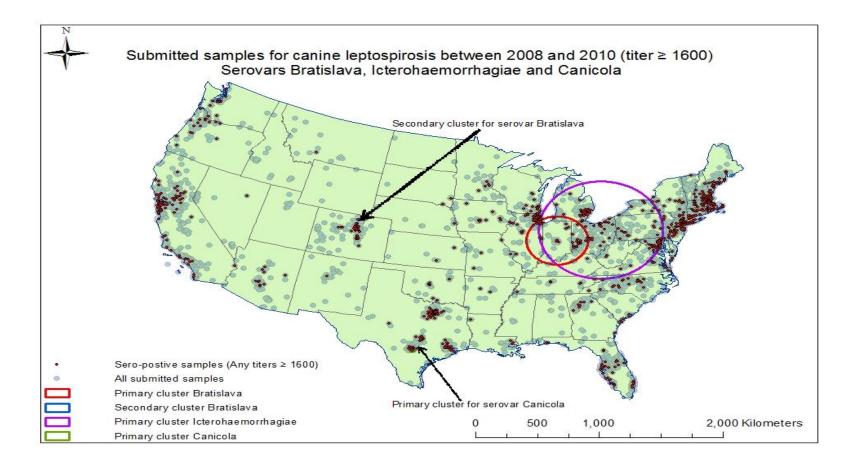
Cluster map from Poisson Model



Discrete Poisson Model



Cluster map from Poisson Model



Spatial clusters of *Leptospira* seropositivity (MAT titers ≥1:1,600) between 2008 and 2010 in dogs in the United States (spatial window: max 20%)

Cluster	Commission	D1-6'	Dedline (less)	Ob a larger an anti-	D 1
	Geographical area	Population	Radius (km)	Obs/exp = ratio	P-value
Any titers ≥ 1600					
N=18,717; cases:1,487	G (177	2 1 2 0	222.07		-0.001
Primary cluster	Central Texas	2,139	232.87	262 / 162.02 = 1.62	< 0.001
Secondary cluster	North-central Colorado	83	7.94	22 / 6.29 = 3.50	0.011
Autumnalis					
N=18,632; cases:949					
Primary cluster	Central Oklahoma	2,164	667.02	156 / 102.99 = 1.51	0.011
		-,			
Grippotyphosa					
N=18,595; cases:795					
Primary cluster	Central Oklahoma	5,970	941.48	358 / 234.14 = 1.53	< 0.001
Pomona					
N=18,646; cases:494					
Primary cluster	Central Kansas	2,358	675.28	115/59.58=1.93	< 0.001
		-,			
Bratislava					
N=18,678; cases:425					
Primary cluster	Central Indiana	919	249.73	44 / 18.69 = 2.35	0.006
Secondary cluster	North-central Colorado	65	4.86	10/1.32 = 7.56	0.020
Secondary cluster	Norm-central Colorado	05	4.00	1071.52 - 7.50	0.020
Icterohaemorrhagiae					
N=18,699; cases:172					
Primary cluster	North-central Ohio	5,111	501.41	72 / 38.75 = 1.86	0.001
Filliary cluster	Norm-central Onio	5,111	501.41	72758.75 - 1.80	0.001
Canicola					
N=18,698; cases:118					
	South-central Texas	116	22.42	12/0.57 = 21.21	<0.001
Primary cluster	South-central Texas	110	22.42	12/0.57 - 21.21	~0.001

Example: FMD

FMD

 Foot-and-mouth disease is a highly contagious, viral disease of domestic cloven-hoofed and many wild animals

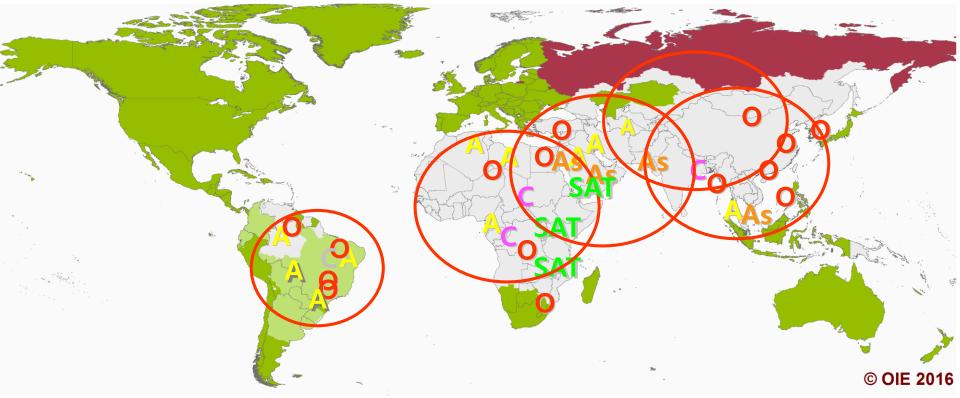


Worldwide occurrence of FMD

OIE Member Countries' official FMD status map

Last update October 2016

Click on a specific region to zoom in



Member Countries and zones recognised as free from FMD without vaccination

Member Countries and zones recognised as free from FMD with vaccination

Countries and zo

Countries and zones without an OIE official status for FMD

Suspension of FMD free status without vaccination

 Data (2010-2015): South Korea http://www.kahis.go.kr/home/lkntscrinfo/selec

tLkntsOccrrncList.do?openFlag=Y

• Data (2010-2015): North Korea

http://www.oie.int/wahis_2/public/wahid.php/Wahid home/Home

- Number of infected/susceptible animal at farm levels (with geographical information)
- Outbreak date

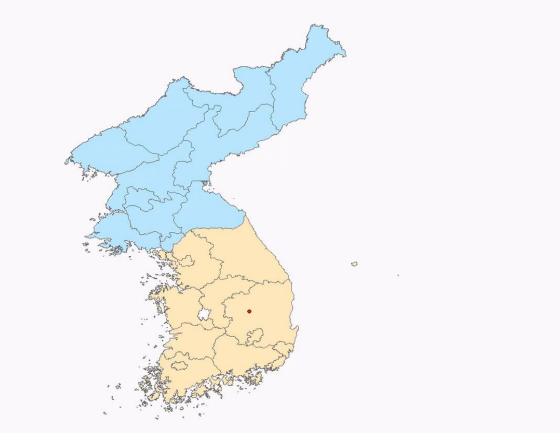
Discrete Poisson Model

				G	eo_FMD	- Notepad
File	Edit	Format	View	Help		
Loc	atio	n	Lat	itude		Longitude
1		36.401	54		128.4522	2166
2		36.401	681		128.4530	014
3		36.365			128.415	
4		36.394			128.4717	
5		36.392			128.4749	
6		36.354			128.423	
7		36.381			128.4343	
8		36.382			128.4304	
9		36.375			128.4221	
10		36.381			128.430	
11		36.383			128.4213	
12		36.345			128.4122	
13		36.381			128.4351	
14 15		36.373			128.424	
16		36.361			128.4120	
17		36.382			128.4352	
18		36.371			128.4411	
19		36.383			128.4234	
20		36.383			128.431	
21		36.382			128.4720	
22		36.343			128.3910	
23		36.381			128.4548	
24		36.404			128,423	
25		36.375	689		128.4530	509
26		36.398	099		128.4159	968
27		36.401	094		128.4219	913
28		36.315	029		128.474	543
29		36.351	712		128.2890	034
30		36.381	163		128.5552	229
31		36.371	847		128.3940	019
32		36.404	778		129.0329	
33		36.470			128.3748	
34		36.322			128.2308	392
35		37.823		.0082		
36		38.026				
37		37.828				
38		39.883				
39		37.713	3 126	.6985		
-						

Results

1 Location IDS incloses: Space(50%)-time(50%) 379, 50, 395, 398, 157, 351, 357, 139, 318, 53, 98, 339, 366, 407 348, 353, 324, 383, 183, 375, 72, 54, 414, 62, 346, 454, 370, 341 367, 307, 340, 337, 347, 342, 352, 349, 363, 362, 328, 336, 369, 64, 338, 331, 368, 425, 359, 314, 179, 365, 106, 335, 312, 424, 3 423, 435, 57, 409, 329, 356, 309, 358, 125, 350, 301, 303, 305, 3 300, 302, 401, 317, 396, 164, 315, 299, 298, 313, 297, 61, 147, 4 135, 316, 310, 132, 144, 133, 321, 344, 345, 332, 408, 381, 142, 126, 404, 394, 420, 410, 419, 161, 155, 306, 467, 308, 311, 361, 219, 322, 163, 140, 433, 478, 304, 261, 173, 320, 63, 83, 33, 323 280, 374, 417, 360, 124, 472, 119, 327, 371, 42, 250, 481, 468, 2 27, 26, 439, 1, 2, 440, 247, 11, 19, 20, 8, 4, 10, 29, 31, 7, 17, 5, 245, 9, 23, 14, 21, 25, 3, 185, 16, 445, 34, 6, 22, 47, 441 12, 30, 121, 91, 376, 102, 334, 447, 28, 441, 399, 46, 196, 453, 411, 172, 460, 249, 466, 479, 99, 465, 442, 35, 459, 448, 109, 28 127, 473, 474, 156, 100, 326, 248, 343, 476, 145, 43, 409, 457, 3 44, 469, 246, 462, 378, 416, 470, 354, 377, 60, 52, 279, 97, 95, 422 104, 137, 96, 276, 458, 84, 283, 45, 265, 456, 138, 438, 446, 319 451, 480, 437, 443, 450, 426, 429, 235, 475, 452, 55, 403, 436, 4 427, 413, 389, 415, 406, 382, 418, 397, 434, 384, 391, 428, 393, 4 402, 388, 380, 385, 390, 392, 464, 387, 386, 373, 471, 444, 234, 110, 288, 92, 431, 287, 289, 82, 229, 208, 432 Coordinates / radius: (37.224964 N, 128.081455 E) / 167.39 km Time frame: 2014/12/1 to 2015/4/30 Population: 2172 Number of cases: 155767 Expected cases: 155767	<pre>300, 302, 401, 317, 396, 164, 315, 299, 298, 313, 297, 61, 147, 49, 135, 316, 310, 132, 144, 133, 321, 344, 345, 332, 408, 381, 142, 355, 126, 404, 394, 420, 410, 419, 161, 155, 306, 467, 308, 311, 361, 325, 219, 322, 163, 140, 433, 478, 304, 261, 173, 320, 63, 83, 33, 323, 280, 374, 417, 360, 124, 472, 119, 327, 371, 42, 250, 481, 468, 24, 27, 26, 439, 1, 2, 440, 247, 11, 19, 20, 8, 4, 10, 29, 31, 7, 17, 13, 5, 245, 9, 23, 14, 21, 25, 3, 18, 15, 16, 445, 34, 6, 22, 47, 421, 12, 30, 121, 91, 376, 102, 334, 447, 28, 441, 399, 46, 196, 453, 412, 411, 172, 460, 249, 466, 479, 99, 465, 442, 35, 459, 448, 109, 281, 127, 473, 474, 156, 100, 326, 248, 343, 476, 145, 43, 449, 457, 32, 44, 469, 246, 462, 378, 416, 470, 354, 377, 182, 37, 477, 40, 430, 59, 56, 274, 36, 120, 143, 39, 180, 277, 60, 52, 279, 97, 95, 422, 104, 137, 96, 276, 458, 84, 283, 45, 265, 456, 138, 438, 446, 319, 451, 480, 437, 443, 450, 426, 429, 235, 475, 452, 55, 403, 436, 455, 427, 413, 389, 415, 406, 382, 418, 397, 434, 384, 391, 428, 393, 461, 402, 388, 380, 385, 390, 392, 464, 387, 386, 373, 471, 444, 234, 101, 110, 288, 92, 431, 287, 289, 82, 229, 208, 432</pre>
Expected cases: 6523.33 Annual cases / 100000.: 8923072.9 Observed / expected: 23.88	Annual cases / 100000.: 8923072.9 Observed / expected: 23.88
Relative risk: 254.51 Log likelihood ratio.:: 457679.909303	Relative risk: 254.51 Log likelihood ratio: 457679.909303
P-value: < 0.00000000000000000	P volue
2.Location IDs included.: 165, 129, 225, 105, 223, 232, 166, 206, 222, 184, 226, 254, 195, 273, 257, 130, 216, 220, 210, 136, 186, 215, 189, 188, 268, 153, 150, 152, 259, 160, 159, 117, 148, 77, 256, 149, 213, 192, 123, 1 294, 193, 66, 94, 212, 86	294, 193, 66, 94, 212, 86
Coordinates / radius.: (38.733300 N, 126.120000 E) / 34.16 km Time frame 2010/12/1 to 2011/1/31 Population 20 Number of cases: 6172 Expected cases: 73.77 Annual cases / 100000.: 31265973.8 Observed / expected: 83.67 Relative risk: 86.76 Log likelihood ratio: 21334.342555 P-value: < 0.000000000000001	Coordinates / Padlus.: (36.735300 N, 120.120000 E) / 34.10 Km Time frame 2010/12/1 to 2011/1/31 Population 20 Number of cases 6172 Expected cases 73.77 Annual cases / 100000.: 31265973.8 Observed / expected 83.67 Relative risk

ArcGIS - import (outbreak locations)



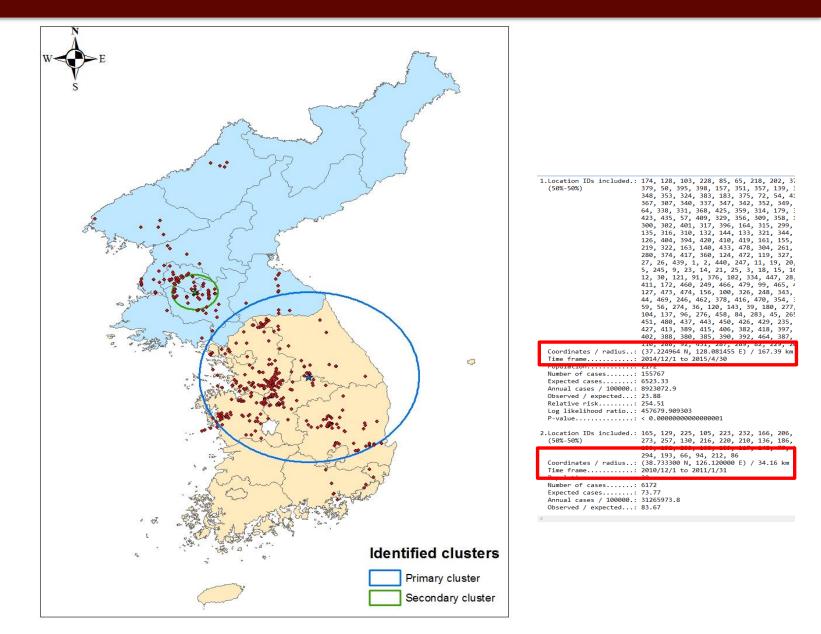
Outbreak locations (2010 Nov-2015 April)



ArcGIS – buffer (50%-50%)

e Edit View Bookmarks	Insert Selection Geoprocessing Customize	e Windows Help				
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 XVPrim_cluster_50_50 ✓ prim_50_50 C:\Users\del\Desktop\Cl Prim_cluster_50_50.bt Outbreak_location.tbt Prim_cluster_50_50.bt C:\Users\del\Desktop\M ✓ Korea polbnda2013 	(50%-50%) 379, 348, 347, 348, 347, 348, 348, 348, 347, 348, 348, 348, 348, 348, 348, 348, 349, 129, 280, 27, 280, 27, 280, 27, 280, 27, 280, 280, 27, 280, 290, 294, 273, 294, 2	33 372.9 36 31 99.909303 9000000000000001 129, 225, 105, 223, 232, 166, 206, 257, 130, 216, 220, 210, 136, 186, 152, 259, 160, 159, 117, 148, 77, ; 193, 66, 94, 212, 86 133300 N, 126.120000 E) / 34.16 km 12/1 to 2011/1/31		Input Features [XYSecond_cluster_50_50 Output Feature Class [C:\Jsers\dell'Desktop\Cluster\Kor Distance [value or field] ① Linear unit [] ① Field [] Side Type (optional) FULL End Type (optional) PLANAR Dissolve Type (optional) NONE []]	34.16 Kilometers	×

Map with identified clusters



Time clusters (2010-2015)

18000	Number of cases
16000	Number of cases
14000	
12000	
10000	
8000	
6000 4000 2000 0	
261212010 2610312012 2610512012 2610112012 26101120	$\frac{1}{26012603} \frac{1}{2603} \frac{1}{$
Coordinates / radias Time frame	. (37.224964 N, 120.00145 5 E) / 167.39 km : 2014/12/1 to 2015/4/30 - 2172
Number of cases Expected cases Annual cases / 100000. Observed / expected Relative risk Log likelihood ratio P-value	: 6523.33 : 8923072.9 : 23.88 : 254.51
Coordinates / radius	<pre>: 165, 129, 225, 105, 223, 232, 166, 206, 222, 184, 226, 254, 195, 122, 273, 257, 130, 216, 220, 210, 136, 186, 215, 189, 188, 268, 153, 253, 150, 152, 259, 160, 159, 117, 148, 77, 256, 149, 213, 192, 123, 197, 294, 193, 66, 94, 212, 86 . (30.755500 N, 120.12000 E) / 34.16 km : 2010/12/1 to 2011/1/31</pre>
Number of cases Expected cases Annual cases / 100000. Observed / expected Relative risk Log likelihood ratio P-value	: 73.77 : 31265973.8 : 83.67 : 86.76

Exercises (50 mins)

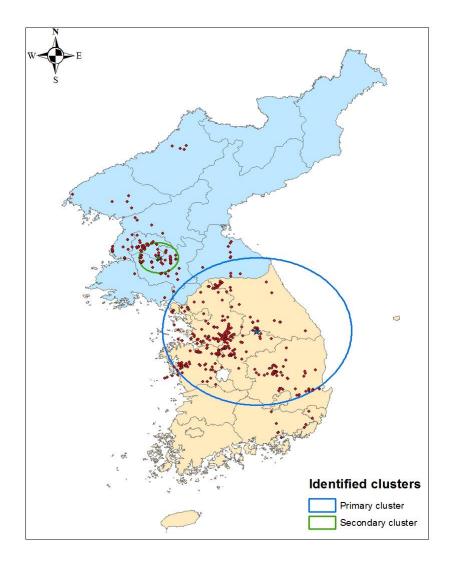
Leptospirosis data

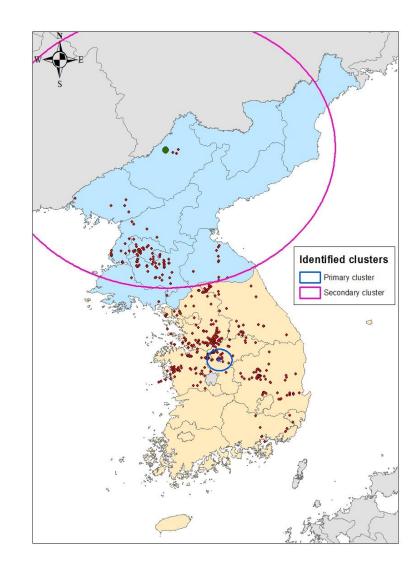
- Which model? (Bernoulli vs Poisson)
- Create files (I already created for you)
- Import data and Output
- Determine: cluster size (from zero to 50%)
- Comparison among different size (10%, 30% and 50%)

FMD data

- Which model? Space or time ?
 (Poisson vs Space-time permutation)
- Create files (You need to create files your own) Notepad
- Import data and Output
- Determine: cluster size (from zero to 50%)
- Comparison among different size (50%-50%, 50%-10%, 10%-50% and 10%-10%)

Poisson vs Permutation models (50%-50%)





Seasonality analysis

Introduction

• Procedures (seasonality using R)

J Vet Intern Med 2014

Regional and Temporal Variations of *Leptospira* Seropositivity in Dogs in the United States, 2000–2010

H.S. Lee, M. Levine, C. Guptill-Yoran, A.J. Johnson, P. von Kamecke, and G.E. Moore

Background: Previous studies have reported a seasonal increased risk for leptospirosis, but there is no consistent seasonality reported across regions in the United States.

Objectives: To evaluate and compare seasonal patterns in seropositivity for leptospirosis in dogs for 4 US regions (northeast [NE], midwest [MW], south-central [SC], and California-southern west coast [CS]).

Animals: Forty four thousand nine hundred and sixteen canine serum samples submitted to a commercial laboratory for microscopic agglutination tests (MAT) from 2000 through 2010.

Methods: In this retrospective study, positive cases were defined as MAT titers ≥ 1 : 3,200 for at least one of 7 tested serovars. Four geographic regions were defined, and MAT results were included in regional analyses based on hospital zip-

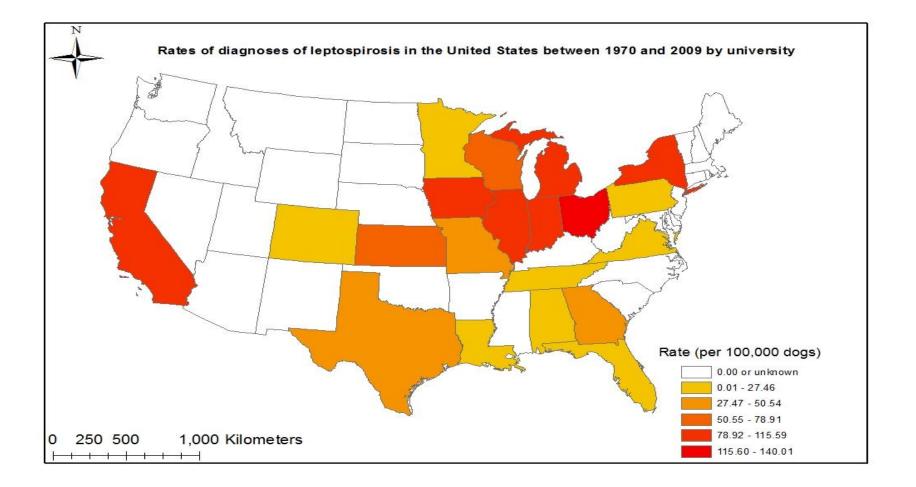
code. A seasonal-trend decomposition method for times series was utilized for the analysis. Monthly variation in the seropositive rate was evaluated using a seasonal cycle subseries plot and logistic regression.

Results: Two thousand and twelve of 44,916 (4.48%) samples were seropositive. Compared to seropositive rates for February, significantly higher monthly rates occurred during the 2nd half of the year in the MW (OR 3.92–6.35) and NE (OR 2.03–4.80) regions, and only in January (OR 2.34) and December (OR 1.74) in the SC region. Monthly seropositive rates indicative of seasonality were observed earlier in the calendar year for both CS and SC regions.

Conclusions and Clinical Importance: Seasonal patterns for seropositivity to leptospires differed by geographic region. Although risk of infection in dogs can occur year round, knowledge of seasonal trends can assist veterinarians in formulating differential diagnoses and evaluation of exposure risk.

Key words: Dogs; *Leptospira*; Microscopic agglutination tests; Seasonal cycle subseries plot; Seasonal-trend decomposition procedure based on loess; Seropositive.

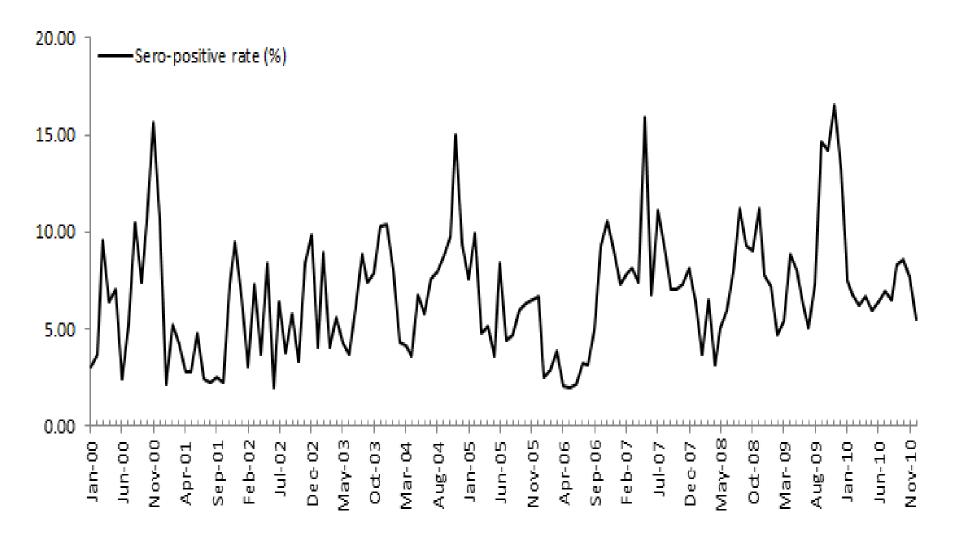
Risk map



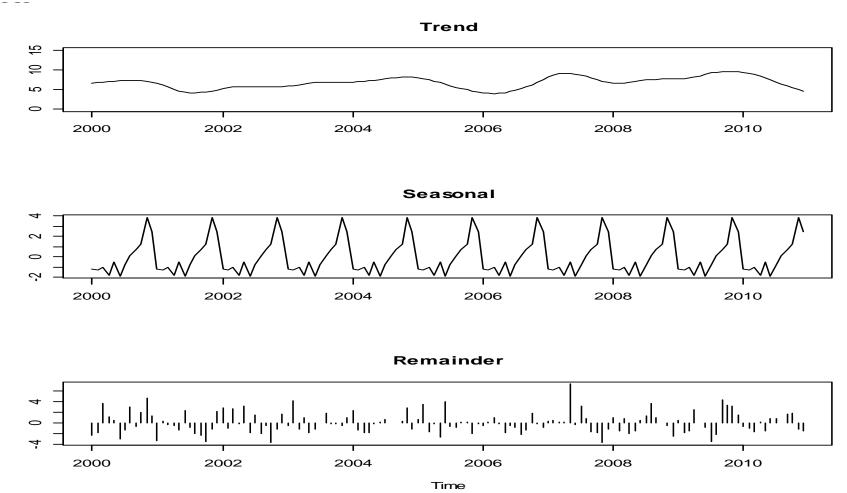
What is STL and SCS ?

- STL is a useful tool to visualize time series datasets that consists of
 - Trend, seasonal and remainder components
- SCS helps to visualize patterns both between and with groups that consists of
 Horizontal lines: average for each month
 - Vertical lines: individual pattern for the same month in each year

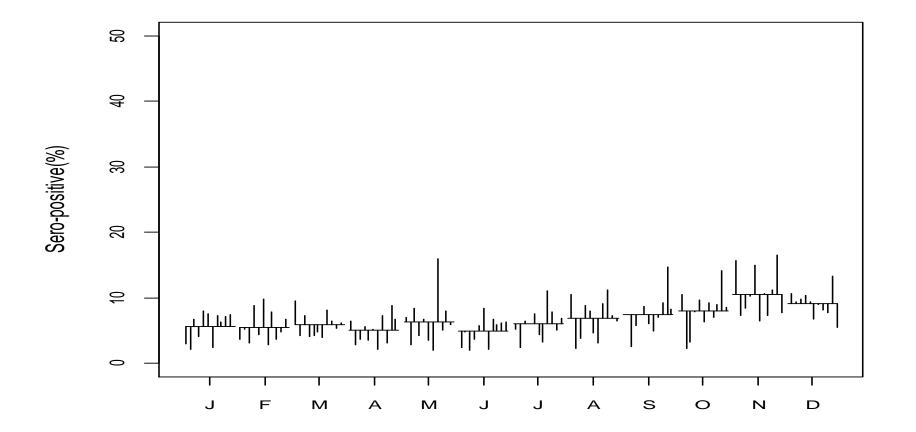
Time series plot



STL plot



Seasonal cycle subseries plot

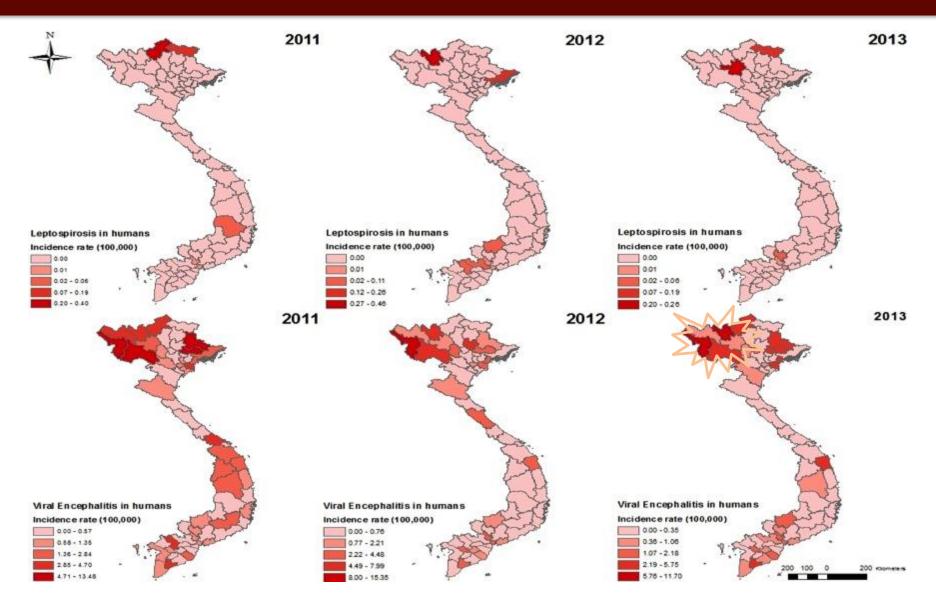


Regression model

Odds ratios with 95% Confidence interval

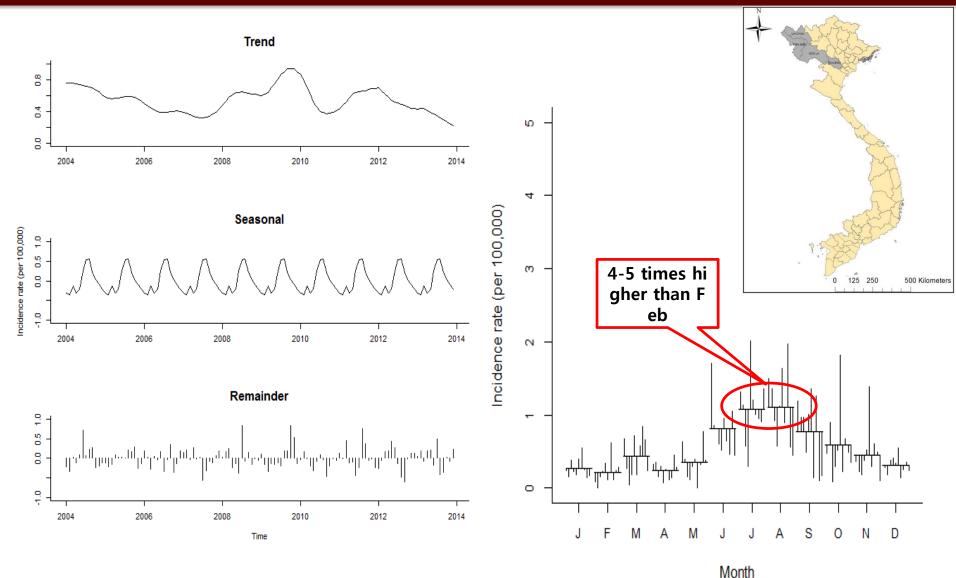
Month	Northeast	Mid-west	South central	California
Jan	1.30 (0.86-1.98)	1.18 (0.60-2.33)	1.32 (0.76-2.27)	Reference: 1
Feb	Reference: 1	Reference: 1	Reference: 1	1.93 (1.17-3.18)
Mar	0.71 (0.44-1.15)	1.26 (0.65-2.46)	1.08 (0.62-1.88)	1.98 (1.22-3.23)
Apr	0.91 (0.59-1.41)	0.87 (0.42-1.79)	1.28 (0.73-2.23)	1.60 (0.96-2.66)
Мау	1.06 (0.69-1.62)	1.70 (0.90-3.20)	1.45 (0.85-2.45)	1.49 (0.87-2.53)
Jun	0.90 (0.58-1.39)	1.16 (0.60-2.24)	1.08 (0.62-1.88)	1.22 (0.71-2.11)
Jul	1.23 (0.82-1.86)	1.42 (0.74-2.70)	1.35 (0.79-2.30)	0.95 (0.53-1.70)
Aug	1.61 (1.09-2.38)	1.59 (0.84-3.01)	1.07 (0.61-1.85)	1.02 (0.58-1.80)
Sep	2.04 (1.40-2.98)	1.59 (0.85-2.96)	1.30 (0.77-2.20)	0.76 (0.41-1.42)
Oct	2.29 (1.59-3.29)	1.72 (0.94-3.17)	1.30 (0.77-2.19)	0.62 (0.33-1.18)
Nov	3.40 (2.38-4.88)	2.31 (1.27-4.19)	1.47 (0.88-2.46)	1.60 (0.96-2.68)
Dec	2.39 (1.65-3.46)	2.00 (1.09-3.69)	1.74 (1.04-2.90)	1.31 (0.77-2.24)

Annual incidence rates for lepto and VE in humans



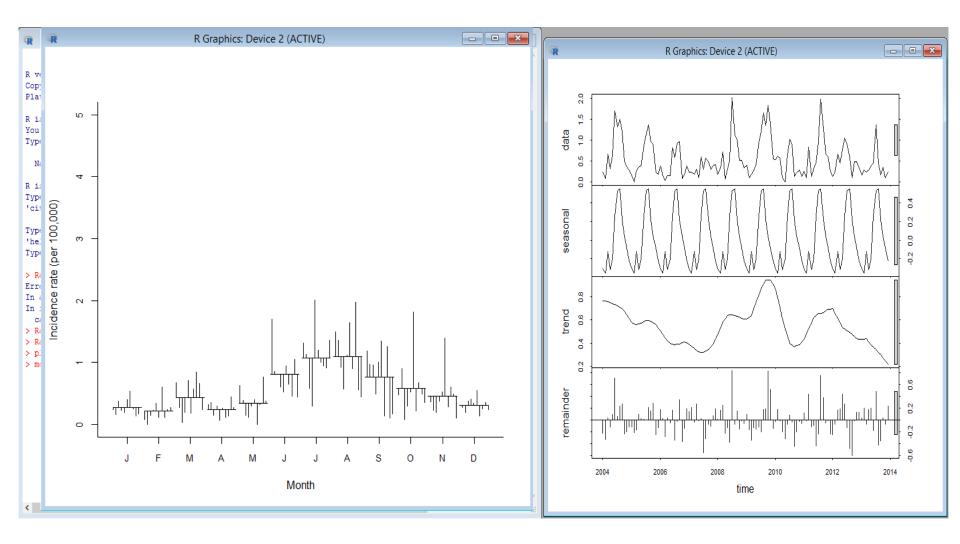
*Previous study showed that 17~71% of VE were caused by JE in Vietnam

Seasonality of VE in humans between 2004 and 2013 (Dien Bien, Hoa Binh, Lai Chau and Son La)



*Previous study showed that 17~71%

Screenshot (R package)



Exercises (15 mins)

- VE monthly incidence rate from 2004 to 2013
 - Open file: seasonality_VE and R_code_VE
 - Import into R:
 - Run the model
 - Interpretation

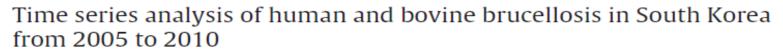
Time series analysis



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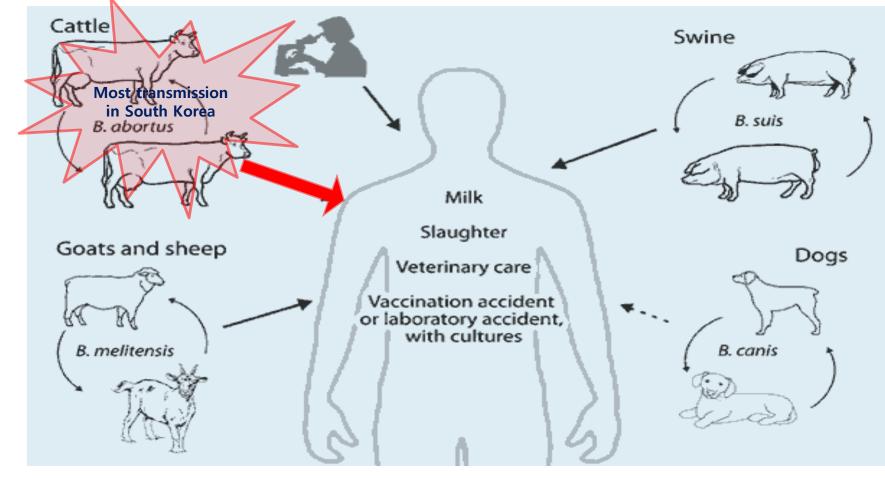
ABSTRACT

Brucellosis is considered to be one of the most important zoonotic diseases in the world, affecting underdeveloped and developing countries. The primary purpose of brucellosis control is to prevent the spread of disease from animals (typically ruminants) to humans. The main objective of this study was to retrospectively develop an appropriate time series model for cattle-to-human transmission in South Korea using data from independent national surveillance systems. Monthly case counts for cattle and people as well as national population data were available for 2005–2010. The temporal relationship was evaluated using an autoregressive integrated moving average with exogenous input (ARIMAX) model [notated as ARIMA(p, d, q) – AR(p)] and a negative binomial regression (NBR) model.

Human incidence rate was highly correlated to cattle incidence rate in the same month and the previous month (both r-0.82). In the final models, ARIMA (0, 1, 1) – AR (0, 1) was determined as the best fit with 191.5% error in the validation phase, whereas the best NBR model including lags (0, 1 months) for the cattle incidence rate yielded a 131.9% error in the

Brucellosis

Transmission to Humans



Introduction of project

Rationale

 Most human cases are related to not wearing protection, such as gloves and protective clothing when in contact with suspected cattle or materials

Objective

 To develop an appropriate time series model for cattle and humans in South Korea using data from independent national surveillance systems

Materials

- Study period: Jan 1, 2005 ~ Dec 31, 2010
- Data sources
 - Data collected on a yearly & quarterly basis:
 - **KOSIS** : national total population for human & cattle (Korean Statistical Information Service)
 - Data collected on a monthly basis:
 - **KCDC :** number of human cases (Korea Centers for Disease Control and Prevention)
 - **AIMS** : number of cattle cases (Animal Infectious Disease Data Management System)

Methods

- Human & cattle incidence rates were calculate d on a monthly basis

 Cases / national total population
 The ARIMAX model is an extension of ARIMA (autoregressive integrated moving average) model
 the ARIMA [Constant + Y_{t-1} + \$\phi((Y_{t-1} - Y_{t-2})) - \$\theta\$ et..] + AR[\$\beta\$_{t-1}X_{t-1} +
- ARIMAX (autoregressive integrated moving average with exogenous input) model
 - Cross correlation between human and cattle incidence rates
 - Value of P < 0.05 was considered significant
- Model validation
 - Divided into model: construction (2005-2007) and validation (2008-2010) phases

Methods

 Conducted simulation intervention scenarios (50% and 75% reductions in cattle cases)

• *MAPE =
$$\left(\frac{100\%}{n}\sum_{t=1}^{n} \left| \frac{\text{Actual cases-predicted cases}}{\text{Actual cases}} \right| \right)$$

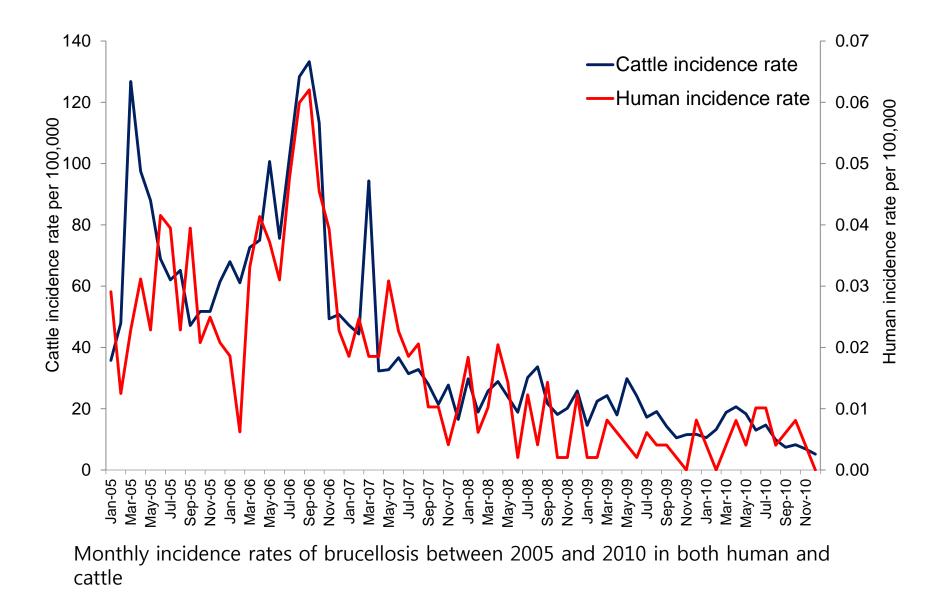
*Mean Absolute Percentage Error

- Software programs:
 - Microsoft Excel (Redmond, WA, USA)
 - STATA version 11.2 (Stata Corp., College Station, TX, USA)

Actual cases



Descriptive statistics



Descriptive statistics

Month	Human cases	Human (% of total)	Cattle cases	Cattle (% of total)
Jan	44	7.50	5,150	6.91
Feb	25	4.26	5,184	6.96
Mar	47	8.01	8,764	11.76
Apr	61	10.39	6,878	9.23
May	55	9.37	7,460	10.01
Jun	53	9.03	6,032	8.10
Jul	65	11.07	6,676	8.96
Aug	56	9.54	7,427	9.97
Sep	66	11.24	6.431	8.63
Oct	43	7.33	5,725	7.69
Nov	36	6.13	4,345	5.83
Dec	36	6.13	4,421	5.93
Total	587	100	74,493	100

Cases of human and cattle brucellosis reported to the KCDC and AIMS between Jan 1 2005 and Dec 31 2010

Cross-correlation results

human and cattle incidence rates between 2005 and 2010 on a monthly basis with lags of 0-12 months

Lags*(months)	Correlation(r)
0	0.82
1	0.82
2	0.79
3	0.74
4	0.67
5	0.57
6	0.52
7	0.47
8	0.44
9	0.41
10	0.40
11	0.35
12	0.40

* The time periods between two observations. For example, lag 1 is between Y_t and Y_{t-1} . Lag 2 is between Y_t and Y_{t-2} . Time series can also be lagged forward, Y_t and Y_{t+1} .

Variable	Lags(months)	Coefficient	Standard Error	P-value	MAPE		
ARIMAX best model (First difference) ^a : ARIMA (0, 1, 1) – AR (0, 1)							
Unpredictable factors	1	-0.86	0.10	<0.001	65.86%		
Cattle incidence rate	0	1.84 x 10-4	0.5 x 10-5	<0.001			
Cattle incidence rate	1	1.85 x 10-4	0.5 x 10-5	0.001			
Constant	N/A	8.13 x 10-11	1.66 x 10-9	0.961			

^aThe first difference of $Y_t = Y_{t-1} - Y_{t-2}$ (Y_t = human incidence rate at time t)

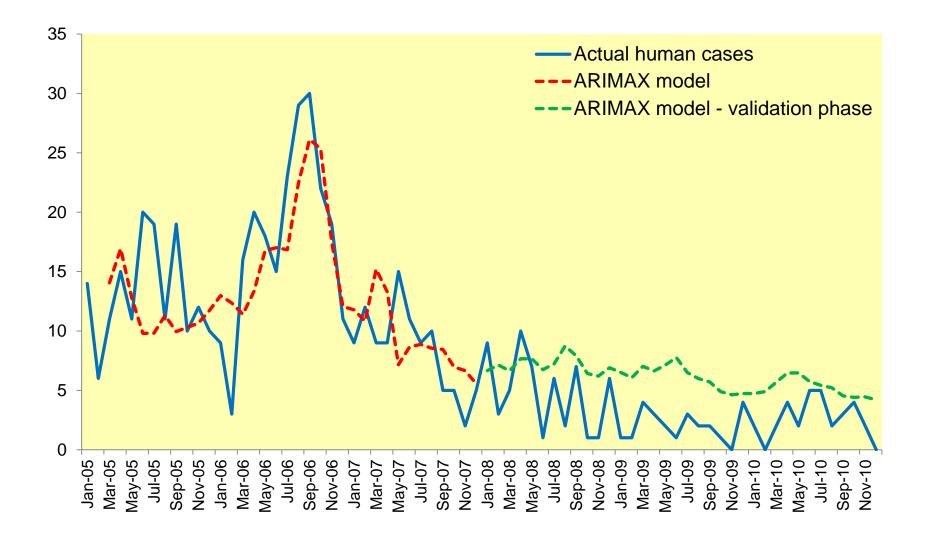
$\hat{Y}_{t} = [8.13 \times 10^{-11} + Y_{t-1} + 1.84 \times 10^{-4} X_{t} + 1.85 \times 10^{-4} X_{t-1} + 0.86 e_{t-1}] \times Y_{popt}$

 \hat{Y}_t = the predicted number of human cases at time t, X_t = the cattle incidence rate at time t e_{t-1} = unpredictable factors at time t-1, Y_{popt} = the human population at time t

Examples

- Model Calculations
 - Actual human cases: 30
 - Predicted human cases in Sep 2006 : 26.12
- 26.12= $[8.13 \times 10^{-11} + 6.00 \times 10^{-7} + 1.84 \times 10^{-4} \times 1.28 \times 10^{-3} + 1.85 \times 10^{-4} \times 1.33 \times 10^{-3} + 0.86 \times e_{Aug2006}] \times 48,372,000$
 - Actual human cases: 5
 - Predicted human cases in July 2010: 2.12
- $3.95 = [8.13 \times 10^{-11} + 1.01 \times 10^{-7} + 1.84 \times 10^{-4} \times 1.30 \times 10^{-4} + 1.85 \times 10^{-4} \times 0.15 \times 10^{-4} + 0.86 \times e_{Jun2010}] \times 49,410,000$

Predicted versus actual human cases



Limitations

- A limitation of the study potentially lies in not being able to utilize data at the individual province level
- Exposure history of human brucellosis was not available
 - Restricting our ability to adjust for delayed recognition
- Other potential risk factors were not taken into account in the model
 - Surveillance of the wildlife has not been actively implemented to date
 - Farm levels and / or environmental factors

Thank you





