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Department of Statistics Papers

Title

CGM and insulin pump data to introduce classical and machine learning time series analysis concepts to students

Permalink

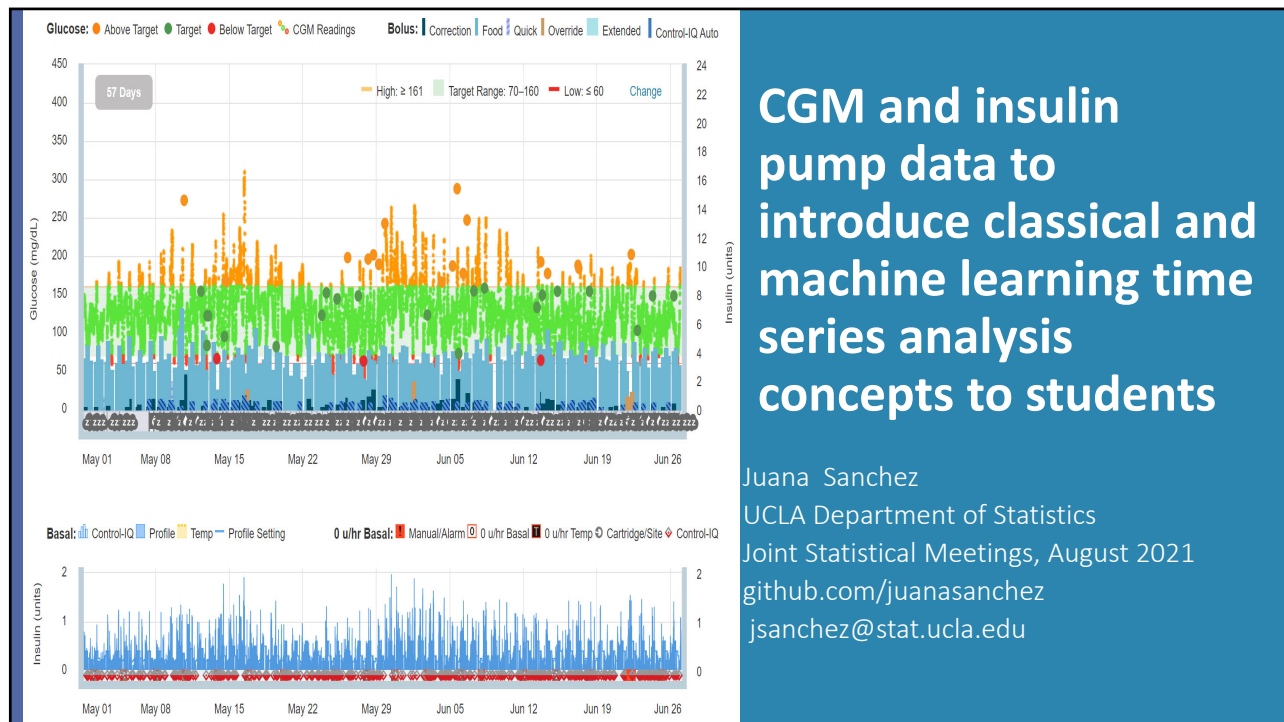
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Author

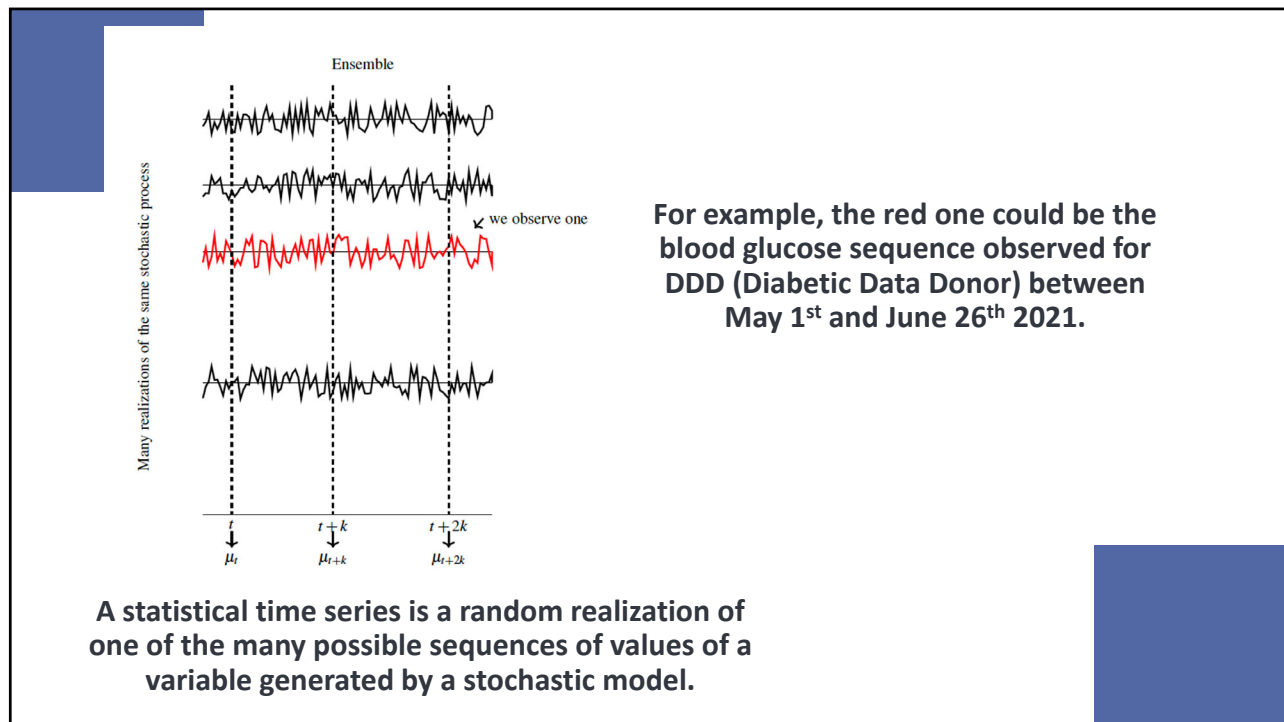
Sanchez, Juana

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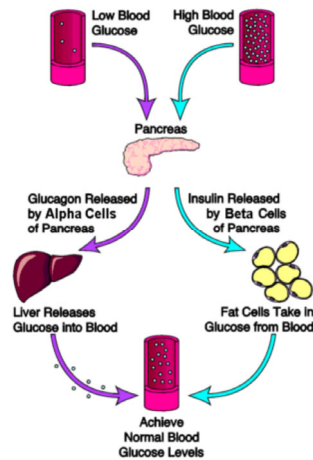
- Time series data analysis: rarely taught to undergraduates in college.
- Many students graduate with:
 - the i.i.d mindset,
 - without ever having corrected for autocorrelation in regression,
 - without ever using their regression models to predict out-of-sample.
- In the rare cases where an upper division elective time series course is offered to juniors and seniors,
 - the iid mindset interferes with the learning of basic time series concepts,
 - students have to invest a large amount of time learning concepts that could have easily been taught at the intro stats level.
- The volume, velocity and variety of timestamped data (smart cities, medical devices, finance, economy, climate, water quality, energy...) is making it increasingly necessary to include at least some basic time series education in the intro stats course.



This talk is a case study involving data produced by closed-loop technology for the management of T1D of the DDD. Follows GAISE recommendations.

1. **The context:** Blood glucose regulation for a T1D (Type I Diabetic) person
2. **The multivariate and real timestamped sensor data donated by a DDD** is produced by the technology with a purpose: to help non-statistician health care providers routinely guide DDD's health care management.
3. **Our objective is** to use this context and data to engage students in the discovery of time series features by means of basic graphs and summary statistics that they learn in an introductory statistics course, by investigations and critical thinking.

1. The context: blood glucose regulation

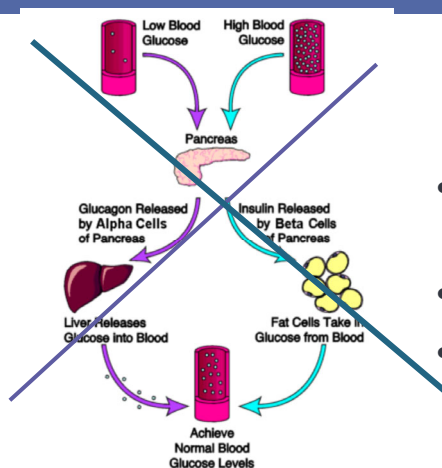


Low and high blood sugar are self-regulated by the pancreas. When the pancreas functions well:

- **If high blood sugar level:** Insulin released by the Beta Cells of Pancreas makes sugar go to cells to produce energy. This helps maintain normal sugar levels in the blood.
- **If low blood sugar level:** Glucagon released by Alpha Cells of Pancreas makes the liver release glucose to the blood. This helps maintain normal sugar levels in the blood. Not desirable even in normal persons. Better eat that cookie.

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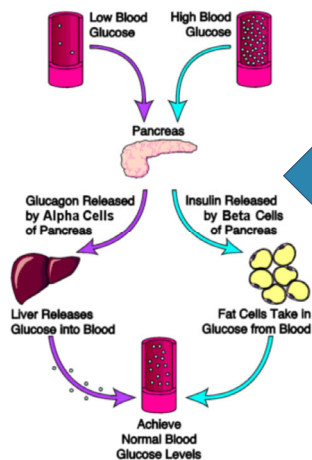
For a T1D (for 1.6 million people in the US of every age, race, shape, and size)



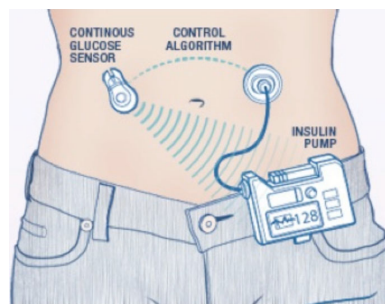
- **Normal blood glucose self regulation does not work.**
- **All the blood sugar stays in the blood.**
- **Artificial insulin is needed.**

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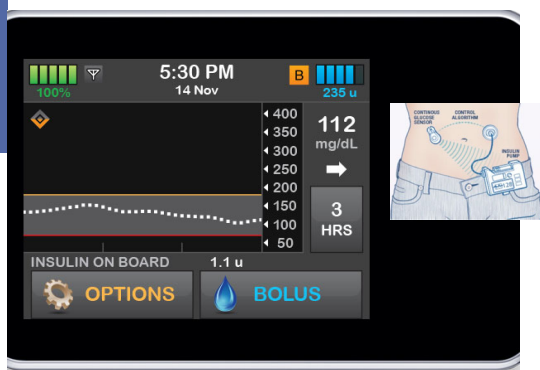
Since Jan 2020 close-loop system technology adjusts insulin levels and provides readings of interstitial blood glucose. The DDD in this case study uses the technology.



Objective: mimic normal blood glucose control



DDD's view of the data produced by the technology



DDD can view sensor glucose readings in the insulin pump display up to 24 hours



DDD can also view sensor glucose readings in the smart phone

DDD can see more than glucose readings

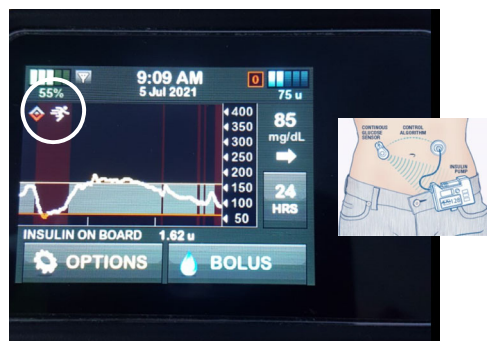


DDD also validates cgm readings by periodically checking “true” blood sugar manually and periodically intervenes.



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The algorithm is unknown



The pump seems to act based on predictions of blood glucose for the next half hour. When sleeping, that works very well. When awake, with unexpected events, body reactions, food, etc. it gets complicated. DDD would like to understand the secret algorithm inside the technology better.

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EventDateTime	Readings (CGM / BGM)
2021-05-01T00:02:00	122
2021-05-01T00:07:00	125
2021-05-01T00:12:00	127
2021-05-01T00:17:00	129
2021-05-01T00:22:00	132
2021-05-01T00:27:00	134
2021-05-01T00:32:00	133
2021-05-01T00:37:00	132
2021-05-01T00:42:00	133
2021-05-01T00:47:00	133
2021-05-01T00:52:00	133
2021-05-01T00:57:00	
2021-05-01T01:02:00	
2021-05-01T01:07:00	
2021-05-01T01:12:00	
2021-05-01T01:17:00	

The technology stores DDD's timestamped data and plots.

IOB	81	2021-05-01T05:40:17	0.28
IOB	81	2021-05-01T05:50:18	0.25
IOB	81	2021-05-01T06:00:17	0.21
IOB	9	2021-05-01T06:05:40	0.1
IOB	9	2021-05-01T06:05:49	0.1
IOB	81	2021-05-01T06:10:17	0.19
IOB	9	2021-05-01T06:15:22	0.1
IOB	16	2021-05-01T06:15:23	0.19
IOB	64	2021-05-01T06:15:41	0.19
IOB	55	2021-05-01T06:15:56	0.19
IOB	20	2021-05-01T06:17:12	3.8
IOB	81	2021-05-01T06:17:12	0.19
IOB	81	2021-05-01T06:20:17	3.82
IOB	81	2021-05-01T06:30:17	3.75
IOB	81	2021-05-01T06:40:17	3.64
IOB	81	2021-05-01T06:50:17	3.43
IOB	81	2021-05-01T07:00:17	3.22
IOB	81	2021-05-01T07:10:18	2.97
IOB	81	2021-05-01T07:20:17	2.68

The purpose of the data storage: to allow non-statistician nurses and doctors inspect all the plots and summaries every three months to guide DDD's adjustments to therapy.

Low Summary

- High (>=181 mg/dL): 14% (2279 readings)
- Above Target (181-180 mg/dL): 0% (0 readings)
- Within Target (70-180 mg/dL): 81% (12921 readings)
- Below Target (60-69 mg/dL): 2% (438 readings)
- Low (<=60 mg/dL): 2% (345 readings)

Average CGM: 125 mg/dL
Standard Deviation: 35.3 mg/dL

Insulin Delivery Summary

- Avg. Daily Basal Units: 35% (7.10 units)
- Avg. Daily Food Bolus: 55% (11.77 units)
- Avg. Daily Correction Bolus: 8% (1.54 units)
- Avg. Daily Quick Bolus: 0% (0.32 units)
- Avg. Daily Override Bolus: 0% (0.19 units)

Average Total Daily Dose: 20.23 units/day
Average Daily Carbs: 188 grams/day

High Summary

- High (>=181 mg/dL): 38% (18 readings)
- Above Target (181-180 mg/dL): 0% (0 readings)
- Within Target (70-180 mg/dL): 55% (26 readings)
- Below Target (60-69 mg/dL): 6% (3 readings)
- Low (<=60 mg/dL): 0% (0 readings)

Average BG: 150 mg/dL
Standard Deviation: 54 mg/dL

Bolus Usage Summary

- Food Only Boluses: 52% (111 boluses)
- Correction + Food Boluses: 32% (67 boluses)
- Correction Boluses: 7% (15 boluses)
- Quick Boluses: 7% (15 boluses)
- Override Boluses: 2% (4 boluses)

Average Boluses: 5.37/day
Skipped Boluses: 0.00 boluses

Time Period	Low	Below	Target	Above	High
Night 12am - 6am	0%	0%	91%	0%	9%
Morning 6am - 12pm	3%	1%	76%	0%	20%
Afternoon 12pm - 6pm	6%	4%	59%	0%	32%
Evening 6pm - 12am	8%	7%	60%	0%	25%

This graph shows your data averaged over 90 days

Graphs and summary statistics are not quite those taught at the intro stats level.

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Lots of time series graphs and data stored in the technology's server

Hourly average glucose readings summarized by hour of the day.
Something close to seasonal box plots

Some kind of trends
Something close to trends during the day

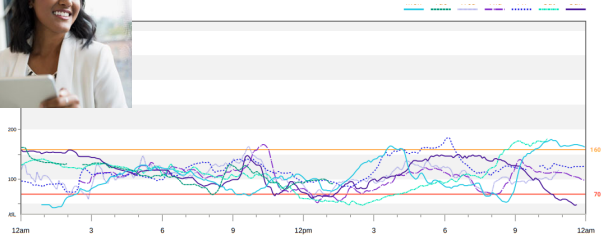
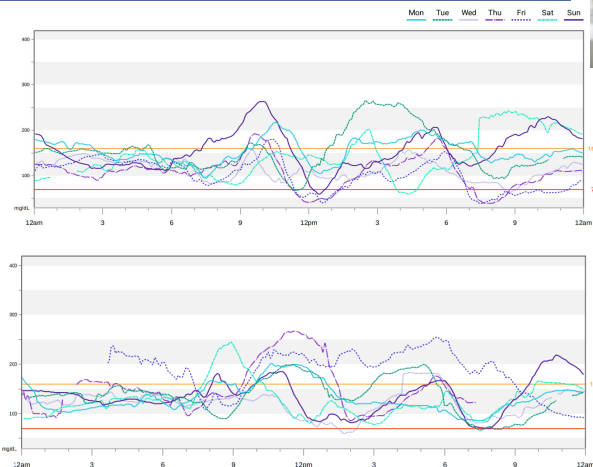
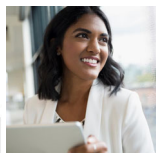
Spaghetti plots
Something close to spaghetti plots

Streaming data.
Something like multivariate time plots, customized, but not line plots.

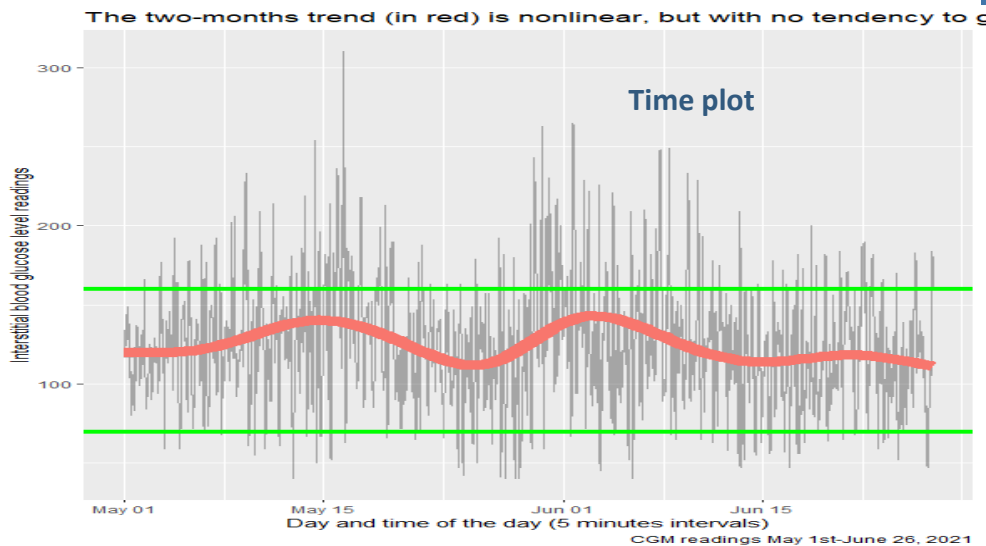
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Not all weeks are the same, not all days are the same

In some weeks, all days are roller-coasters



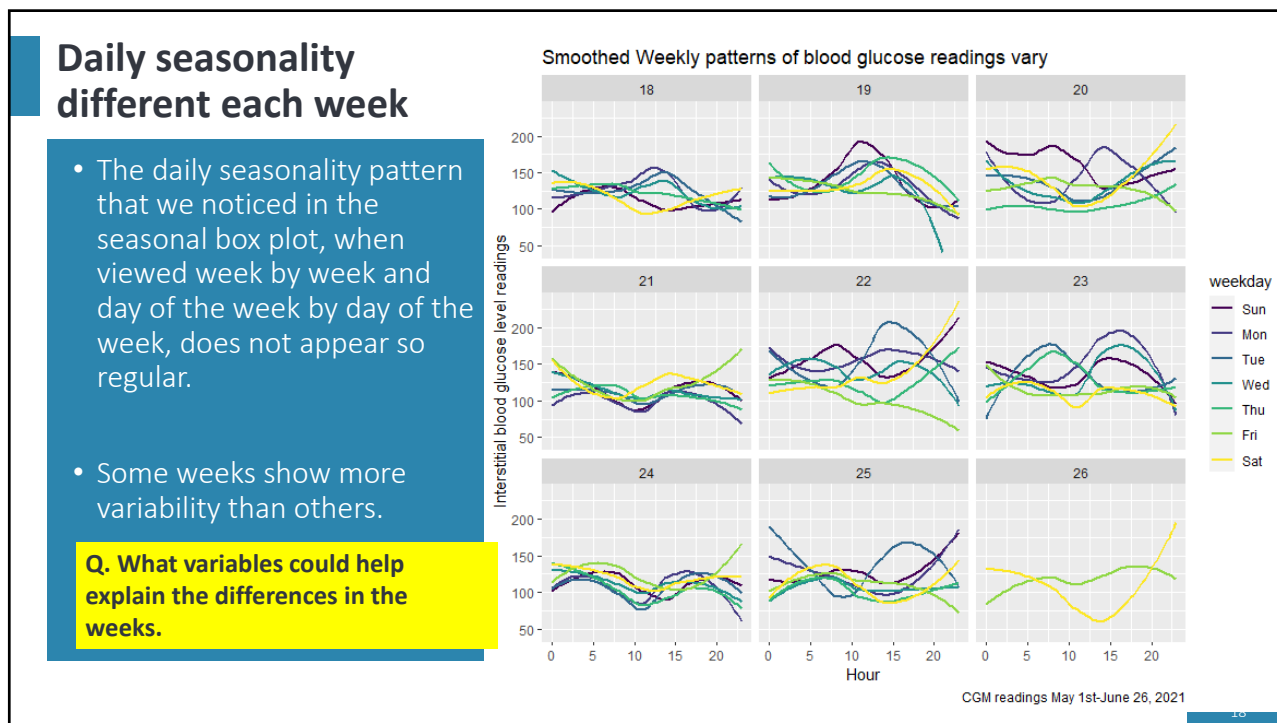
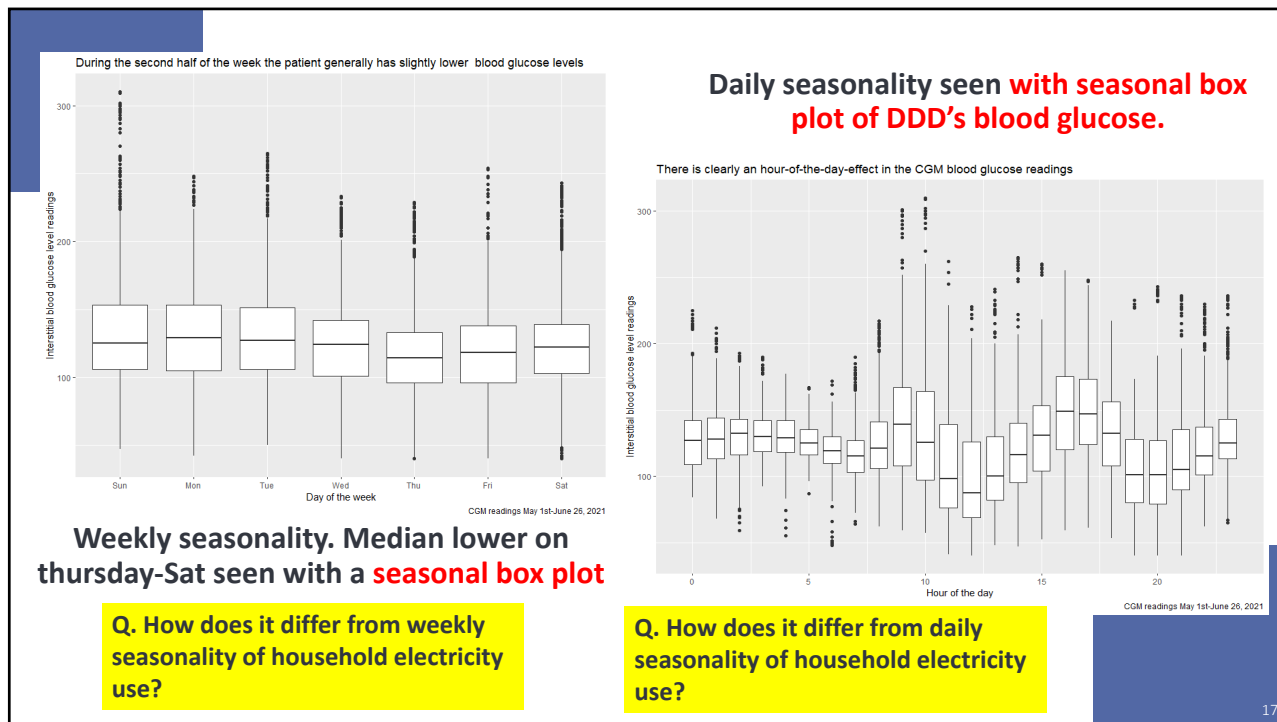
There is a rich story to tell students with just blood glucose built-in graphs. We analyze 16017 observations of blood glucose of May 1st-June 26, 2021 and 9055 observations of insulin interventions during the same period.



Non-linear trend according to a moving average smoother

Q. What does this plot reveal about the technology? About DDD? Would you expect this trend in a time series of household electricity use recorded at 5 minute intervals?

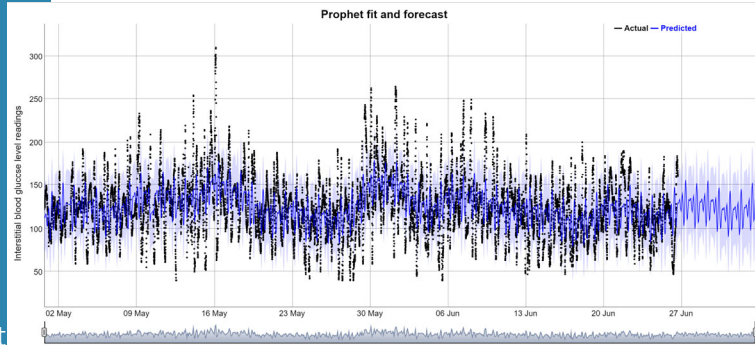
3. Teaching basic time series concepts in the intro stats class with R



Forecasting like analysts in the loop with Prophet

- Cannot expect intro stats students to forecast with sophisticated time series models.
- But students without any time series training work sometimes as analysts in the loop (use a proprietary automated forecasting routine). Facebook's prophet is an example.

Q. What do you observe about the forecasts out-of-sample as compared with the training data?



Glucose = trend + hour +day of week+holidays

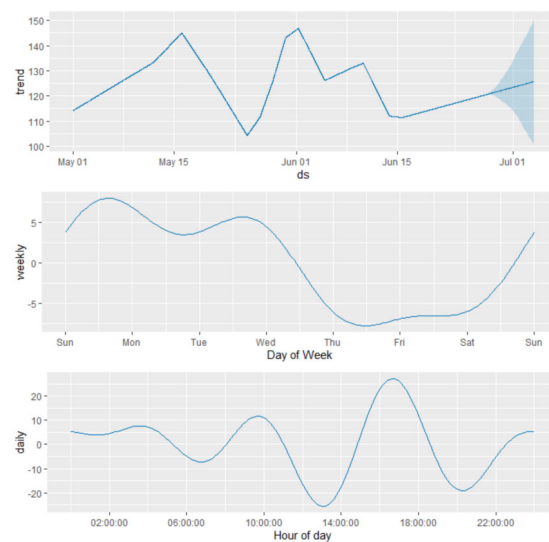
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Forecasting like analysts in the loop with Prophet

- Prophet models the components of the time series that we analyzed with our plots.
- Glucose = trend + hour +day of week+holidays

Q. What is happening to our uncertainty when we forecast farther ahead?

Another way to do time series decomposition.



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Machine learning: features engineering and clustering to be able to combine insulin and glucose data

- We involve now the insulin interventions by the pump and DDD, but because the recording time interval is different than for glucose we use features engineering :


- Use summaries familiar to the intro stats student: interquartile range, min, max, median, frequencies of different types of boluses.

- By the hour.

```
# A tibble: 1,367 x 20
# Groups:   day [57]
  day hour iob_med_hour iob_sd_hour iob_IQR_hour iob_min_hour iob_max_hour iobmonth iobday n_81 n_9 n_16 n_20 n_55 n_64 cgm_med_hour
<dbl> <int> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl> <dbl>
1 121 0 0.1 0.0742 0.128 0.01 0.18 5 1 6 0 0 0 0 0 132
2 121 1 0.15 0.0186 0.0225 0.12 0.17 5 1 6 0 0 0 0 0 128.
3 121 2 0.26 0.0235 0 0.21 0.28 5 1 6 0 0 0 0 0 138
4 121 3 0.325 0.00816 0.0100 0.32 0.34 5 1 6 0 0 0 0 0 143
5 121 4 0.385 0.0387 0.0600 0.32 0.41 5 1 6 0 0 0 0 0 148.
6 121 5 0.33 0.0520 0.0775 0.25 0.38 5 1 6 0 0 0 0 0 138.
7 121 6 0.19 1.76 3.40 0.1 3.82 5 1 7 3 1 1 1 1 135
8 121 7 2.54 0.505 0.697 1.89 3.22 5 1 6 0 0 0 0 0 122.
9 121 8 1.01 0.420 0.595 0.53 1.62 5 1 6 0 0 0 0 0 106
10 121 9 0.19 0.136 0.18 0.03 0.37 5 1 5 0 0 0 0 0 105
# ... with 1,357 more rows, and 4 more variables: cgm_sd_hour <dbl>, cgm_IQR_hour <dbl>, cgm_min_hour <dbl>, cgm_max_hour <dbl>
```

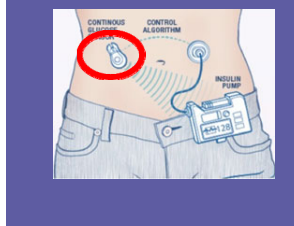
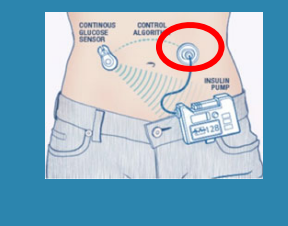

Clustering of features

Q. What explains the differences in the clusters of the hours? Investigate using data not used in clustering.

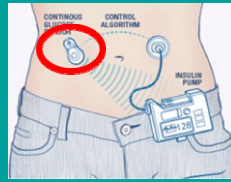


iob_med_hour
0.8557059
0.6131099
1.1281836

cgm_med_hour
88.41529
127.66491
177.18164

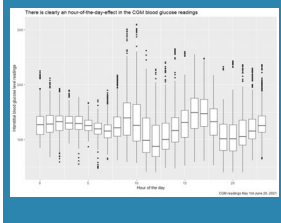



iob_IQR_hour
0.6495647
0.4184902
0.4602930

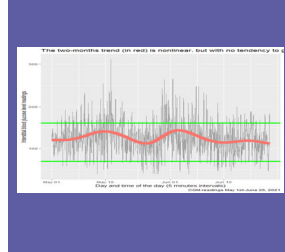


cgm_IQR_hour
10.304118
9.724774
13.486328

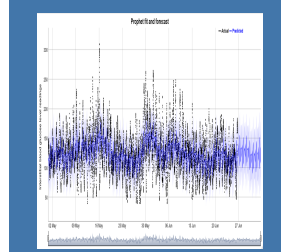
Conclusion: the case study engages students and makes them use the tools they know to investigate a complex process. At the same time, they learn basic time series concepts using only their intro stats tools.



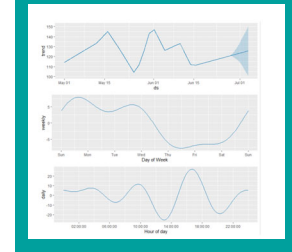
The concept of seasonality



The concept of long term trend



The concept of forecast



The concept of decomposition of a time series into its components

And with all of the above, an intuition for the concept of autocorrelation is gained before introducing the ACF and PACF. Of course, having time series for more than one DDD would make the study more interesting. After all, the technology designers are inspired by time series like ours of thousands of individuals and design to target the average individual.

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Thank you for your attention

The paper (with references), simple R programs and data for this talk can be found at [github/juanasanchez](https://github.com/juanasanchez) shortly after the JSM meetings.
jsanchez@stat.ucla.edu

I thank the DDD for the data and for all the information provided that helped me understand the data. Without the DDD's help I would not have been able to complete this presentation.

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