Simulate of Gestational Periods (Binomial), Conception Dates (Mixture), and Temperature Exposures (Sine wave and AR(1)) during Pregnancy

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1 Conception, Birth and Extreme Temperature

The various simulations on this page are done in support of Same Environment, Stratified Impacts? Air Pollution, Extreme Temperatures, and Birth Weight in South China.

Go to the RMD, R, PDF, or HTML version of this file. Go back to fan's REconTools Package, R Code Examples Repository (bookdown site), or Intro Stats with R Repository (bookdown site).

1.1 Gestational Age at Birth Distribution (Binomial)

Suppose the number of weeks that an individual is pregnant follows the binomial distributions. According to data from the Right from the Start study from West Virginia, published in Hoffman et al. (2008), median gestational age at birth was 276/7 weeks and the standard deviation was around 14/7 weeks. We use binomial to approximate normal rules to generate the relevant binomial parameters.

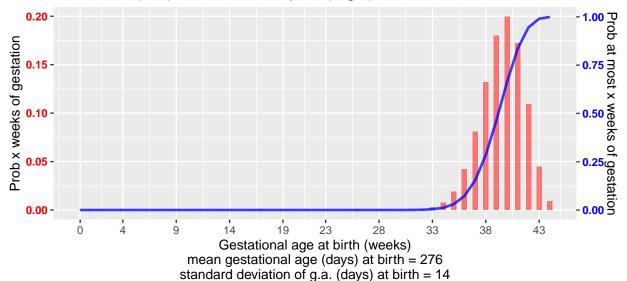
On each date after conception, there is some chance to give birth, due to various purely random factors, denote this chance as $p_{\tau,i}$, where τ is discrete time since conception, $p_{\tau,i}$ follows the just defined binomial distribution. The distribution might or might not be individual specific. They would be individual-specific if individual experiences during the course of pregnancy, such as exposures to extreme temeprature, matter for gestational age at birth. The probability that an individual i gives birth during the week of conception is $p_{0,i}$. Conditional on proceeding to the day after conception, the probability of giving birth is $\frac{p_{1,i}}{1-p_{0,i}}$. Similarly for future dates. The conditional birth probability is the Hazard rate in this scenario.

Below, we formulate the gestational age at birth distribution function ffi_gestation_age_at_birth_dist().

```
# This function generates the distribution of gestational weeks at birth
ffi_gestation_age_at_birth_dist <- function(</pre>
  mu_gabirth_days = 276,
  sd_gabirth_days = 14
){
  # # Parameters
  # # gabirth = gestational age at birth
  # mu_gabirth_days <- 276</pre>
  # sd_gabirth_days <- 14</pre>
  # from https://fanwangecon.github.io/R4Econ/statistics/discrandvar/htmlpdfr/fs disc approx cts.html
  it_binom_n <- round((mu_gabirth_days / 7)^2 / (mu_gabirth_days / 7 - (sd_gabirth_days / 7)^2))
  fl_binom_p <- 1 - (sd_gabirth_days / 7)^2 / (mu_gabirth_days / 7)</pre>
  # Same graphing code as from: https://fanwangecon.github.io/Stat4Econ/probability_discrete/htmlpdfr/b
  # Generate Data
  ar_grid_gabirth <- 0:it_binom_n</pre>
  ar_pdf_gabirth <- dbinom(ar_grid_gabirth, it_binom_n, fl_binom_p)</pre>
  ar_cdf_gabirth <- pbinom(ar_grid_gabirth, it_binom_n, fl_binom_p)</pre>
  df_dist_gabirth <- tibble(gabirth = (ar_grid_gabirth), prob = ar_pdf_gabirth, cum_prob = ar_cdf_gabir
  # Two axis colors
  axis_sec_ratio <- max(ar_cdf_gabirth) / max(ar_pdf_gabirth)</pre>
  right_axis_color <- "blue"
  left_axis_color <- "red"</pre>
  # Probabilities
  plt_dist_gabirth <- df_dist_gabirth %>%
    ggplot(aes(x = gabirth)) +
    geom_bar(aes(y = prob),
      stat = "identity", alpha = 0.5, width = 0.5, fill = left_axis_color
    )
  # Cumulative Probabilities
  plt_dist_gabirth <- plt_dist_gabirth +</pre>
    geom_line(aes(y = cum_prob / axis_sec_ratio),
      alpha = 0.75, size = 1, color = right_axis_color
  # Titles Strings etc
```

```
graph_title <- paste0("Gestational age at birth (weeks)\n",</pre>
      "Prob mass (Left) and cumulative prob (Right)")
  graph\_caption \leftarrow pasteO("Assuming the binomial properties apply\n",
      "fl_binom_p = ", fl_binom_p, ", it_binom_n = ", it_binom_n)
  graph_title_x <- paste0("Gestational age at birth (weeks)\n",</pre>
    "mean gestational age (days) at birth = ", mu_gabirth_days, "\n",
    "standard deviation of g.a. (days) at birth = ", sd_gabirth_days)
  graph title y axisleft <- "Prob x weeks of gestation"</pre>
  graph_title_y_axisright <- "Prob at most x weeks of gestation"</pre>
  # Titles etc
  plt_dist_gabirth <- plt_dist_gabirth +</pre>
    labs(
      title = graph_title,
      x = graph_title_x,
      y = graph_title_y_axisleft,
      caption = graph_caption
    scale_y_continuous(
      sec.axis =
        sec_axis(~ . * axis_sec_ratio, name = graph_title_y_axisright)
    scale_x_continuous(
      labels = ar_grid_gabirth[floor(seq(1, it_binom_n, length.out = 10))],
      breaks = ar grid gabirth[floor(seq(1, it binom n, length.out = 10))]
    ) +
    theme(
      axis.text.y = element_text(face = "bold"),
      axis.text.y.right = element_text(color = right_axis_color),
      axis.text.y.left = element_text(color = left_axis_color)
    )
  # Print
  return(list(
    df_dist_gabirth=df_dist_gabirth,
    plt_dist_gabirth=plt_dist_gabirth
  ))
# Test the function
ls_gsbirth <- ffi_gestation_age_at_birth_dist(mu_gabirth_days = 276, sd_gabirth_days = 14)</pre>
# Figure
print(ls_gsbirth$plt_dist_gabirth)
```

Gestational age at birth (weeks) Prob mass (Left) and cumulative prob (Right)



Assuming the binomial properties apply fl_binom_p = 0.898550724637681, it_binom_n = 44

# Table	
df_dist_gabirth <- ls_gsbirth\$df_dist_gabirth	
<pre>kable(df_dist_gabirth %>% filter(prob >= 0.01)) %>% kable_styling_fc()</pre>	

gabirth	prob	cum_prob
35	0.0190913	0.0303855
36	0.0422736	0.0726591
37	0.0809563	0.1536154
38	0.1320866	0.2857020
39	0.1799862	0.4656882
40	0.1992704	0.6649586
41	0.1721918	0.8371504
42	0.1089377	0.9460881
43	0.0448780	0.9909661

Some research on the distribution of gestational ages at birth:

- Gage (2002)
- Nassar et al. (2013)
- Lei et al. (2016)
- Pereira et al. (2021)

1.2 Conception Dates Distribution (Mixture Binomials + Random)

Denote week of the year with w. There are potentially 52 possible weeks of the year, starting at week 1, w = 1, and ending with w = 52. Conception can happen at any month.

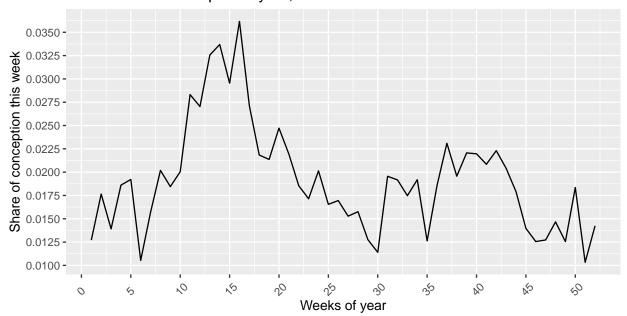
Suppose the conception week of the year distribution is bimodal, we will pick two max conception count weeks, out of 52 weeks. Suppose week 15 and week 40 are two peak conception months, with week 15 more heavily weighted. What is the distribution of conception weeks? We will mix two binomial distribution with a largely-uniform distribution together. We will assume later that the day (of the week) of conception is uniformly distributed.

Below, we formulate the conception date distribution function ffi_concept_distribution_year().

```
# This function generates the distribution of conception weeks across
# the year, with two peak months, and some randomness
ffi_concept_distribution_year <- function(it_max_weeks = 52,</pre>
                                            it peak wk 1st = 15,
                                            it_peak_wk_2nd = 40,
                                            fl_binom_1st_wgt = 0.150,
                                            fl_binom_2nd_wgt = 0.075,
                                            it runif seed = 123,
                                            df_dist_conception_exo = NULL) {
  # # Peak (local) months and weights
  # it_max_weeks <- 52
  # it_peak_wk_1st <- 15
  # it_peak_wk_2nd <- 40
  # # Weights for the two binomial and the remaining weight is for an uniform distribution
  # fl_binom_1st_wqt <- 0.25
  \# fl\_binom\_2nd\_wgt \leftarrow 0.10
  if (is.null(df_dist_conception_exo)) {
    # Discrete random variables
    ar_fl_binom_1st <- dbinom(</pre>
      0:(it_max_weeks - 1), (it_max_weeks - 1),
      (it_peak_wk_1st - 1) / (it_max_weeks - 1)
    ar_fl_binom_2nd <- dbinom(</pre>
      0:(it_max_weeks - 1), (it_max_weeks - 1),
      (it_peak_wk_2nd - 1) / (it_max_weeks - 1)
    set.seed(it_runif_seed)
    ar_random_base <- runif(it_max_weeks, min = 0.5, max = 1)</pre>
    ar_random_base <- ar_random_base / sum(ar_random_base)</pre>
    # Mix two binomials and a uniform
    ar_fl_p_concept_week <- ar_fl_binom_1st * fl_binom_1st_wgt +</pre>
      ar_fl_binom_2nd * fl_binom_2nd_wgt +
      ar_random_base * (1 - fl_binom_1st_wgt - fl_binom_2nd_wgt)
    # Dataframe
    df_dist_conception <- tibble(conception_calendar_week = 1:it_max_weeks,</pre>
                                 conception_prob = ar_fl_p_concept_week)
  } else {
    df_dist_conception <- df_dist_conception_exo</pre>
  }
  # Line plot
  # Title
  st_title <- paste0(</pre>
      "Distribution of conception week of birth\n",
      "over weeks of one specific year, seed=", it_runif_seed
```

```
# Display
  plt_concept_week_of_year <- df_dist_conception %>%
    ggplot(aes(x = conception_calendar_week, y= conception_prob)) +
    geom_line() +
    labs(
      title = st_title,
      x = 'Weeks of year',
      y = 'Share of conception this week'
      ) +
    scale_x_continuous(n.breaks = 12) +
    scale_y_continuous(n.breaks = 10) +
    theme(
      axis.text.x = element_text(angle = 45, vjust = 0.1, hjust = 0.1)
    )
  # Return
  return(list(
    df_dist_conception = df_dist_conception,
    plt_concept_week_of_year = plt_concept_week_of_year
  ))
# Call function with defaults
ls_concept <- ffi_concept_distribution_year(it_max_weeks = 52,</pre>
                                             it_peak_wk_1st = 15,
                                             it_peak_wk_2nd = 40,
                                             it_runif_seed = 123)
ls_concept$plt_concept_week_of_year
```

Distribution of conception week of birth over weeks of one specific year, seed=123



```
df_dist_conception <- ls_concept$df_dist_conception
kable(df_dist_conception) %>% kable_styling_fc()
```

1.3 Temperature Process Sine Wave and AR(1)

Let t index day of survey. We model temperature in Fahrenheit as F_t using an AR(1) persistent process and a sine function. According to Wikipedia, which reports Climate data for Guangzhou, China, daily mean temperature varies between 57F (Jan) and 84F (June). Record high temperature was reported in July at 102.4F. Record low temperature was reported in December at 32.0F. We will mark out these temperature levels visually in graphs below.

Specifically, temperature in Fahrenheit is modeled as:

$$F_t = \exp\left(\mu + \epsilon_t + \gamma \sin\left(\frac{W_t}{52} \cdot 2 \cdot \pi + \frac{3 - \frac{1}{6}}{2} \cdot \pi\right)\right)$$

where γ scales the sine curve, and the shock process ϵ follows a AR(1) process:

$$\epsilon_t = \rho \cdot \epsilon_{t-1} + \nu_t$$

with $0 < \rho < 1$, and $\nu \sim N(0, \sigma_{\nu})$. Using the function above, the coldest month will be somewhere around january.

Below, we formulate the temperature simulation function ffi_daily_temp_simulation().

```
ffi daily temp simulation <- function(
  fl_mthly_mean_lowest = 57,
  fl_mthly_mean_highest = 84,
  fl_record_lowest = 32,
  fl_record_highest = 102.4,
  it_weeks_in_year = 52,
  it_days_in_week = 7,
  it_years = 6,
  fl_mu = 4.15,
  fl_sin_scaler = 0.20,
  fl_sigma_nv = 0.15,
  fl_rho_persist = 0.40,
  it rand seed = 123,
  st_extreme_cold_percentile = "p05",
  st_extreme_heat_percentile = "p95"
  # # Guangzhou temp info
  # fl_mthly_mean_lowest <- 57
  # fl mthly mean highest <- 84
  # fl_record_lowest <- 32
  # fl_record_highest <- 102.4
  # # Total number of periods (over three years)
  # it_weeks_in_year <- 52</pre>
  # it_days_in_week <- 7</pre>
  # it years <- 6
  it_max_days <- it_weeks_in_year*it_days_in_week</pre>
  T <- it_max_days * it_years</pre>
  # # Mean temp
  # fl_mu <- 4.15
```

```
# fl_sin_scaler <- 0.20
 # # AR 1 parameter
 # fl sigma nv <- 0.15
 # fl_rho_persist <- 0.40
 # Generate a vector of shocks
 set.seed(it_rand_seed)
 ar_nv_draws <- rnorm(T, mean = 0, sd = fl_sigma_nv)</pre>
 # Generate a vector of epsilons
 ar_epsilon_ar1 <- vector("double", length=T)</pre>
 ar_epsilon_ar1[1] <- ar_nv_draws[1]</pre>
 for (it_t in 2:T) {
   ar_epsilon_ar1[it_t] <- fl_rho_persist*ar_epsilon_ar1[it_t-1] + ar_nv_draws[it_t]</pre>
 # Generate week by week sin curve values
 ar_day_at_t <- rep(1:it_max_days, it_years)</pre>
 ar_year_at_t <- as.vector(t(matrix(data=rep(1:it_years, it_max_days),</pre>
                                     nrow=it_years, ncol=it_max_days)))
 ar_base_temp \leftarrow sin((ar_day_at_t/it_max_days)*2*pi + ((3-1/6)/2)*pi)
 # Generate overall temperature in Fahrenheit
 ar_fahrenheit_city_over_t <-
   exp(fl_mu + ar_epsilon_ar1 + fl_sin_scaler*ar_base_temp)
 # Dataframe with Temperatures
 mt_fahrenheit_info <- cbind(ar_day_at_t, ar_year_at_t, ceiling(ar_day_at_t/it_days_in_week),</pre>
   ar_fahrenheit_city_over_t,
   exp(ar_base_temp), ar_epsilon_ar1, ar_nv_draws)
 ar_st_varnames <- c('survey_t','day_of_year', 'year', 'week_of_year',</pre>
   'Fahrenheit', 'FnoShock', 'AR1Shock', 'RandomDraws')
 # Combine to tibble, add name col1, col2, etc.
 df_fahrenheit <- as_tibble(mt_fahrenheit_info) %>%
   rowid_to_column(var = "t") %>%
   rename_all(~c(ar_st_varnames))
 # Generate extreme temperatures
 df_stats_fahrenheit <- REconTools::ff_summ_percentiles(df_fahrenheit, FALSE)</pre>
 # Add Extreme Thresholds
 fl_lowF_threshold <- df_stats_fahrenheit %>% filter(var == "Fahrenheit") %>% pull(st_extreme_cold_per
 fl_highF_threshold <- df_stats_fahrenheit %% filter(var == "Fahrenheit") %>% pull(st_extreme_heat_pe
 df_fahrenheit <- df_fahrenheit %>%
   mutate(extreme_cold = case_when(Fahrenheit <= fl_lowF_threshold ~ 1, TRUE ~ 0)) %>%
   mutate(extreme_hot = case_when(Fahrenheit >= fl_highF_threshold ~ 1, TRUE ~ 0))
 # REconTools::ff_summ_percentiles(df_fahrenheit, FALSE)
 # Title
 st_title <- pasteO('Simulated Temperature for Guangzhou (Sine Wave + AR(1))\n',
      'Each subplot is a different year\n',
      'RED = Guangzhou Temp 1971-2000 lowest and highest monthly averages \n',
      'BLUE = Guangzhou Temp 1961-2000 record lows and highs')
```

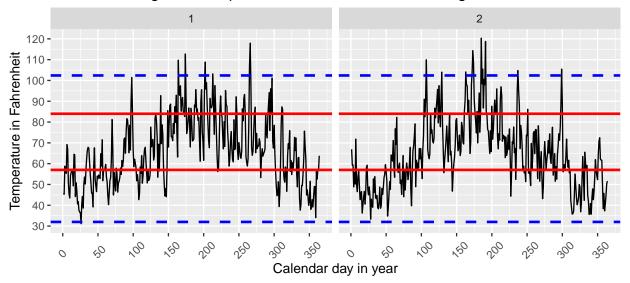
```
# Display
  plt_fahrenheit <- df_fahrenheit %>%
   ggplot(aes(x = ar_day_at_t, y=Fahrenheit)) +
   geom_hline(yintercept = fl_mthly_mean_lowest, linetype = "solid", colour = "red", size = 1) +
   geom_hline(yintercept = fl_mthly_mean_highest, linetype = "solid", colour = "red", size = 1) +
   geom_hline(yintercept = fl_record_lowest, linetype = "dashed", colour = "blue", size = 1) +
   geom hline(vintercept = fl record highest, linetype = "dashed", colour = "blue", size = 1) +
   facet_wrap(~ year) +
   labs(
     title = st_title,
     x = 'Calendar day in year',
     y = 'Temperature in Fahrenheit'
    scale_x_continuous(n.breaks = 12) +
   scale_y_continuous(n.breaks = 10) +
   theme(
      axis.text.x = element_text(angle = 45, vjust = 0.1, hjust = 0.1)
   )
  # Return
  return(list(
   df fahrenheit = df fahrenheit,
   plt_fahrenheit = plt_fahrenheit
  ))
}
```

Having constructed the temperature simulation function, first we call it with AR1 shock + sin curve, persistence is high at $\rho = 0.7$.

```
# Test 1: Call function with AR1 + Since Curve
ls_fahrenheit <- ffi_daily_temp_simulation(
  fl_mthly_mean_lowest = 57,
  fl_mthly_mean_highest = 84,
  fl_record_lowest = 32,
  fl_record_highest = 102.4,
  it_weeks_in_year = 52,
  it_days_in_week = 7,
  it_years = 2,
  fl_mu = 4.15,
  fl_sin_scaler = 0.25,
  fl_sigma_nv = 0.15,
  fl_rho_persist = 0.70,
  it_rand_seed = 123)
print(ls_fahrenheit$plt_fahrenheit)</pre>
```

Simulated Temperature for Guangzhou (Sine Wave + AR(1)) Each subplot is a different year

RED = Guangzhou Temp 1971–2000 lowest and highest monthly averages BLUE = Guangzhou Temp 1961–2000 record lows and highs

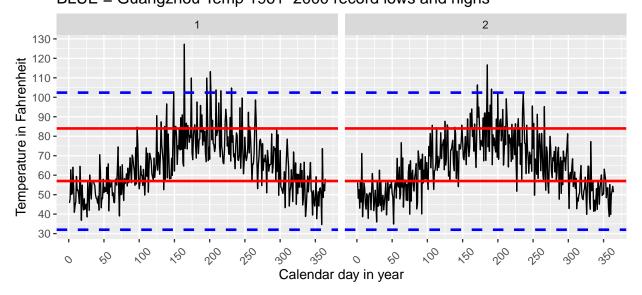


```
# df_fahrenheit <- ls_fahrenheit$df_fahrenheit
# kable(df_fahrenheit) %>% kable_styling_fc()
```

Second, we call the same temperature function, but we reduce shock persistence to 0.

```
# Test 2: Call function with defaults
ls_fahrenheit <- ffi_daily_temp_simulation(</pre>
 fl_mthly_mean_lowest = 57,
 fl_mthly_mean_highest = 84,
 fl_record_lowest = 32,
 fl_record_highest = 102.4,
  it_weeks_in_year = 52,
  it_days_in_week = 7,
 it_years = 2,
 fl_mu = 4.15,
 fl_sin_scaler = 0.25,
 fl_sigma_nv = 0.15,
 fl_rho_persist = 0.0,
  it_rand_seed = 123)
# Show
print(ls_fahrenheit$plt_fahrenheit)
```

Simulated Temperature for Guangzhou (Sine Wave + AR(1))
Each subplot is a different year
RED = Guangzhou Temp 1971–2000 lowest and highest monthly averages
BLUE = Guangzhou Temp 1961–2000 record lows and highs



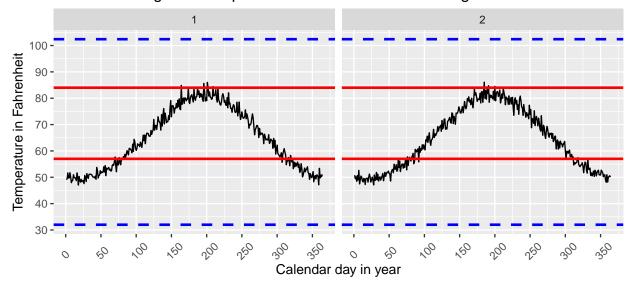
```
# df_fahrenheit <- ls_fahrenheit$df_fahrenheit
# kable(df_fahrenheit) %>% kable_styling_fc()
```

Third, we call the same temperature function, but we reduce shock persistence to 0 and shock variance to almost zero.

```
# Test 2: Call function with defaults
ls_fahrenheit <- ffi_daily_temp_simulation(</pre>
 fl_mthly_mean_lowest = 57,
 fl_mthly_mean_highest = 84,
 fl_record_lowest = 32,
 fl_record_highest = 102.4,
  it_weeks_in_year = 52,
  it_days_in_week = 7,
  it_years = 2,
 fl_mu = 4.15,
 fl_sin_scaler = 0.25,
 fl_sigma_nv = 0.025,
 fl_rho_persist = 0.0,
  it_rand_seed = 123)
# Show
print(ls_fahrenheit$plt_fahrenheit)
```

Simulated Temperature for Guangzhou (Sine Wave + AR(1)) Each subplot is a different year

RED = Guangzhou Temp 1971–2000 lowest and highest monthly averages BLUE = Guangzhou Temp 1961–2000 record lows and highs



df_fahrenheit <- ls_fahrenheit\$df_fahrenheit
kable(df_fahrenheit) %>% kable_styling_fc()

1.4 Distributions of Conception, Birth, Temperature and Pre-Term

We will simulate several datasets on birth and relate to temperature distributions. Below we do that, we go through several key points that will be confirmed/demonstrated by the simulations.

1.4.1 Conception and Birth Marginal and Joint Distributions

There are three relevant distributional concepts here:

- $P(C_i)$: From $ffi_concept_distribution_year()$ function generated earlier, this is the marginal discrete distribution of conception by calendar week of the year (perhaps some months are more popular for conception than others). C stands for conception.
- $P(B_i)$: From $ffi_gestation_age_at_birth_dist()$ function generated earlier, $P(B_i C_i)$ is the marginal discrete distribution of gestational age by birth by day/week of gestation. B stands for birth.
- $P(C_i, B_i C_i)$: the joint distribution of C_i and B_i. If they are unrelated, than at each conception day, the conditional distribution of gestational age at birth is identical. To allow for arbitrary correlation between conception and gestational-age, we can draw from some mixture of joint normal distribution with some set of variance-covariance matrixes, invert to the draws to quantiles along each dimension, and then map the quantiles to discrete outcomes along C and B dimensions.

1.4.2 Extreme Temperature and Pre-Term Births

Pre-term birth or not is determined jointly by the gestational age at birth and conception date distributions, which determine the conception and birth dates. We establish a common threshold below which a birth is considered pre-term.

Extreme temperature and whether a child is pre-term or not can be related.

Under **scenario** (A), conception dates are uniformly distribution across the year, and gestational age at birth is unrelated to conception time. Given this, extreme temperature or not has nothing to do with gestational

age at birth.

Under scenario (B), we can spuriously have pre-term experiencing less extreme-cold/heat when $P(C_i)$ is non-uniform/random, meaning $P(C_i) \neq \frac{1}{\text{CountTotalDatesOfConception}}$. This could go in several directions:

- 1. Under **scenario** (**B.1**), suppose all conception starts in Jan. and all extreme temp happen in Oct. Suppose pre-term means birth in Sep. rather than Nov.. If all extreme cold takes place in Oct. Then all pre-term births will not have experienced extreme cold. So the potential direction for spuriousness here is that pre-term will be correlated with less extreme temp. This is fixed by a conception month fixed effect.
- 2. Under scenario (B.2), suppose all conception starts in August. and all extreme temp happen in Oct. The number of extreme cold days will be the same under both pre-term and full-term birth groups, but the percentage of days exposed to extreme cold will be greater for the pre-term individuals, a spurious relationship again.
- 3. Under **Scenario** (**B.3**), we use the actual distribution of conception from Guangzhou China. Will the results be more similar to **scenarior** (**A**), where conception was uniformly distribution? Or more similar to **scenario B.1** with a greater concentration of conception earlier in the year, or **scenario B.2** with a greater concentration of conception later in the year?

Additionally, pre-term could experience more extreme-cold under **scenario** (C.1), where extreme temperature has a causal impact on gestational-age, causally leading to more pre-term. This is the research hypothesis of interest.

But pre-term could also experience more extreme-cold under **scenario** (C.2) for another reason. Suppose the distribution of high-pregnancy-risk parents/mothers across calendar month in each year is non-random: if all high risk mother get pregnant in August, and low risk mother get pregnant in March, we might end up with full-term births not experiencing extreme temperature, but pre-term births experiencing a lot of extreme-cold days. The pre-term births (likely with lower birthweight) is correlated with cold exposures, but this correlation due to the conception timing of high-risk pregnancies. In this situation, cold might not real effects on whether a birth is pre-term or not.

1.4.3 Days and Percent of Gestational Days Exposed to Cold

Under the various scenarios, the relationship between the *number* of days and *percentage* of conception days to pre-term birth of not will be different. They are:

- Under Scenario (A):
 - the share of days exposed to extreme cold will be identical for pre-term and full-term births.
 - the number of days exposed to extreme cold is less for pre-term birth compared to full-term.
- Under Scenario (B.1):
 - the share of days exposed to extreme cold will be less for pre-term and full-term births.
 - the number of days exposed to extreme cold is *less* for pre-term birth compared to full-term.
- Under Scenario (B.2):
 - the share of days exposed to extreme cold will be the same for pre-term and full-term births.
 - the number of days exposed to extreme cold is *more* for pre-term birth compared to full-term.
- Under Scenario (B.3):
 - Will this be like (A) or (B.1) or (B.3)? It turns out this is very similar to (A).
- Under Scenario (C.1) and (C.2):
 - the share of days exposed to extreme cold is *more* for pre-term birth compared to full-term.
 - the number of days exposed to extreme cold might generally be *less* although potentially *more* for pre-term and full-term births.

The share/percentage of days exposed to exposure cold is the appropriate measure to use. As seen in the scenarios above, it does not generate "spurious relationships" due to shorter time spans of pre-term births.

In the exercises below, we only simulate Scenarios (A) and (B) for now.

1.5 Simulating Datasets on Extreme Temperature Exposures Across Pregnancies

1.5.1 Simulate and Compute Extreme Temperature Exposure

We generate a dataframe with N individuals, each with conception day C_i , birth day B_i , and whether in the birth was full-term or not F_i . For simplicity, assume that there are 7 days per week, 4 weeks per month, and 13 months per year (to reach 52 weeks per year). This means there are 336 days of possible birth in a year.

Our gestational age distribution function can take any mean and standard deviation, we rescale the information mentioned prior from Right from the Start under the assumption of 336 days per year. (Plug in mean and sd as if there are 365 days into the function below, it will rescale that based on the actual number of days in the simulated year).

- Conception draws $P(C_i)$: For each of the N individuals, we first draw week of conception from the week of conception distribution function (adjust distribution depending on number of years and the starting month), then we draw the day of week of conception randomly, and the year of conception randomly as well.
- Gestation draws $P(B_i C_i)$: For each of the N individuals, we draw from the gestational week at the time of birth distribution the number of weeks of gestation, multiply this by seven and add a random number between 1 to 7.
- Joint distribution $P(C_i, B_i C_i)$: For the simulation below, the draws for day of birth is independent from the draws for gestational age at time of birth.

First, we create a function for generating birth conception and gestational distributions.

```
ffi_pop_concept_birth_simu <- function(</pre>
  it pop n = 50000,
  it_days_in_week = 7,
  it_weeks_in_month = 4,
  it months in year = 12,
  it years = 3,
  it_rng_seed = 123,
  fl_pre_term_ratio = 0.84,
  fl_peak_concept_frac_of_year_1st = 0.3,
  fl_peak_concept_frac_of_year_2nd = 0.9,
  fl_binom_1st_wgt = 0.15,
  fl_binom_2nd_wgt = 0.05,
  fl_mu_gabirth_days_365 = 276,
  fl_sd_gabirth_days_365 = 14,
  df_dist_conception_exo = NULL
) {
  # # 1. Define parameters
  # # 1.a Number of individuals of interest
  # it_pop_n <- 50000
  # # 1.b Number of days per week, week per month
  # # for simplicity, 7 days per week, 4 weeks per month, 12 months
  # it_days_in_week <- 7
  # it_weeks_in_month <- 4</pre>
  # it_months_in_year <- 12</pre>
  # it_years <- 3
  # # 1.c random draw seed
  # it_rnq_seed <- 123
  # # 1.d pre-term threshold
  # # what fraction of maximum pregnancy length is pre-term?
 # # Max length 44 weeks for example, 44*0.85 is about 37 months.
```

```
# fl_pre_term_ratio <- 0.84
# 1.e Other dates
it_weeks_in_year <- it_weeks_in_month*it_months_in_year</pre>
it days in month <- it days in week*it weeks in month
it_days_in_year <- it_days_in_week*it_weeks_in_month*it_months_in_year
\# 1.f Month of conception distribution
it_peak_wk_1st <- round(it_weeks_in_year*fl_peak_concept_frac_of_year_1st)</pre>
it peak wk 2nd <- round(it weeks in year*fl peak concept frac of year 2nd)
# fl_binom_1st_wqt <- 0.15
# fl_binom_2nd_wqt <- 0.05
# # 1.q Gestational age at birth distribution parameters
mu_gabirth_days <- round((fl_mu_gabirth_days_365/365)*it_days_in_year)</pre>
sd_gabirth_days <- round((fl_sd_gabirth_days_365/365)*it_days_in_year)</pre>
# 2. Date of conception random draws
# 2.a Week of conception distribution
ls_concept_fc <- ffi_concept_distribution_year(</pre>
 it_max_weeks = it_weeks_in_year,
 it_peak_wk_1st = it_peak_wk_1st, it_peak_wk_2nd = it_peak_wk_2nd,
 fl_binom_1st_wgt = fl_binom_1st_wgt, fl_binom_2nd_wgt = fl_binom_2nd_wgt,
  it runif seed = it rng seed*210,
  df_dist_conception_exo = df_dist_conception_exo)
df_dist_conception_week <- ls_concept_fc$df_dist_conception</pre>
# 2.b.1 Randomly (uniformly) drawing the day of birth
set.seed(it rng seed*221)
ar_draws_conception_day_of_week <- sample(</pre>
  it_days_in_week, it_pop_n, replace=TRUE)
# 2.b.2 Week of conception draws
set.seed(it_rng_seed*222)
ar_draws_conception_week <- sample(</pre>
  df_dist_conception_week$conception_calendar_week,
  it pop n,
  prob=df_dist_conception_week$conception_prob,
  replace=TRUE)
# 2.b.3 Randomly (uniformly) drawing the year of conception
set.seed(it rng seed*223)
ar draws conception year <- sample(
  it years, it pop n, replace=TRUE)
# 2.c Date of birth
ar_draws_concept_date <- (ar_draws_conception_year-1)*it_days_in_year +</pre>
  (ar_draws_conception_week-1)*it_days_in_week +
  ar_draws_conception_day_of_week
ar_draws_conception_day_of_year <- (ar_draws_conception_week-1)*it_days_in_week +
  ar_draws_conception_day_of_week
# 3. Gestational age at birth distribution simulation
# 3.a Gestational age distribution
ls_gsbirth_fc <- ffi_gestation_age_at_birth_dist(</pre>
 mu_gabirth_days = mu_gabirth_days, sd_gabirth_days = sd_gabirth days)
df_dist_gabirth <- ls_gsbirth_fc$df_dist_gabirth</pre>
# 3.b.1 Gestational day of week draws (random)
set.seed(it_rng_seed*321)
```

```
ar_draws_gsbirth_day_of_week <- sample(</pre>
  it_days_in_week, it_pop_n, replace=TRUE)
# 3.b.2 Gestational week draws
set.seed(it rng seed*322)
ar_draws_gsbirth_week <- sample(</pre>
 df_dist_gabirth$gabirth,
 it_pop_n,
 prob=df_dist_gabirth$prob,
 replace=TRUE)
# 3.c Gestational days at birth
ar_draws_gsbirth_day <- ar_draws_gsbirth_week*it_days_in_week + ar_draws_gsbirth_day_of_week
ar_draws_birth_date <- ar_draws_concept_date + ar_draws_gsbirth_day</pre>
# 4. Create dataframe
# 4.a Variables and labels
mt_birth_data <- cbind(</pre>
  ar_draws_concept_date, ar_draws_birth_date, ar_draws_gsbirth_day,
  ar_draws_conception_year, ar_draws_conception_week,
  ar_draws_conception_day_of_week, ar_draws_conception_day_of_year,
  ar_draws_gsbirth_week, ar_draws_gsbirth_day_of_week)
ar st varnames <- c('id',
  'survey_date_conception', 'survey_date_birth', 'gestation_length_in_days',
  'conception_year', 'conception_week',
  'conception_day_of_week', 'concept_day_of_year',
  'gestational_week_at_birth', 'gestational_day_of_week_at_birth')
# 4.b tibble with conception and birth data
df birth data <- as tibble(mt birth data) %>%
 rowid to column(var = "id") %>%
 rename_all(~c(ar_st_varnames)) %>%
 arrange(survey_date_conception, survey_date_birth)
# 4.c generate cut-off for preterm
it_pre_term_threshold <- round(fl_pre_term_ratio*length(df_dist_gabirth$gabirth)*it_days_in_week)
df_birth_data <- df_birth_data %>% mutate(
 preterm = case_when(it_pre_term_threshold >= gestation_length_in_days ~ 1,
                      TRUE \sim 0)
# 5. Display data
# Display
st_title <- pasteO('Day of year of conception and gestational age at birth\n',
  'pop=', it_pop_n, ', days-in-year=', it_days_in_year, ', seed=', it_rng_seed, '\n',
  'mean-ga-at-birth-in-month=', round(mu_gabirth_days/it_days_in_month, 3),
  ', sd-ga-at-birth-in-month=', round(sd_gabirth_days/it_days_in_month, 3))
plt concept birth <- df birth data %>%
 mutate(preterm = factor(preterm)) %>%
  ggplot(aes(x = concept_day_of_year, y=gestation_length_in_days, color=preterm)) +
 facet_wrap(~ conception_year) +
 geom_point() +
  labs(
   title = st_title,
   x = 'Calendar day in year',
   y = 'Gestational age in days at birth'
  scale_x_continuous(n.breaks = 12) +
```

```
scale_y_continuous(n.breaks = 10) +
theme(
    axis.text.x = element_text(angle = 45, vjust = 0.1, hjust = 0.1)
)

# print(plt_concept_birth)
# kable(df_birth_data) %>% kable_styling_fc_wide()
# plot(df_birth_data$survey_date_conception, df_birth_data$gestation_length_in_days)

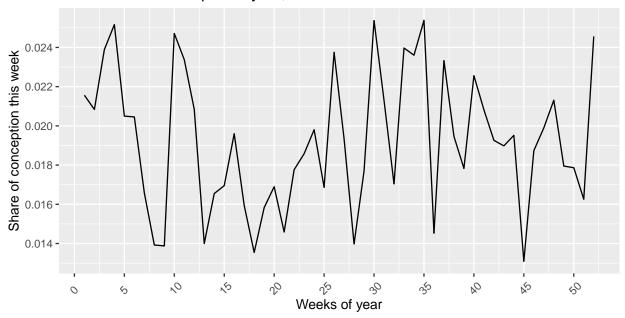
# Return
return(list(
    df_birth_data = df_birth_data,
    plt_concept_birth = plt_concept_birth,
    ls_concept_fc = ls_concept_fc,
    ls_gsbirth_fc = ls_gsbirth_fc
))
}
```

1.5.2 Scenario (A) Simulation, Uniform Conception

We generate a dataframe where the conception months are randomly/uniformly distributed, and there is no correlation between conception and gestation. This is **Scenario** (A).

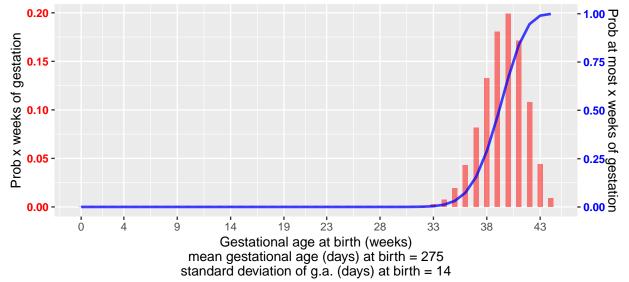
```
# Define some dates
it_days_in_week <- 7
it_weeks_in_month <- 4
it_months_in_year <- 13</pre>
it_years <- 3</pre>
# Simulate
ls_concept_birth <- ffi_pop_concept_birth_simu(</pre>
 it_pop_n = 10000,
 it_days_in_week = it_days_in_week,
 it_weeks_in_month = it_weeks_in_month,
 it_months_in_year = it_months_in_year,
 it_years = it_years,
  it_rng_seed = 999,
 fl_pre_term_ratio = 0.84,
 fl_peak_concept_frac_of_year_1st = 0.3,
 fl_peak_concept_frac_of_year_2nd = 0.9,
 fl_binom_1st_wgt = 0.00,
 fl_binom_2nd_wgt = 0.00,
 fl_mu_gabirth_days_365 = 276,
 fl_sd_gabirth_days_365 = 14
# Get dataframe and print distribution
df_birth_data_rand_cor0 <- ls_concept_birth$df_birth_data</pre>
print(ls_concept_birth$ls_concept_fc$plt_concept_week_of_year)
```

Distribution of conception week of birth over weeks of one specific year, seed=209790



print(ls_concept_birth\$ls_gsbirth_fc\$plt_dist_gabirth)

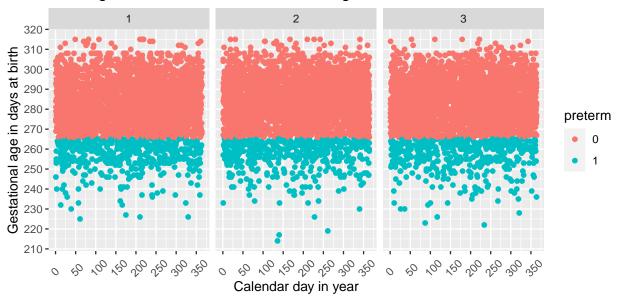
Gestational age at birth (weeks) Prob mass (Left) and cumulative prob (Right)



Assuming the binomial properties apply fl_binom_p = 0.898181818181818, it_binom_n = 44

print(ls_concept_birth\$plt_concept_birth)

Day of year of conception and gestational age at birth pop=10000, days-in-year=364, seed=999 mean-ga-at-birth-in-month=9.821, sd-ga-at-birth-in-month=0.5

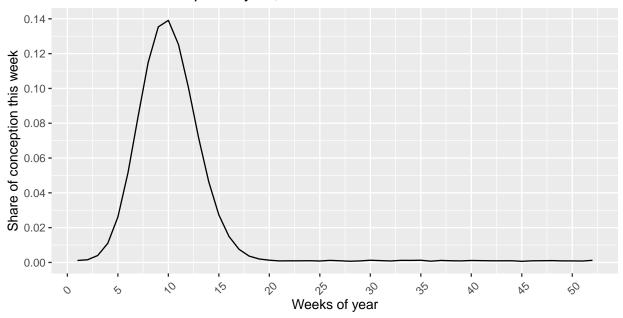


1.5.3 Scenario (B.1) Simulation, Nearly All Conceptions in Feb.

We generate a dataframe where the conception distribution has a high peak around Feb, and there is no correlation between conception and gestation. This is **Scenario** (B.1).

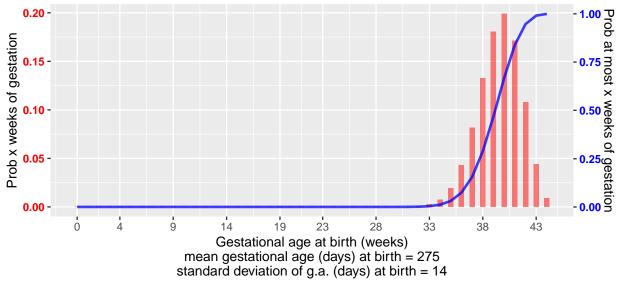
```
# Simulate
ls_concept_birth <- ffi_pop_concept_birth_simu(</pre>
  it_{pop_n} = 10000,
  it_days_in_week = it_days_in_week,
  it_weeks_in_month = it_weeks_in_month,
  it_months_in_year = it_months_in_year,
  it_years = it_years,
  it_rng_seed = 999,
  fl_pre_term_ratio = 0.84,
  fl_peak_concept_frac_of_year_1st = 0.2,
  fl_peak_concept_frac_of_year_2nd = 0.9,
  fl_binom_1st_wgt = 0.95,
 fl binom 2nd wgt = 0.00,
 fl_mu_gabirth_days_365 = 276,
  fl_sd_gabirth_days_365 = 14
# Get dataframe and print distribution
df_birth_data_CFeb_cor0 <- ls_concept_birth$df_birth_data</pre>
print(ls_concept_birth$ls_concept_fc$plt_concept_week_of_year)
```

Distribution of conception week of birth over weeks of one specific year, seed=209790



print(ls_concept_birth\$ls_gsbirth_fc\$plt_dist_gabirth)

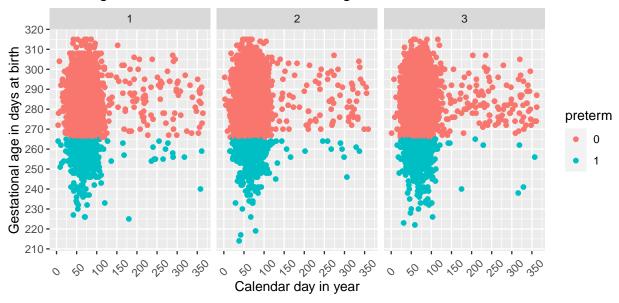
Gestational age at birth (weeks) Prob mass (Left) and cumulative prob (Right)



Assuming the binomial properties apply fl_binom_p = 0.898181818181818, it_binom_n = 44

print(ls_concept_birth\$plt_concept_birth)

Day of year of conception and gestational age at birth pop=10000, days-in-year=364, seed=999 mean-ga-at-birth-in-month=9.821, sd-ga-at-birth-in-month=0.5

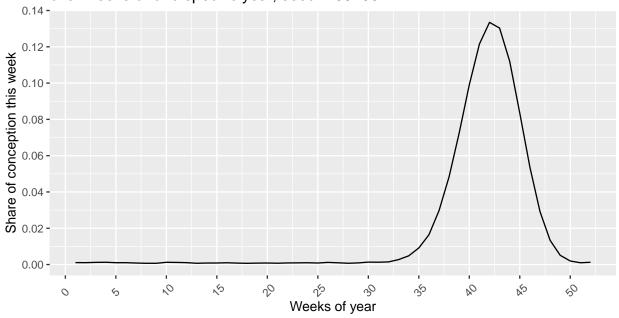


1.5.4 Scenario (B.2) Simulation, Nearly All Conceptions in Oct.

We generate a dataframe where the conception distribution has a high peak around October, and there is no correlation between conception and gestation. This is **Scenario** (B.2).

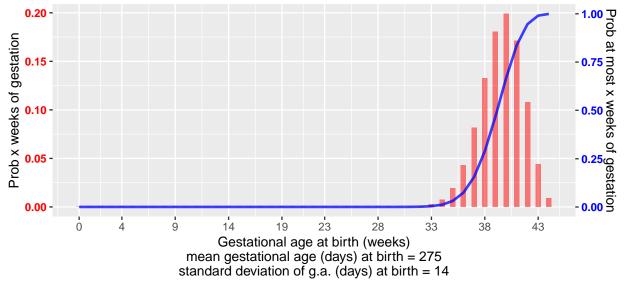
```
# Simulate
ls_concept_birth <- ffi_pop_concept_birth_simu(</pre>
  it_{pop_n} = 10000,
  it_days_in_week = it_days_in_week,
  it_weeks_in_month = it_weeks_in_month,
  it_months_in_year = it_months_in_year,
  it_years = it_years,
  it_rng_seed = 999,
  fl_pre_term_ratio = 0.84,
  fl_peak_concept_frac_of_year_1st = 0.2,
  fl_peak_concept_frac_of_year_2nd = 0.8,
  fl_binom_1st_wgt = 0.0,
 fl binom 2nd wgt = 0.95,
 fl_mu_gabirth_days_365 = 276,
  fl_sd_gabirth_days_365 = 14
# Get dataframe and print distribution
df_birth_data_COct_cor0 <- ls_concept_birth$df_birth_data</pre>
print(ls_concept_birth$ls_concept_fc$plt_concept_week_of_year)
```

Distribution of conception week of birth over weeks of one specific year, seed=209790



print(ls_concept_birth\$ls_gsbirth_fc\$plt_dist_gabirth)

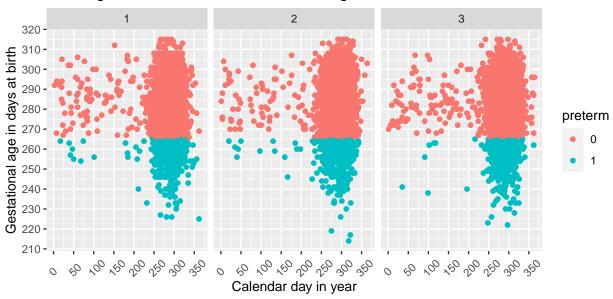
Gestational age at birth (weeks) Prob mass (Left) and cumulative prob (Right)



Assuming the binomial properties apply fl_binom_p = 0.898181818181818, it_binom_n = 44

print(ls_concept_birth\$plt_concept_birth)

Day of year of conception and gestational age at birth pop=10000, days-in-year=364, seed=999 mean-ga-at-birth-in-month=9.821, sd-ga-at-birth-in-month=0.5



1.5.5 Scenario (B.3) Simulation, Guangzhou Actual from Liu et al. Conception Distribution

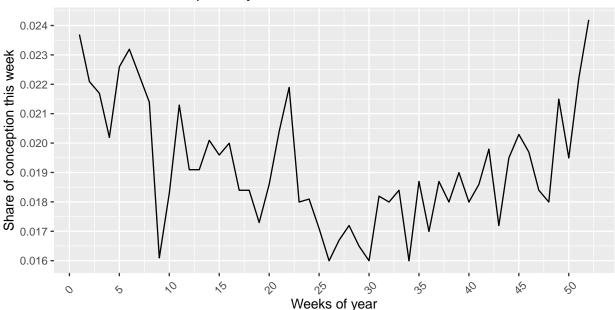
We generate a dataframe based on the actual conception distribution in Guangzhou, China from a particular survey year. Is this more similar to Scenarios **A, B.1 or B.2**? The data is from Same Environment, Stratified Impacts? Air Pollution, Extreme Temperatures, and Birth Weight in South China.

```
st_dist_conception_lbhwz_actual <- "conception_calendar_week, conception_prob
    1, 0.0237
       0.0221
    3,
       0.0217
        0.0202
        0.0226
    5,
    6,
        0.0232
        0.0223
    7,
       0.0214
    8,
       0.0161
    10, 0.0183
    11, 0.0213
    12, 0.0191
    13, 0.0191
    14, 0.0201
    15, 0.0196
    16, 0.0200
    17, 0.0184
    18, 0.0184
    19, 0.0173
    20, 0.0186
    21, 0.0204
    22, 0.0219
    23, 0.0180
    24, 0.0181
```

```
25, 0.0171
    26, 0.0160
    27, 0.0167
    28, 0.0172
    29, 0.0165
    30, 0.0160
    31, 0.0182
    32, 0.0180
    33, 0.0184
    34, 0.0160
    35, 0.0187
    36, 0.0170
    37, 0.0187
    38, 0.0180
    39, 0.0190
    40, 0.0180
    41, 0.0186
    42, 0.0198
    43, 0.0172
    44, 0.0195
    45, 0.0203
    46, 0.0197
    47, 0.0184
   48, 0.0180
    49, 0.0215
    50, 0.0195
    51, 0.0222
    52, 0.0242"
# Raw probability data to table
df_dist_conception_lbhwz_actual = read.csv(text=st_dist_conception_lbhwz_actual, header=TRUE)
ar_st_varnames <- c('conception_calendar_week',</pre>
                    'conception_prob')
tb_dist_conception_lbhwz_actual <- as_tibble(df_dist_conception_lbhwz_actual) %>%
  rename_all(~c(ar_st_varnames)) %>%
  mutate(conception_prob = conception_prob/sum(conception_prob))
# Summarize
summary(tb_dist_conception_lbhwz_actual)
## conception_calendar_week conception_prob
## Min.
          : 1.00
                           Min.
                                   :0.01600
## 1st Qu.:13.75
                            1st Qu.:0.01799
## Median :26.50
                            Median :0.01869
## Mean
          :26.50
                             Mean
                                   :0.01923
## 3rd Qu.:39.25
                             3rd Qu.:0.02032
## Max.
           :52.00
                             Max.
                                    :0.02419
# Check Probability sums to 1
sum(tb_dist_conception_lbhwz_actual$conception_prob)
## [1] 1
# Simulate
ls_concept_birth <- ffi_pop_concept_birth_simu(</pre>
it_pop_n = 10000,
```

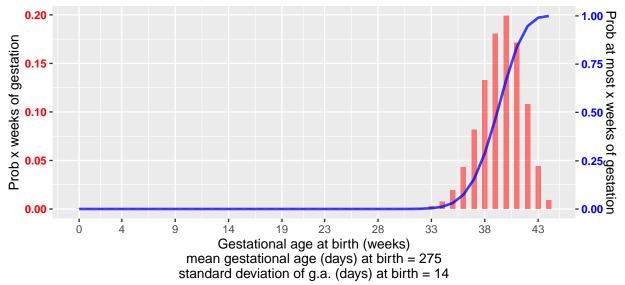
```
it_days_in_week = it_days_in_week,
  it_weeks_in_month = it_weeks_in_month,
  it_months_in_year = it_months_in_year,
  it_years = it_years,
  it rng seed = 999,
 fl_pre_term_ratio = 0.84,
 fl_peak_concept_frac_of_year_1st = 0.2,
 fl_peak_concept_frac_of_year_2nd = 0.9,
 fl_binom_1st_wgt = 0.95,
 fl_binom_2nd_wgt = 0.00,
 fl_mu_gabirth_days_365 = 276,
 fl_sd_gabirth_days_365 = 14,
 df_dist_conception_exo = tb_dist_conception_lbhwz_actual
\# Get dataframe and print distribution
df_birth_data_Clbhwz_cor0 <- ls_concept_birth$df_birth_data</pre>
print(ls_concept_birth$ls_concept_fc$plt_concept_week_of_year)
```

Distribution of conception week of birth over weeks of one specific year, seed=209790



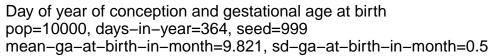
print(ls_concept_birth\$ls_gsbirth_fc\$plt_dist_gabirth)

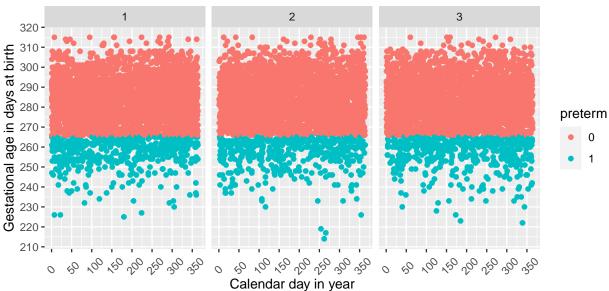
Gestational age at birth (weeks)
Prob mass (Left) and cumulative prob (Right)



Assuming the binomial properties apply fl_binom_p = 0.898181818181818, it_binom_n = 44

print(ls_concept_birth\$plt_concept_birth)





1.6 Compute Extreme Temperature Exposure

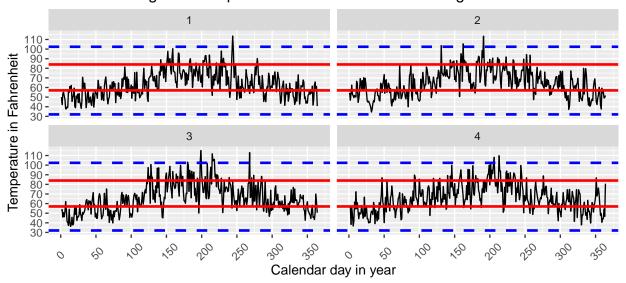
we simulate temperature corresponding to the span of data of interest that covers all individuals' potential course of pregnancy. We loop over each individual, and calculate the individual-specific extreme temperature exposure. Specifically, we store the percentage of days during pregnancy exposed to extreme cold or extreme

hot. We use as threshold 5 percent temperature extreme tails. We also store the number of days as well as the percent of gestational days that are exposed to extreme cold or hot.

First, we generate the temperature distribution along with extreme cold and hot days.

```
# Simulate the temperature distribution using just define parameters
ls_fahrenheit <- ffi_daily_temp_simulation(
  fl_mthly_mean_lowest = 57,
  fl_mthly_mean_highest = 84,
  fl_record_lowest = 32,
  fl_record_highest = 102.4,
  it_weeks_in_year = it_months_in_year*it_weeks_in_month,
  it_days_in_week = it_days_in_week,
  it_years = it_years+1,
  it_rand_seed = 999,
  st_extreme_cold_percentile = "p05",
  st_extreme_heat_percentile = "p95")
print(ls_fahrenheit$plt_fahrenheit)</pre>
```

Simulated Temperature for Guangzhou (Sine Wave + AR(1))
Each subplot is a different year
RED = Guangzhou Temp 1971–2000 lowest and highest monthly averages
BLUE = Guangzhou Temp 1961–2000 record lows and highs



```
df_fahrenheit <- ls_fahrenheit$df_fahrenheit
summary(df_fahrenheit$Fahrenheit)

## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 34.46 54.57 63.42 65.07 74.42 115.24

df_stats_fahrenheit <- REconTools::ff_summ_percentiles(df_fahrenheit, FALSE)</pre>
```

Second, for one individual, we test computing the number of days exposed to extreme cold.

```
# Define the extreme cold function
ffi_extreme_cold_percent_gestation <- function(df_fahrenheit, it_date_conception, it_date_birth){
    # get extreme cold</pre>
```

```
ar_extreme_cold <- df_fahrenheit %>%
    filter(survey_t >= it_date_conception & survey_t <= it_date_birth) %>% pull(extreme_cold)

# extreme cold days
it_extreme_cold_days <- sum(ar_extreme_cold)

return(it_extreme_cold_days)
}

# Test the function
it_extreme_cold_days <- ffi_extreme_cold_percent_gestation(df_fahrenheit, 11, 200)
print(it_extreme_cold_days)</pre>
```

[1] 9

Third, we create a function that takes the birth dataframe and temperature dataframe as inputs and generate extreme cold days exposure for each pregancy in the birth dateaframe using apply with an anonymous function

```
# Given two dataframes with birth data and temperature data, find cold exposure
ffi_birth_extreme_exposure <- function(df_birth_data, df_fahrenheit){</pre>
  # apply row by row, anonymous function
  # see: https://fanwangecon.github.io/R4Econ/function/noloop/htmlpdfr/fs_apply.html#122_anonymous_func
  mt_birth_cold <- apply(df_birth_data, 1, function(row) {</pre>
      id <- row[1]
      it_date_conception <- row[2]</pre>
      it_date_birth <- row[3]</pre>
      it_week_of_year_conception <- row[6]</pre>
      it_preterm <- row[11]</pre>
      it_extreme_cold_days <- ffi_extreme_cold_percent_gestation(</pre>
        df_fahrenheit, it_date_conception, it_date_birth)
      mt_all_res <- cbind(id, it_date_conception, it_week_of_year_conception,</pre>
                           it_date_birth,
                           it_preterm, it_extreme_cold_days)
      return(mt_all_res)
    })
  # Column Names
  ar st varnames <- c('id',
                       'survey_date_conception', 'week_of_year_conception',
                       'survey_date_birth',
                       'preterm', 'days_extreme_cold')
  # Combine to tibble, add name col1, col2, etc.
  tb_birth_cold <- as_tibble(t(mt_birth_cold)) %>%
    rename_all(~c(ar_st_varnames)) %>%
    mutate(days_extreme_cold_percent = days_extreme_cold/(survey_date_birth-survey_date_conception)) %>
    mutate(month_of_year_conception = round(week_of_year_conception/it_weeks_in_month))
  # Show Results
  # kable(tb_birth_cold[1:20,]) %>% kable_styling_fc()
  return(tb_birth_cold)
```

```
}
```

Fourth, we generate cold exposures for the datasets we generated using the same temperature dataframe by different birth/conception dataframes created under different assumptions. We use the <u>ffi_birth_extreme_exposure</u> function just created.

```
# Scenario (A)
tb_birth_cold_rand_cor0 <- ffi_birth_extreme_exposure(df_birth_data_rand_cor0, df_fahrenheit)
# Scenario (B.1)
tb_birth_cold_CFeb_cor0 <- ffi_birth_extreme_exposure(df_birth_data_CFeb_cor0, df_fahrenheit)
# Scenario (B.2)
tb_birth_cold_COct_cor0 <- ffi_birth_extreme_exposure(df_birth_data_COct_cor0, df_fahrenheit)
# Scenario (B.3)
tb_birth_cold_Clbhwz_cor0 <- ffi_birth_extreme_exposure(df_birth_data_Clbhwz_cor0, df_fahrenheit)</pre>
```

1.6.1 Regression pre-term Births and Extreme Temperature Exposures

In the four datasets that we have generated, each under a different set of conception date distribution assumptions, we have information on the binary prer-term outcome and extreme cold exposure for each individual. We now regress pre-term on extreme cold exposures.

In the regression under **Scenarior** (A), percent of extreme cold days is unrelated to pre-term.

```
# Scenarior (A) regression based on percent of days, with month of year conception fixed effects
re_esti_cold_rand_cor0 <- lm(preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
  data=tb birth cold rand cor0)
summary(re_esti_cold_rand_cor0)
##
## Call:
  lm(formula = preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
##
       data = tb_birth_cold_rand_cor0)
##
## Residuals:
##
      Min
                1Q Median
                                30
                                       Max
  -0.1833 -0.1525 -0.1451 -0.1375 0.8941
##
## Coefficients:
##
                                       Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                       0.151680
                                                  0.021404
                                                             7.086 1.47e-12 ***
                                                                     0.7931
## days_extreme_cold_percent
                                       0.082741
                                                  0.315434
                                                             0.262
## factor(month_of_year_conception)1
                                      -0.007843
                                                  0.022386 -0.350
                                                                     0.7261
## factor(month_of_year_conception)2
                                       0.001869
                                                  0.022307
                                                             0.084
                                                                     0.9332
## factor(month_of_year_conception)3
                                       0.026794
                                                  0.023217
                                                                     0.2485
                                                             1.154
## factor(month_of_year_conception)4
                                      -0.009478
                                                  0.021501
                                                           -0.441
                                                                     0.6593
## factor(month_of_year_conception)5
                                                                     0.8868
                                      -0.003390
                                                  0.023813 -0.142
## factor(month_of_year_conception)6
                                      -0.010144
                                                  0.021580
                                                           -0.470
                                                                     0.6383
## factor(month_of_year_conception)7
                                      -0.002499
                                                  0.024223
                                                           -0.103
                                                                     0.9178
## factor(month_of_year_conception)8
                                      -0.013379
                                                  0.021352
                                                            -0.627
                                                                     0.5309
## factor(month_of_year_conception)9
                                      -0.003886
                                                  0.023437
                                                            -0.166
                                                                     0.8683
## factor(month_of_year_conception)10 -0.012146
                                                  0.021819 -0.557
                                                                     0.5778
## factor(month_of_year_conception)11 -0.018703
                                                  0.024094
                                                            -0.776
                                                                     0.4376
## factor(month_of_year_conception)12 -0.049246
                                                            -2.304
                                                                     0.0212 *
                                                  0.021375
## factor(month_of_year_conception)13 -0.011213
                                                  0.024470 -0.458
                                                                     0.6468
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

```
##
## Residual standard error: 0.3528 on 9985 degrees of freedom
## Multiple R-squared: 0.001927, Adjusted R-squared: 0.0005276
## F-statistic: 1.377 on 14 and 9985 DF, p-value: 0.1548
```

In the regression under **Scenarior** (**B.1**), even controlling for conception month fixed effects, those experiencing extreme cold are associated with full-term birth. Since the outcome is equal to 1 if preterm happens, the coefficient for extreme-code below is negative. More extreme cold seems to make pre-term less likely, this is a spurious relationship due the conception been concentrated around Feburary. (Spurious because gestational age at birth is i.i.d. draw for all individual birth.)

```
# Scenarior (B.1) regression based on percent of days, with month of year conception fixed effects
re_esti_cold_CFeb_cor0 <- lm(preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
  data=tb_birth_cold_CFeb_cor0)
summary(re esti cold CFeb cor0)
##
## Call:
  lm(formula = preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
       data = tb_birth_cold_CFeb_cor0)
##
##
##
  Residuals:
                       Median
                                    3Q
##
                  1Q
   -0.47445 -0.17960 -0.13325 -0.02082
##
##
## Coefficients:
                                       Estimate Std. Error t value Pr(>|t|)
##
                                                             8.946 < 2e-16 ***
## (Intercept)
                                        0.67722
                                                   0.07570
## days_extreme_cold_percent
                                      -12.66697
                                                   0.36116 -35.073 < 2e-16 ***
## factor(month_of_year_conception)1
                                                   0.07639 -3.139 0.001702 **
                                       -0.23975
## factor(month_of_year_conception)2
                                       -0.31858
                                                   0.07498
                                                            -4.249 2.17e-05 ***
## factor(month_of_year_conception)3
                                                   0.07496
                                                            -3.440 0.000583 ***
                                       -0.25788
## factor(month_of_year_conception)4
                                       -0.10998
                                                   0.07521
                                                            -1.462 0.143698
## factor(month_of_year_conception)5
                                        0.17635
                                                   0.09135
                                                             1.931 0.053561 .
## factor(month_of_year_conception)6
                                        0.18984
                                                   0.09010
                                                             2.107 0.035152 *
## factor(month_of_year_conception)7
                                        0.06726
                                                   0.09927
                                                             0.678 0.498076
## factor(month_of_year_conception)8
                                        0.12532
                                                   0.08690
                                                              1.442 0.149310
## factor(month_of_year_conception)9
                                        0.28153
                                                   0.09647
                                                             2.918 0.003529 **
## factor(month_of_year_conception)10
                                        0.26670
                                                   0.08831
                                                             3.020 0.002533 **
## factor(month of year conception)11
                                        0.17190
                                                   0.10026
                                                             1.715 0.086462 .
## factor(month_of_year_conception)12
                                        0.15399
                                                   0.08724
                                                             1.765 0.077585 .
## factor(month_of_year_conception)13
                                       -0.03690
                                                   0.10290 -0.359 0.719887
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.333 on 9985 degrees of freedom
## Multiple R-squared: 0.1108, Adjusted R-squared: 0.1095
## F-statistic: 88.83 on 14 and 9985 DF, p-value: < 2.2e-16
```

In the regression under **Scenarior** (**B.2**), even controlling for conception month fixed effects, those experiencing extreme cold are associated with pre-term birth. Since the outcome is equal to 1 if preterm happens, the coefficient for extreme-code below is positive. More extreme cold seems to make pre-term more likely, this is a spurious relationship due the conception been concentrated around October. (Spurious because gestational age at birth is i.i.d. draw for all individual birth.)

```
# Scenarior (B.2) regression based on percent of days, with month of year conception fixed effects
re_esti_cold_COct_cor0 <- lm(preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
  data=tb_birth_cold_COct_cor0)
summary(re_esti_cold_COct_cor0)
##
## Call:
## lm(formula = preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
       data = tb_birth_cold_COct_cor0)
##
## Residuals:
                1Q Median
##
      Min
                                       Max
## -0.2987 -0.1992 -0.1118 -0.0833 1.0466
##
## Coefficients:
##
                                        Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                      -0.1840200 0.0827025
                                                             -2.225 0.02610 *
                                                             17.136
## days_extreme_cold_percent
                                       4.8908282
                                                  0.2854113
                                                                    < 2e-16 ***
## factor(month_of_year_conception)1
                                       0.1729513 0.1000501
                                                              1.729 0.08390
## factor(month_of_year_conception)2
                                       0.2536674
                                                  0.0951003
                                                              2.667 0.00766 **
## factor(month_of_year_conception)3
                                       0.1208876
                                                  0.1009776
                                                              1.197 0.23127
## factor(month_of_year_conception)4
                                       0.1373929
                                                  0.0973410
                                                              1.411 0.15814
                                                              0.009 0.99249
## factor(month_of_year_conception)5
                                       0.0009896 0.1051860
## factor(month_of_year_conception)6
                                      -0.0128848 0.0988770
                                                             -0.130 0.89632
## factor(month_of_year_conception)7
                                       0.0900028
                                                  0.1053556
                                                              0.854 0.39297
## factor(month_of_year_conception)8
                                                              0.412 0.68068
                                       0.0368224 0.0894734
## factor(month_of_year_conception)9
                                       0.0395859
                                                  0.0835103
                                                              0.474 0.63549
## factor(month_of_year_conception)10
                                       0.0287383
                                                  0.0824476
                                                              0.349 0.72742
## factor(month_of_year_conception)11
                                       0.0280835
                                                              0.340 0.73358
                                                  0.0825082
## factor(month_of_year_conception)12
                                                              0.650 0.51551
                                       0.0538927
                                                  0.0828729
## factor(month_of_year_conception)13
                                                              1.257 0.20870
                                       0.1479429
                                                  0.1176737
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
\#\# Residual standard error: 0.3479 on 9985 degrees of freedom
## Multiple R-squared: 0.02959,
                                    Adjusted R-squared: 0.02823
## F-statistic: 21.75 on 14 and 9985 DF, p-value: < 2.2e-16
In the regression under Scenarior (B.3), we have a very weak negative and insignificant correlation between
```

In the regression under **Scenarior** (B.3), we have a very weak negative and insignificant correlation between preterm and extreme-cold exposure in percentage of days. Given our empirical conception distribution, our result is more similar to **Scenarior A** and not close to **Scenarior B.1** and **Scenarior B.2**. This means that if indeed gestational age at birth are i.i.d. draws and have nothing to do with cold exposures, our empirical conception distribution will not generate a spurious correlation between extreme-cold exposure and pre-term birth (and lower birth-weight).

```
# Scenarior (B.3) regression based on percent of days, with month of year conception fixed
re_esti_cold_Clbhwz_cor0 <- lm(preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
    data=tb_birth_cold_Clbhwz_cor0)

##
## Call:
## Im(formula = preterm ~ days_extreme_cold_percent + factor(month_of_year_conception),
##
    data = tb_birth_cold_Clbhwz_cor0)
##</pre>
```

```
## Residuals:
##
      Min
                1Q Median
                                30
                                       Max
## -0.1764 -0.1524 -0.1435 -0.1331 0.8863
##
## Coefficients:
##
                                       Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                       0.156453
                                                  0.020274
                                                             7.717 1.31e-14 ***
## days_extreme_cold_percent
                                      -0.559244
                                                  0.312876 - 1.787
                                                                     0.0739 .
## factor(month_of_year_conception)1
                                       0.007591
                                                  0.021583
                                                             0.352
                                                                     0.7251
## factor(month_of_year_conception)2
                                      -0.003988
                                                  0.020844 -0.191
                                                                     0.8483
## factor(month_of_year_conception)3
                                      -0.007072
                                                  0.022648 -0.312
                                                                     0.7549
## factor(month_of_year_conception)4
                                       0.024037
                                                  0.019699
                                                             1.220
                                                                     0.2224
## factor(month_of_year_conception)5
                                       0.016194
                                                  0.022347
                                                             0.725
                                                                     0.4687
## factor(month_of_year_conception)6
                                       0.008069
                                                  0.020596
                                                             0.392
                                                                     0.6952
                                       0.037003
## factor(month_of_year_conception)7
                                                  0.023068
                                                             1.604
                                                                     0.1087
## factor(month_of_year_conception)8
                                       0.008074
                                                  0.021107
                                                             0.383
                                                                     0.7021
## factor(month_of_year_conception)9
                                       0.028895
                                                  0.023264
                                                             1.242
                                                                     0.2142
## factor(month_of_year_conception)10
                                       0.024185
                                                  0.020911
                                                                     0.2475
                                                             1.157
## factor(month_of_year_conception)11
                                       0.036997
                                                  0.022806
                                                             1.622
                                                                     0.1048
## factor(month_of_year_conception)12
                                       0.029644
                                                  0.020268
                                                             1.463
                                                                     0.1436
## factor(month_of_year_conception)13  0.007813
                                                  0.023647
                                                             0.330
                                                                     0.7411
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.353 on 9985 degrees of freedom
## Multiple R-squared: 0.001145,
                                    Adjusted R-squared: -0.0002554
## F-statistic: 0.8176 on 14 and 9985 DF, p-value: 0.6506
```

1.7 Visualize Pre-term, Full-term and Days of Extreme Temperature Exposures

1.7.1 Analysis and Visualization Functions

Using the data generated prior, we generate distribution of days exposed to extreme cold temperature.

First, create a function for generating statistics and visualization for the relationship between extreme cold days and pre-term or not.

```
# Create a function.
ffi_cold_days_preterm_analyze <- function(</pre>
  tb_birth_cold,
  st_title = pasteO('Scenario (A), Extreme cold DAYS Distribution\n',
                     'Uniform Conception\n',
                     'Conception and Birth uncorrelated')){
  # summarize
  str_stats_group <- 'allperc'</pre>
  ar_perc \leftarrow c(0.05, 0.25, 0.5, 0.75, 0.95)
  # For tb birth cold
  ls_summ_by_group <- REconTools::ff_summ_bygroup(</pre>
    tb_birth_cold, c('preterm'),
    'days_extreme_cold', str_stats_group, ar_perc)
  df_table_grp_stats_rand_cor0 <- ls_summ_by_group$df_table_grp_stats
  print(df table grp stats rand cor0)
  # Visualize
 plt_rand_cor0_level <- tb_birth_cold %>%
```

```
mutate(preterm = factor(preterm)) %>%
    group_by(preterm) %% mutate(days_extreme_cold_mean = mean(days_extreme_cold)) %>% ungroup() %>%
   ggplot(aes(x=days_extreme_cold, color=preterm)) +
   geom_density() +
    geom_vline(aes(xintercept=days_extreme_cold_mean, color=preterm), linetype="dashed") +
    labs(
      title = st_title,
     x = 'Days exposed to extreme cold',
     y = 'Density'
    scale_x_continuous(n.breaks = 10) +
   scale_y_continuous(n.breaks = 10) +
   theme(
      axis.text.x = element_text(angle = 45, vjust = 0.1, hjust = 0.1)
    )
  return(list(
   df_temp_preterm_stats = df_table_grp_stats_rand_cor0,
   plt_temp_preterm = plt_rand_cor0_level))
}
```

Second, create a function for generating statistics and visualization for the relationship between extreme cold percent of gestational days and pre-term or not.

```
ffi_cold_percent_days_preterm_analyze <- function(</pre>
  tb_birth_cold,
  st_title = pasteO('Scenario (A), Extreme cold PERCENT of DAYS Distribution\n',
                     'Uniform Conception\n',
                     'Conception and Birth uncorrelated')){
  # summarize
  str_stats_group <- 'allperc'</pre>
  ar_perc \leftarrow c(0.05, 0.25, 0.5, 0.75, 0.95)
  # For tb birth cold rand cor0
  ls_summ_by_group <- REconTools::ff_summ_bygroup(</pre>
    tb birth cold, c('preterm'),
    'days_extreme_cold_percent', str_stats_group, ar_perc)
  df_table_grp_stats_rand_cor0 <- ls_summ_by_group$df_table_grp_stats
  # Visualize
  plt_rand_cor0_level <- tb_birth_cold %>%
    mutate(preterm = factor(preterm)) %>%
    group_by(preterm) %>% mutate(days_extreme_cold_percent_mean = mean(days_extreme_cold_percent)) %>% "
    ggplot(aes(x=days_extreme_cold_percent, color=preterm)) +
    geom_density() +
    geom_vline(aes(xintercept=days_extreme_cold_percent_mean, color=preterm), linetype="dashed") +
    labs(
      title = st_title,
      x = 'Percent of gestational days exposed to extreme cold',
      y = 'Density'
    scale_x_continuous(n.breaks = 10) +
    scale y continuous (n.breaks = 10) +
    theme(
```

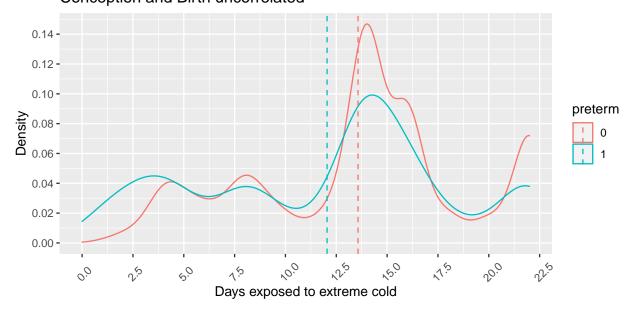
```
axis.text.x = element_text(angle = 45, vjust = 0.1, hjust = 0.1)
)

return(list(
    df_temp_preterm_stats = df_table_grp_stats_rand_cor0,
    plt_temp_preterm = plt_rand_cor0_level))
}
```

1.7.2 Scenario A, number of days and preterm

Scenario A, number of days exposed to extreme cold distribution:

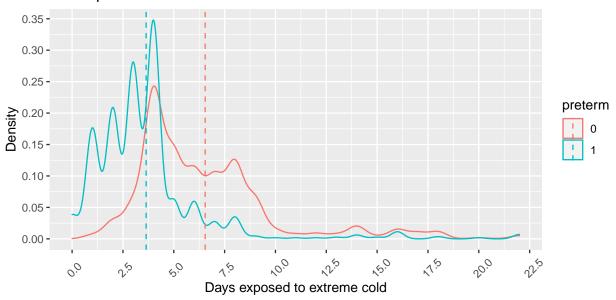
Scenario (A), Extreme cold DAYS Distribution (dashed lines are means) Uniform Conception Conception and Birth uncorrelated



1.7.3 Scenario B.1, number of days and preterm

Scenario B.1, number of days exposed to extreme cold distribution:

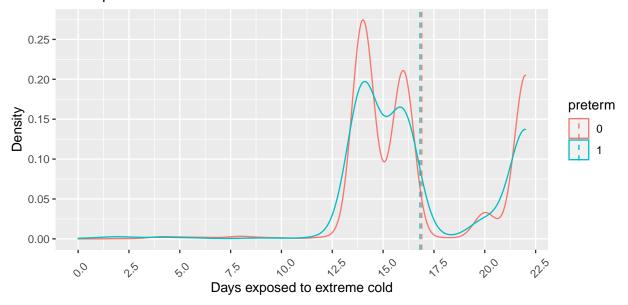
Scenario (B.1), Extreme cold DAYS Distribution (dashed lines are means) Conception Concentrated around Feb Conception and Birth uncorrelated



1.7.4 Scenario B.2, number of days and preterm

Scenario B.2, number of days exposed to extreme cold distribution (average lines largely overlap):

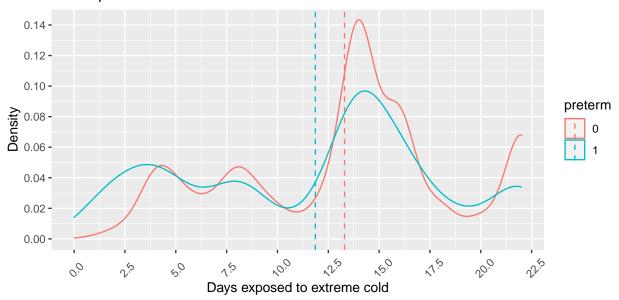
Scenario (B.2), Extreme cold DAYS Distribution (dashed lines are means) Conception Concentrated around Oct Conception and Birth uncorrelated



1.7.5 Scenario B.3, number of days and preterm

Scenario B.3, number of days exposed to extreme cold distribution (average lines largely overlap), note this is the result based on the empirical conception distribution from Guangzhou:

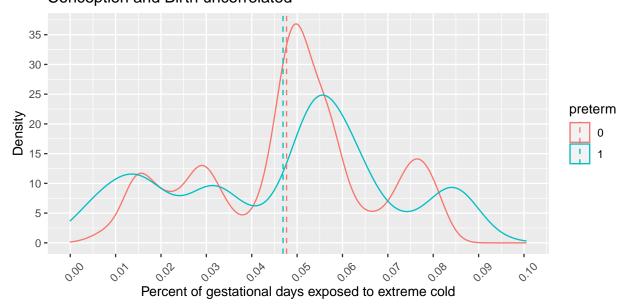
Scenario (B.3), Extreme cold DAYS Distribution (dashed lines are means) Conception Empirical Guangzhou Distribution Conception and Birth uncorrelated



1.7.6 Scenario A, percent of days and preterm

Scenario A, percent of days exposed to extreme cold distribution:

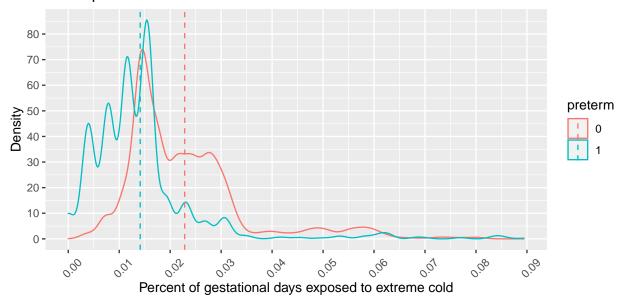
Scenario (A), Extreme cold PERCENT of DAYS Distribution (dashed = means) Uniform Conception Conception and Birth uncorrelated



1.7.7 Scenario B.1, percent of days and preterm

Scenario B.1, percent of days exposed to extreme cold distribution:

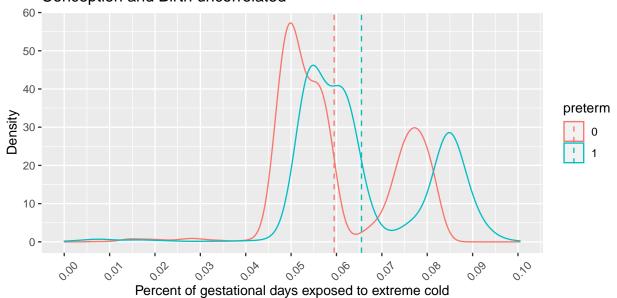
Scenario (B.1), Extreme cold PERCENT of DAYS Distribution (dashed = means) Conception Concentrated around Feb Conception and Birth uncorrelated



1.7.8 Scenario B.2, percent of days and preterm

Fourth, Scenario B.2, percent of days exposed to extreme cold distribution:

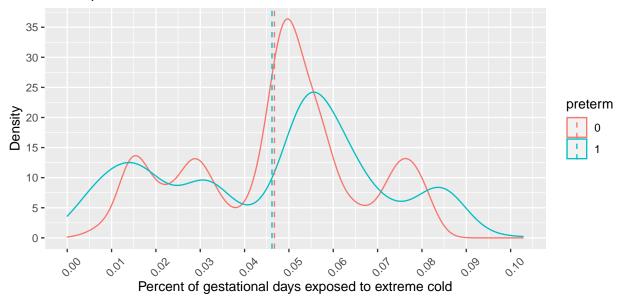
Scenario (B.2), Extreme cold PERCENT of DAYS Distribution (dashed = means) Conception Concentrated around Oct Conception and Birth uncorrelated



1.7.9 Scenario B.3, percent of days and preterm

Scenario B.3, percent of days exposed to extreme cold distribution (average lines largely overlap), note this is the result based on the empirical conception distribution from Guangzhou:

Scenario (B.3), Extreme cold PERCENT of DAYS Distribution (dashed = means) Conception Empirical Guangzhou Distribution Conception and Birth uncorrelated



conception_calendar_week	conception_prob
conception_carendar_week	0.0127130
2	0.0127130
$\frac{2}{3}$	0.0170371
4	0.0139137
5	0.0192192
6	0.0105366
7	0.0157063
8	0.0201894
9	0.0184492
10	0.0200423
11	0.0283162
12	0.0270358
13	0.0325643
14	0.0336904
15	0.0295436
16	0.0361683
17	0.0271255
18	0.0218356
19	0.0213639
20	0.0247214
21	0.0219399
22	0.0185480
23	0.0171439
24	0.0201420
25	0.0165473
26	0.0169517
27	0.0152796
28	0.0157608
29	0.0127605
30	0.0114004
31	0.0195570
32	0.0191672
33	0.0174747
34	0.0191898
35	0.0126323
36	0.0185624
37	0.0230986
38	0.0195683
39	0.0220649
40	0.0219644
41	0.0208423
42	0.0223014
43	0.0204080
44	0.0179026
45	0.0139712
46	0.0125547
47	0.0127300
48	0.0146663
49	0.0125517
50	0.0183537
51	0.0103274
52	0.0142397