# Supplement 2

## Sites of pain

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## **Contents**

Import and check data	2
Demographics	3 4 4
Proportion point estimates with 95% Cls  Process data	7
By sex Process data	16
By age Process data	25
Process data	
Logistic regression (generalised linear mixed model)  Process data	72 73
Session information 7	76
Note: Sites RBP and RISI removed at data cleaning stage because n < 10.	

### Import and check data

```
# Import
data <- read rds('data-cleaned/data-pain-sites.rds')</pre>
demo <- read_rds('data-cleaned/data-demographics.rds')</pre>
# Check
## Pain sites
dim(data)
## [1] 595
names (data)
## [1] "ID"
                                                        "Head"
                                                                                                "Throat"
                                                        "Arms"
                                                                                                "Elbows"
## [4] "Shoulder"
## [7] "Wrists.Hands"
                                                        "Chest"
                                                                                                "Upper_back"
## [10] "Lower_back"
                                                                                                "Cervical_spine"
                                                        "Abdomen"
## [13] "Thoracic_spine"
                                                        "Lumbosacral_spine"
                                                                                               "Groin"
## [16] "Hips"
                                                        "Legs"
                                                                                                "Knees"
## [19] "Ankles.Feet"
                                                        "Buttocks"
glimpse(data)
## Rows: 595
## Columns: 20
## $ ID
                                             <chr> "RPB73", "RPB74", "RPB75", "RPB76", "RPB77", "RPB...
                                              <chr> "No", "No", "No", "Yes", "Yes", "No", "No", "No", ...
## $ Head
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Throat
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Shoulder
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Arms
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Elbows
## $ Wrists.Hands
                                          <chr> "No", "No", "No", "No", "No", "No", "No", "No", "...
                                             <chr> "No", "No", "No", "Yes", "No", "No", "No", "No", ...
## $ Chest
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Upper_back
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Lower back
                                             <chr> "No", "No", "Yes", "Yes", "No", "No", "Yes", "No"...
## $ Abdomen
## $ Cervical_spine
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Thoracic_spine
                                              <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Lumbosacral_spine <chr> "No", "
                                              <chr> "No", "No", "No", "No", "No", "No", "No", "No", "No", "...
## $ Groin
                                             <chr> "No", "No", "No", "No", "No", "No", "No", "Yes", ...
## $ Hips
                                              <chr> "No", "No", "No", "No", "Yes", "No", "Yes",...
## $ Legs
                                              <chr> "No", "No", "No", "No", "Yes", "No", "Yes",...
## $ Knees
## $ Ankles.Feet
                                              <chr> "No", "No", "No", "No", "Yes", "No", "Yes",...
                                              <chr> "No", "No", "No", "No", "No", "No", "No", "Yes", ...
## $ Buttocks
## Demographics
dim(demo)
## [1] 595
names (demo)
## [1] "ID"
                                                      "Sex"
                                                                                              "Age"
## [4] "Employment_status" "CD4_recent"
                                                                                              "ART_currently"
## [7] "Education"
```

#### 

### **Basic descriptive statistics**

#### Pain sites

```
data %>%
    select(-ID) %>%
    mutate_if(is.character, factor) %>%
    skim()
```

Table 1: Data summary

Name	Piped data
Number of rows	595
Number of columns	19
Column type frequency:	_
Column type frequency:	10
factor	19
Group variables	 None
<b>-</b>	

#### Variable type: factor

skim_variable	n_missing	complete_rate	n_unique	top_counts
Head	0	1	2	No: 401, Yes: 194
Throat	0	1	2	No: 576, Yes: 19
Shoulder	0	1	2	No: 547, Yes: 48
Arms	0	1	2	No: 570, Yes: 25
Elbows	0	1	2	No: 573, Yes: 22
Wrists.Hands	0	1	2	No: 560, Yes: 35
Chest	0	1	2	No: 479, Yes: 116
Upper_back	0	1	1	No: 595
Lower back	0	1	2	No: 552, Yes: 43
Abdomen	0	1	2	No: 435, Yes: 160
Cervical_spine	0	1	2	No: 566, Yes: 29
Thoracic_spine	0	1	2	No: 522, Yes: 73
Lumbosacral_spine	0	1	2	No: 505, Yes: 90

skim_variable	n_missing	complete_rate	n_unique	top_counts
Groin	0	1	2	No: 541, Yes: 54
Hips	0	1	2	No: 556, Yes: 39
Legs	0	1	2	No: 497, Yes: 98
Knees	0	1	2	No: 512, Yes: 83
Ankles.Feet	0	1	2	No: 412, Yes: 183
Buttocks	0	1	2	No: 576, Yes: 19

## **Demographics**

```
demo %>%
   select(-ID) %>%
   mutate_if(is.character, factor) %>%
   skim()
```

Table 3: Data summary

Name	Piped data
Number of rows	595
Number of columns	6
	_
Column type frequency:	
factor	4
numeric	2
Group variables	None

#### Variable type: factor

skim_variable	n_missing	complete_rate	n_unique	top_counts
Sex	0	1.00	2	Fem: 479, Mal: 116
Employment_status	49	0.92	4	Une: 329, Ful: 131, Par: 52, Oth: 34
ART_currently	5	0.99	2	Yes: 459, No: 131
Education	37	0.94	3	Sec: 393, Pri: 100, Ter: 65

#### Variable type: numeric

skim_variable	n_missing	complete_rate	mean	sd	p0	p25	p50	p75	p100
Age	8	0.99	37.31	9.06	19	31	36.0	42.00	76
CD4_recent	99	0.83	337.17	400.32	1	155	261.5	437.75	7300

### **Boostrap functions**

```
# Proportion
prop_func <- function(d, i){</pre>
```

```
dat <- d[i, ]
  dat_vec <- dat[[1]]
  dat_prop <- mean(dat_vec == 'Yes')
  dat_prop
}</pre>
```

### **Proportion point estimates with 95% CIs**

#### **Process data**

```
# Set seed
set.seed(2020)
# Remove ID and upper_back (only one outcome -- no pain) columns
prop <- data[, !(names(data) %in% c('ID', 'Upper_back'))]</pre>
# Bootstrap CIs
prop_boot <- prop %>%
    # Pivot to long format
   pivot_longer(cols = everything(),
                 names_to = 'site',
                 values_to = 'pain_present') %>%
    # Add body regions
   mutate(region = case_when(
        site == 'Chest' |
            site == 'Head' |
            site == 'Throat' |
            site == 'Shoulder' ~ 'Head and upper torso',
        site == 'Lower_back' |
           site == 'Abdomen' |
           site == 'Hips' |
            site == 'Buttocks' |
            site == 'Groin' ~ 'Lower torso',
        site == 'Legs' |
           site == 'Knees'
            site == 'Ankles.Feet' ~ 'Lower limbs',
        site == 'Arms' |
            site == 'Elbows' |
            site == 'Wrists.Hands' ~ 'Upper limbs',
        site == 'Cervical_spine' |
            site == 'Thoracic_spine' |
            site == 'Lumbosacral_spine' ~ 'Spinal column',
        TRUE ~ 'other'
    # Nest by body region and body site
   group_by(region, site) %>%
   nest() %>%
    # Boostrap data
   mutate(boot = map(.x = data,
```

```
~ boot(data = .x,
                             statistic = prop_func,
                             R = 999,
                             stype = 'i',
                             parallel = 'multicore',
                             ncpus = 4))) %>%
    # Get CI
   mutate(ci = map(.x = boot,
                  ~ boot.ci(.x, type = 'perc'))) %>%
    # Extract ci data
   mutate(point_est = map(.x = ci,
                          ~ .x$t0),
           lower_ci = map(.x = ci,
                          ~ .x$percent[[4]]),
           upper_ci = map(.x = ci,
                          ~ .x$percent[[5]])) %>%
    # Remove columns
   select(-data, -boot, -ci) %>%
    # Unnest
   unnest(cols = c(point_est, lower_ci, upper_ci))
# Re-nest by body region and generate figures and tables
prop_boot2 <- prop_boot %>%
   group_by(region) %>%
   nest() %>%
    # Fix site labels
   mutate(data = map(.x = data,
                      ~ .x %>%
                          mutate(site = str_replace_all(site,
                                                        pattern = '_',
                                                        replacement = ' '),
                                 site = str_replace_all(site,
                                                        pattern = '\\.',
                                                        replacement = ' & ')))) %>%
    # Re-order sites by point_est
   mutate(data = map(.x = data,
                      ~ .x %>%
                          mutate(site = fct_reorder(site,
                                                    point_est)))) %>%
    # Plot data
   mutate(plots = map2(.x = data,
                        y = region,
                       ~ .x %>%
                           ggplot(data = .) +
                           aes(x = site,
                               y = point_est,
                               ymin = lower_ci,
                               ymax = upper_ci) +
                           geom_linerange(size = 1) +
                           geom_point(size = 6) +
                           coord_flip() +
                            labs(title = .y,
                                 subtitle = '(Point estimate with 95%CI)',
```

```
y = 'Proportion with pain') +
                        scale_y_continuous(limits = c(0, 1)) +
                        theme_minimal(base_size = 18) +
                        theme(plot.title = element_text(size = 18),
                              plot.subtitle = element_text(size = 12),
                              axis.title.y = element_blank(),
                              panel.grid = element_blank(),
                              axis.text = element_text(colour = '#000000'),
                              axis.line = element_line(size = 0.5),
                              axis.ticks = element_line(size = 0.5)))) %>%
# Tabulate data
mutate(tables = map2(.x = data,
                     .y = region,
                     ~ .x %>%
                         kable(caption = .y,
                               digits = 2)))
```

### Tabulated proportions (with 95% Cls), by body region

```
walk(prop_boot2$tables, ~ print(.x))
```

Table 6: Head and upper torso

site	point_est	lower_ci	upper_ci
Head	0.33	0.29	0.37
Throat	0.03	0.02	0.05
Shoulder	0.08	0.06	0.10
Chest	0.19	0.16	0.23

Table 7: Upper limbs

site	point_est	lower_ci	upper_ci
Arms	0.04	0.03	0.06
Elbows	0.04	0.02	0.05
Wrists & Hands	0.06	0.04	0.08

Table 8: Lower torso

site	point_est	lower_ci	upper_ci
Lower back	0.07	0.05	0.09
Abdomen	0.27	0.23	0.30
Groin	0.09	0.07	0.12
Hips	0.07	0.05	0.09
Buttocks	0.03	0.02	0.05

Table 9: Spinal column

site	point_est	lower_ci	upper_ci
Cervical spine	0.05	0.03	0.07
Thoracic spine	0.12	0.10	0.15
Lumbosacral spine	0.15	0.12	0.18

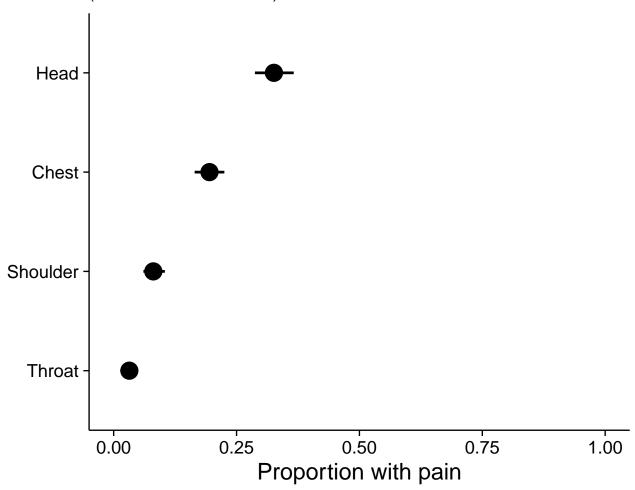
Table 10: Lower limbs

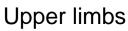
site	point_est	lower_ci	upper_ci
Legs	0.16	0.13	0.20
Knees	0.14	0.11	0.17
Ankles & Feet	0.31	0.27	0.35

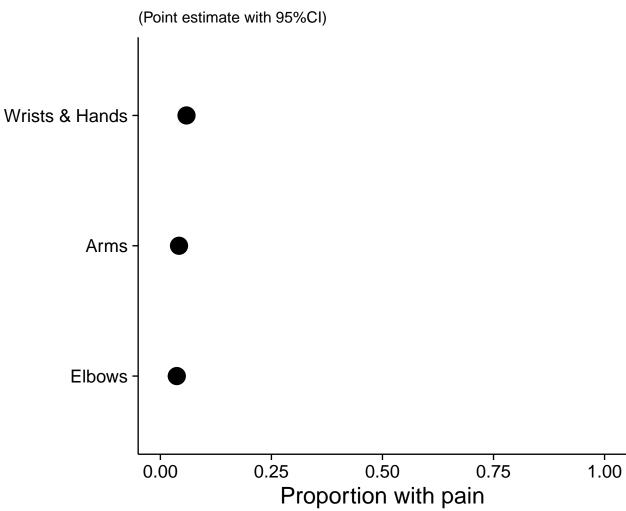
## Plotted proportions (with 95% Cls), by body region

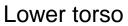
walk(prop\_boot2\$plots, ~ print(.x))

# Head and upper torso

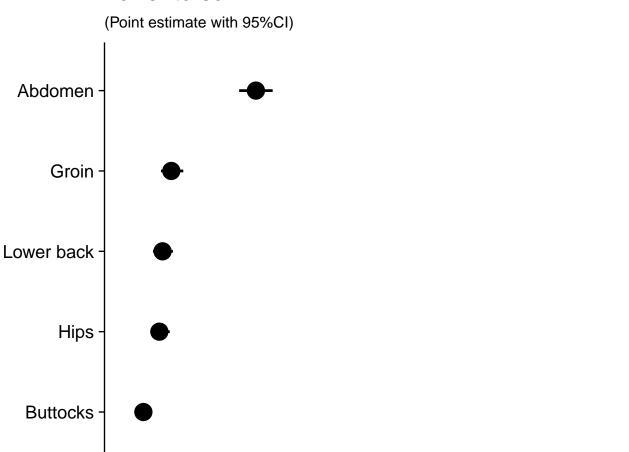








0.00



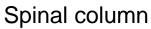
0.50

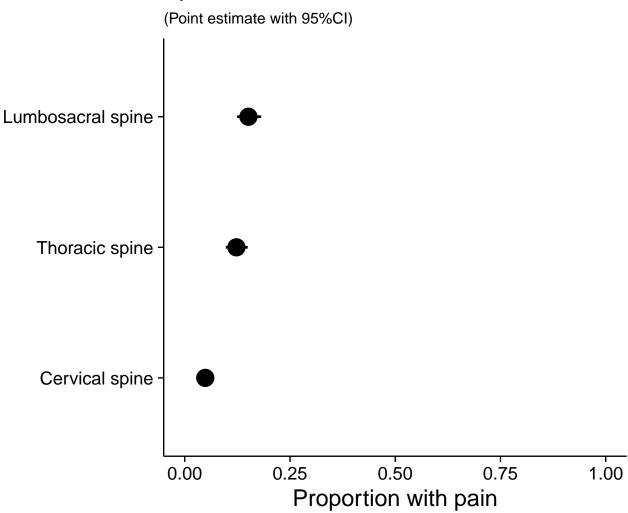
Proportion with pain

0.25

0.75

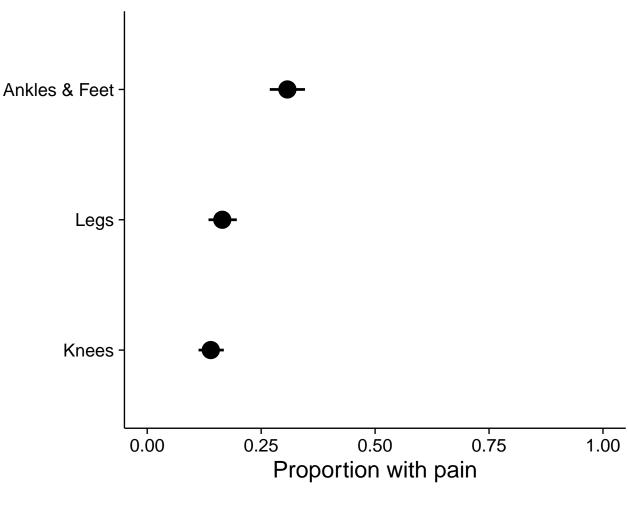
1.00





### Lower limbs





## By sex

#### **Process data**

```
# Set seed
set.seed(2020)

# Select sex data
sex <- demo[, c('ID', 'Sex')]

# Join to boot_data & remove ID and Upper_back (only one outcome -- no pain)
sex <- left_join(data, sex) %>%
    select(-ID, -Upper_back)

# Bootstrap CIs
```

```
sex_boot <- sex %>%
    # Pivot to long format
   pivot_longer(cols = -Sex,
                 names_to = 'site',
                 values_to = 'pain_present') %>%
    # Add body regions
   mutate(region = case_when(
        site == 'Chest'
            site == 'Head' |
            site == 'Throat' |
           site == 'Shoulder' ~ 'Head and upper torso',
        site == 'Lower_back' |
           site == 'Abdomen' |
           site == 'Hips' |
           site == 'Buttocks' |
           site == 'Groin' ~ 'Lower torso',
        site == 'Legs' |
           site == 'Knees' |
           site == 'Ankles.Feet' ~ 'Lower limbs',
        site == 'Arms' |
            site == 'Elbows' |
            site == 'Wrists.Hands' ~ 'Upper limbs',
        site == 'Cervical_spine' |
            site == 'Thoracic_spine' |
           site == 'Lumbosacral_spine' ~ 'Spinal column',
       TRUE ~ 'other'
   )) %>%
    # Nest by body region and body site
   group_by(Sex, region, site) %>%
   nest() %>%
    # Boostrap data
   mutate(boot = map(.x = data,
                      ~ boot(data = .x,
                             statistic = prop_func,
                             R = 999,
                             stype = 'i',
                             parallel = 'multicore',
                             ncpus = 4))) %>%
    # Get CI
   mutate(ci = map(.x = boot,
                   ~ boot.ci(.x, type = 'perc'))) %>%
    # Extract ci data
   mutate(point_est = map(.x = ci,
                           ~ .x$t0),
           lower_ci = map(.x = ci,
                          ~ .x$percent[[4]]),
           upper_ci = map(.x = ci,
                          ~ .x$percent[[5]])) %>%
    # Remove columns
    select(-data, -boot, -ci) %>%
    # Unnest
   unnest(cols = c(point_est, lower_ci, upper_ci))
```

```
# Re-nest by body region and generate figures and tables
sex_boot2 <- sex_boot %>%
   group_by(region) %>%
   nest() %>%
    # Fix site labels
   mutate(data = map(.x = data,
                      ~ .x %>%
                          mutate(site = str_replace_all(site,
                                                         pattern = '_',
                                                         replacement = ' '),
                                 site = str_replace_all(site,
                                                        pattern = '\\.',
                                                        replacement = ' & ')))) %>%
    # Re-order sites by point_est
   mutate(data = map(.x = data,
                      ~ .x %>%
                          mutate(site = fct_reorder(site,
                                                    point_est)))) %>%
    # Plot data
   mutate(plots = map2(.x = data,
                        .y = region,
                       ~ .x %>%
                           ggplot(data = .) +
                           aes(x = site,
                               y = point_est,
                               ymin = lower_ci,
                               ymax = upper ci,
                               fill = Sex) +
                           geom_linerange(position = position_dodge2(width = 0.6),
                                          size = 1,
                                          colour = '#000000') +
                           geom_point(shape = 21,
                                      colour = '#000000',
                                      position = position_dodge2(width = 0.6),
                                      size = 6,
                                      stroke = 1) +
                           coord_flip() +
                           labs(title = .y,
                                subtitle = '(Point estimate with 95%CI)',
                                y = 'Proportion with pain') +
                           scale_y_continuous(limits = c(0, 1)) +
                           scale_fill_manual(values = c('#000000', '#FFFFFF')) +
                           theme_minimal(base_size = 18) +
                           theme(plot.title = element_text(size = 18),
                                 plot.subtitle = element_text(size = 12),
                                 legend.title = element_blank(),
                                 legend.position = 'top',
                                 axis.title.y = element_blank(),
                                 panel.grid = element_blank(),
                                 axis.text = element_text(colour = '#000000'),
                                 axis.line = element_line(size = 0.5),
                                 axis.ticks = element_line(size = 0.5)))) %>%
    # Tabulate data
```

### Tabulated proportions (with 95% Cls), by age and body region

```
walk(sex_boot2$tables, ~ print(.x))
```

Table 11: Head and upper torso

Sex	site	point_est	lower_ci	upper_ci
Female	Head	0.38	0.34	0.42
Female	Throat	0.03	0.01	0.05
Female	Shoulder	0.07	0.05	0.09
Female	Chest	0.19	0.16	0.23
Male	Head	0.12	0.07	0.18
Male	Throat	0.04	0.01	0.08
Male	Shoulder	0.12	0.07	0.18
Male	Chest	0.20	0.13	0.28

Table 12: Upper limbs

Sex	site	point_est	lower_ci	upper_ci
Female	Arms	0.04	0.03	0.06
Female	Elbows	0.03	0.02	0.05
Female	Wrists & Hands	0.06	0.04	0.09
Male	Arms	0.03	0.01	0.07
Male	Elbows	0.05	0.02	0.09
Male	Wrists & Hands	0.03	0.01	0.07

Table 13: Lower torso

Sex	site	point_est	lower_ci	upper_ci
Female	Lower back	0.08	0.05	0.10
Female	Abdomen	0.29	0.24	0.33
Female	Groin	0.10	0.07	0.12
Female	Hips	0.06	0.04	0.08
Female	Buttocks	0.03	0.01	0.05
Male	Lower back	0.06	0.02	0.10
Male	Abdomen	0.20	0.12	0.28
Male	Groin	0.07	0.03	0.12
Male	Hips	0.09	0.04	0.16
Male	Buttocks	0.04	0.01	0.09

Table 14: Spinal column

Sex	site	point_est	lower_ci	upper_ci
Female	Cervical spine	0.05	0.03	0.08
Female	Thoracic spine	0.14	0.11	0.17
Female	Lumbosacral spine	0.15	0.12	0.19
Male	Cervical spine	0.03	0.00	0.06
Male	Thoracic spine	0.07	0.03	0.11
Male	Lumbosacral spine	0.14	0.08	0.21

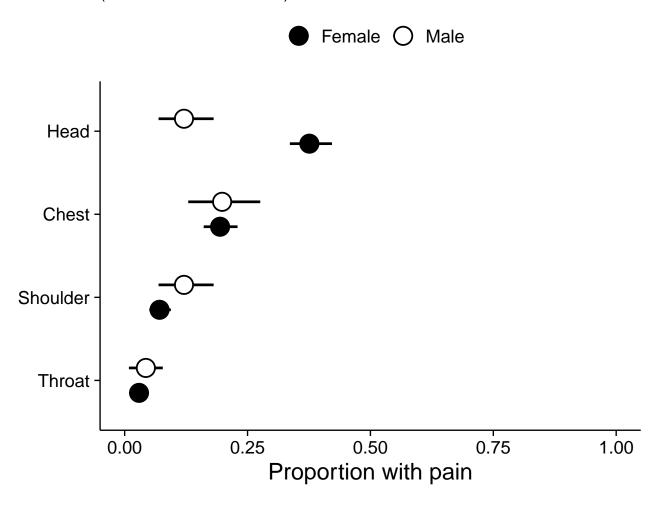
Table 15: Lower limbs

Sex	site	point_est	lower_ci	upper_ci
Female	Legs	0.15	0.12	0.18
Female	Knees	0.13	0.10	0.16
Female	Ankles & Feet	0.28	0.24	0.32
Male	Legs	0.23	0.16	0.31
Male	Knees	0.19	0.12	0.27
Male	Ankles & Feet	0.41	0.32	0.50

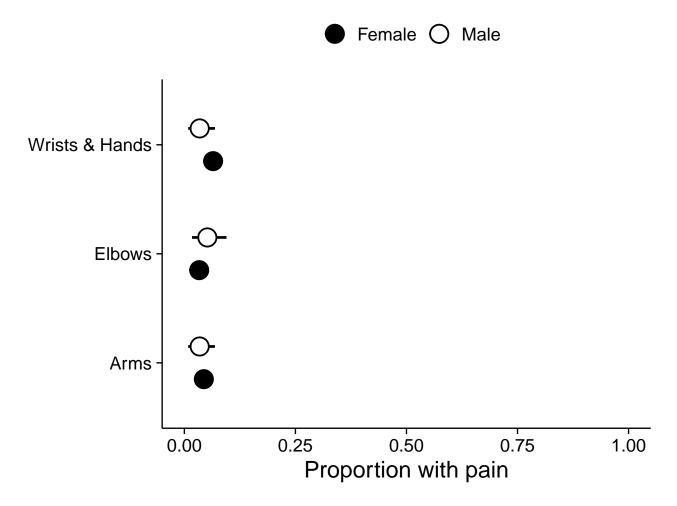
## Plotted proportions (with 95% Cls), by age and body region

walk(sex\_boot2\$plots, ~ print(.x))

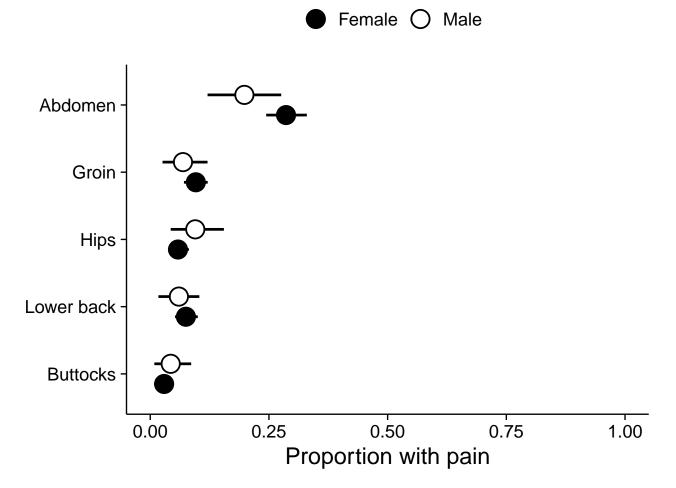
# Head and upper torso



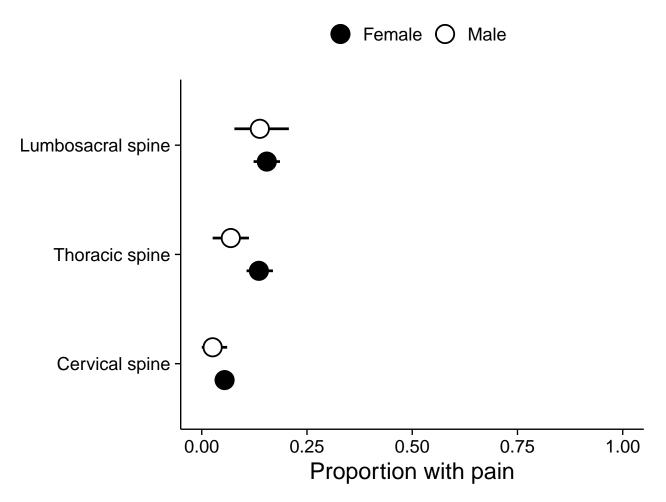
# Upper limbs



# Lower torso



# Spinal column



### Lower limbs

(Point estimate with 95%CI)



### By age

#### **Process data**

```
# Set seed
set.seed(2020)

# Select age data
age <- demo[, c('ID', 'Age')]

# Join to boot_data & remove ID and Upper_back (only one outcome -- no pain)
age <- left_join(data, age) %>%
    select(-ID, -Upper_back)

# Get complete cases
```

```
age <- age[complete.cases(age), ]</pre>
# Pivot and add age group categories (10 year periods)
age_boot <- age %>%
    # Pivot to long format
    pivot_longer(cols = -Age,
                 names_to = 'site',
                 values_to = 'pain_present') %>%
    # Add age categories
    mutate(age_group = case_when(
        Age < 28 \sim '18-27',
        Age \geq 28 \& Age < 38 ~ '28-37',
        Age \geq 38 \& Age < 48 ~ '38-47',
        Age \geq 48 \& Age < 58 ~ '48-57',
        Age >= 58 \& Age < 68 ~ '58-67',
        Age >= 68 & Age < 78 ~ '68-77',
        Age >= 78 & Age < 88 ~ '78-87'
    ))
# Print count per age group
age_boot %>%
    group_by(site, age_group) %>%
    summarise(count = n()) %>%
    filter(site == 'Abdomen') %>%
    ungroup() %>%
    select(-site) %>%
    kable(caption = 'Participant count per age group')
```

Table 16: Participant count per age group

age_group	count
18-27	64
28-37	284
38-47	158
48-57	59
58-67	20
68-77	2

```
stype = 'i',
                             parallel = 'multicore',
                             ncpus = 4))) %>%
    # Get CI
   mutate(ci = map(.x = boot,
                   ~ boot.ci(.x, type = 'perc'))) %>%
    # Extract ci data
   mutate(point_est = map(.x = ci,
                         ~ .x$t0),
           lower_ci = map(.x = ci,
                         ~ .x$percent[[4]]),
           upper_ci = map(.x = ci,
                          ~ .x$percent[[5]])) %>%
    # Remove columns
    select(-data, -boot, -ci) %>%
    # Unnest
   unnest(cols = c(point_est, lower_ci, upper_ci)) %>%
    ungroup()
# Re-nest by body region and generate figures and tables
age_boot2 <- age_boot2 %>%
    # Fix site labels
   mutate(site = str_replace_all(site,
                                  pattern = '_',
                                  replacement = ' '),
           site = str_replace_all(site,
                                  pattern = '\\.',
                                  replacement = ' & ')) %>%
    # Group and nest
   group_by(site) %>%
   nest() %>%
    # Arrange age groups
    # Plot data
   mutate(plots = map2(.x = data,
                        .y = site,
                       ~ .x %>%
                           ggplot(data = .) +
                           aes(x = age_group,
                               y = point_est,
                               ymin = lower_ci,
                               ymax = upper_ci) +
                           geom_linerange(size = 1,
                                          colour = '#000000') +
                           geom point(colour = '#000000',
                                      size = 6) +
                           labs(title = .y,
                                subtitle = '(Point estimate with 95%CI)',
                                caption = 'Age group 68-77 years removed because n = 2',
                                x = 'Age group (Years)',
                                y = 'Proportion with pain') +
                           scale_y_continuous(limits = c(0, 1)) +
                           theme_minimal(base_size = 18) +
                           theme(plot.title = element_text(size = 18),
```

#### Tabulated proportions (with 95% Cls), by age group and body site

```
walk(age_boot2$tables, ~ print(.x))
```

Table 17: Head (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.45	0.33	0.58
28-37	0.33	0.28	0.39
38-47	0.31	0.24	0.39
48-57	0.17	0.08	0.27
58-67	0.35	0.15	0.60

Table 18: Throat (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.05	0.00	0.11
28-37	0.03	0.01	0.05
38-47	0.03	0.01	0.06
48-57	0.05	0.00	0.12
58-67	0.00	0.00	0.00

Table 19: Shoulder (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.12	0.05	0.22
28-37	0.07	0.04	0.10
38-47	0.09	0.05	0.15
48-57	0.07	0.02	0.14
58-67	0.05	0.00	0.15

Table 20: Arms (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.02	0.00	0.05
28-37	0.04	0.02	0.07
38-47	0.06	0.03	0.09
48-57	0.05	0.00	0.12
58-67	0.00	0.00	0.00

Table 21: Elbows (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.00	0.00	0.00
28-37	0.03	0.01	0.05
38-47	0.04	0.01	0.07
48-57	0.05	0.00	0.10
58-67	0.15	0.00	0.30

Table 22: Wrists & Hands (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.02	0.00	0.05
28-37	0.07	0.04	0.10
38-47	0.04	0.01	0.08
48-57	0.07	0.02	0.14
58-67	0.15	0.00	0.30

Table 23: Chest (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.17	0.09	0.27
28-37	0.20	0.16	0.25
38-47	0.20	0.14	0.27
48-57	0.19	0.10	0.29
58-67	0.20	0.05	0.40

Table 24: Lower back (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.09	0.03	0.17
28-37	0.07	0.04	0.10
38-47	0.05	0.02	0.09
48-57	0.10	0.03	0.19

age_group	point_est	lower_ci	upper_ci
58-67	0.10	0.00	0.25

Table 25: Abdomen (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.22	0.12	0.33
28-37	0.32	0.26	0.37
38-47	0.30	0.23	0.37
48-57	0.12	0.03	0.20
58-67	0.05	0.00	0.15

Table 26: Cervical spine (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.12	0.05	0.22
28-37	0.05	0.02	0.08
38-47	0.04	0.01	0.08
48-57	0.00	0.00	0.00
58-67	0.00	0.00	0.00

Table 27: Thoracic spine (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.17	0.09	0.27
28-37	0.14	0.10	0.18
38-47	0.06	0.03	0.11
48-57	0.10	0.03	0.19
58-67	0.25	0.05	0.45

Table 28: Lumbosacral spine (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.17	0.09	0.27
28-37	0.15	0.11	0.19
38-47	0.11	0.06	0.16
48-57	0.15	0.07	0.25
58-67	0.40	0.20	0.60

Table 29: Groin (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.09	0.03	0.17
28-37	0.10	0.07	0.14
38-47	0.09	0.04	0.14
48-57	0.05	0.00	0.12
58-67	0.05	0.00	0.15

Table 30: Hips (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.03	0.00	0.08
28-37	0.05	0.02	0.07
38-47	0.09	0.04	0.13
48-57	0.14	0.05	0.22
58-67	0.10	0.00	0.25

Table 31: Legs (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.12	0.05	0.22
28-37	0.13	0.09	0.17
38-47	0.21	0.15	0.27
48-57	0.29	0.17	0.41
58-67	0.20	0.05	0.40

Table 32: Knees (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.11	0.05	0.19
28-37	0.10	0.07	0.14
38-47	0.16	0.11	0.22
48-57	0.22	0.12	0.32
58-67	0.30	0.10	0.50

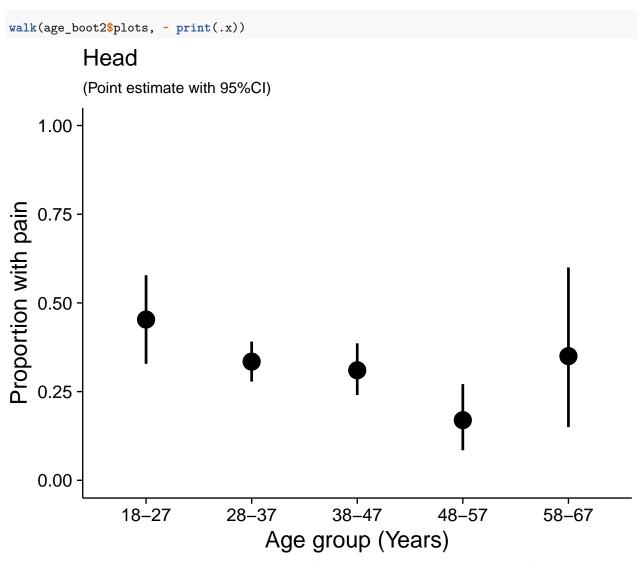
Table 33: Ankles & Feet (Age group 68-77 years removed because n=2)

age_group	point_est	lower_ci	upper_ci
18-27	0.22	0.12	0.33
28-37	0.24	0.19	0.29
38-47	0.37	0.29	0.44
48-57	0.51	0.39	0.63
58-67	0.55	0.35	0.75

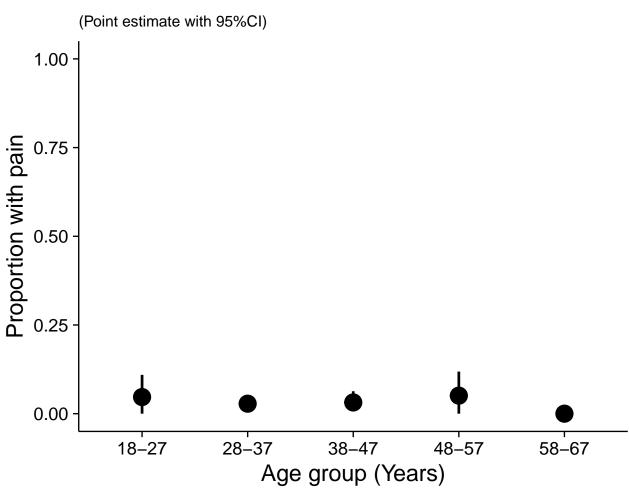
Table 34: Buttocks (Age group 68-77 years removed because n = 2)

age_group	point_est	lower_ci	upper_ci
18-27	0.03	0.00	0.08
28-37	0.04	0.02	0.06
38-47	0.03	0.01	0.05
48-57	0.02	0.00	0.05
58-67	0.05	0.00	0.15

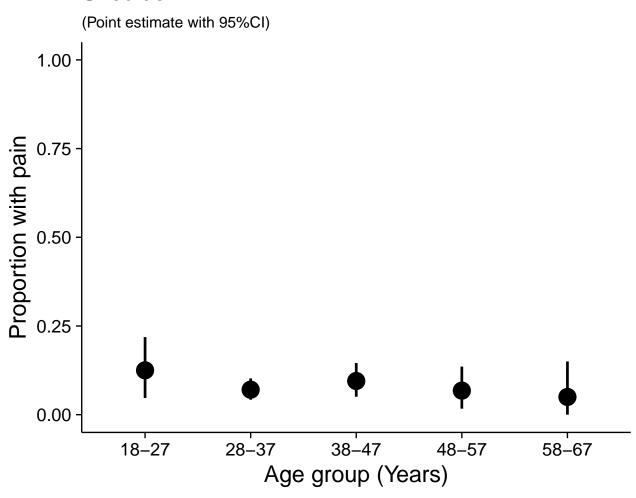
### Plotted proportions (with 95% Cls), by age group and body site





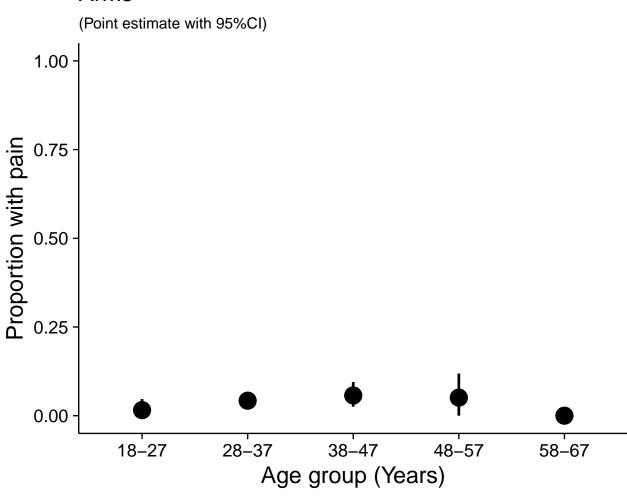


# Shoulder

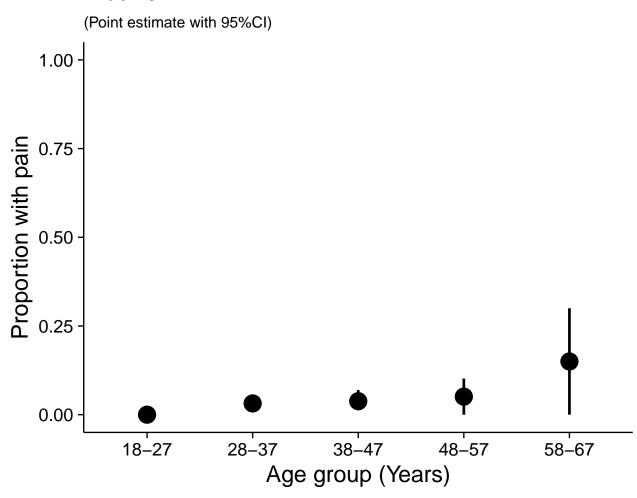


Age group 68-77 years removed because n = 2



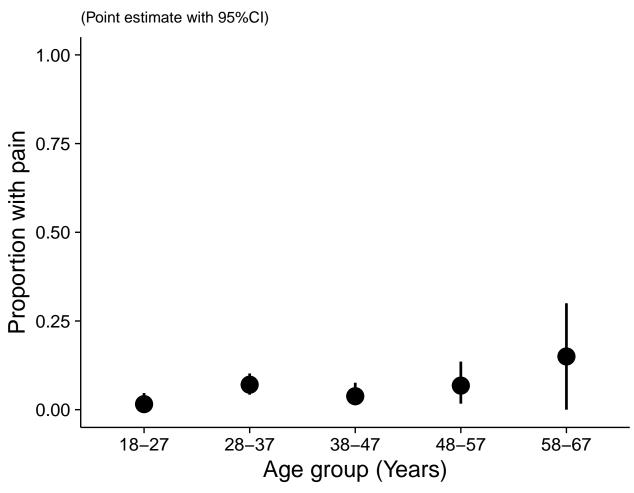


## **Elbows**



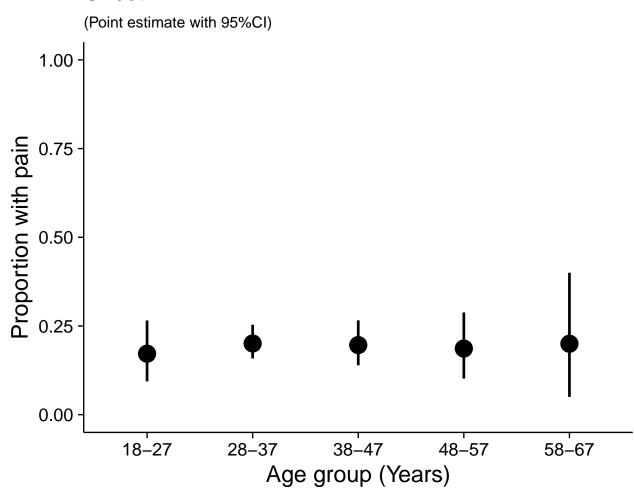
Age group 68-77 years removed because n = 2

## Wrists & Hands



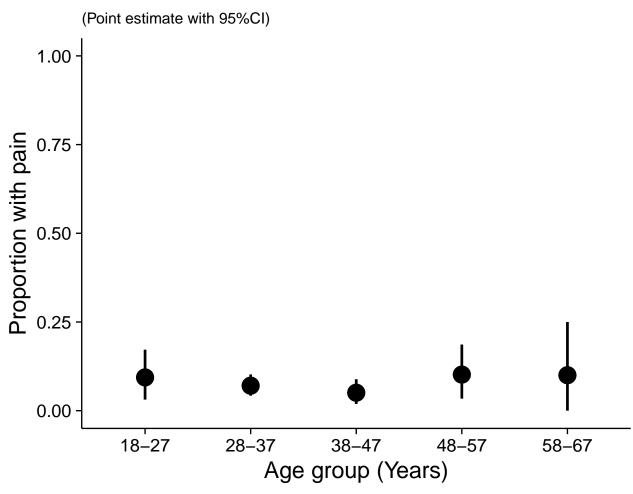
Age group 68-77 years removed because n = 2





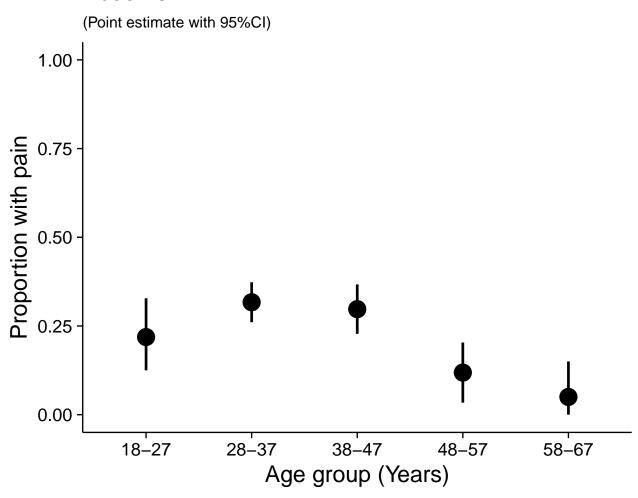
Age group 68-77 years removed because n = 2

## Lower back



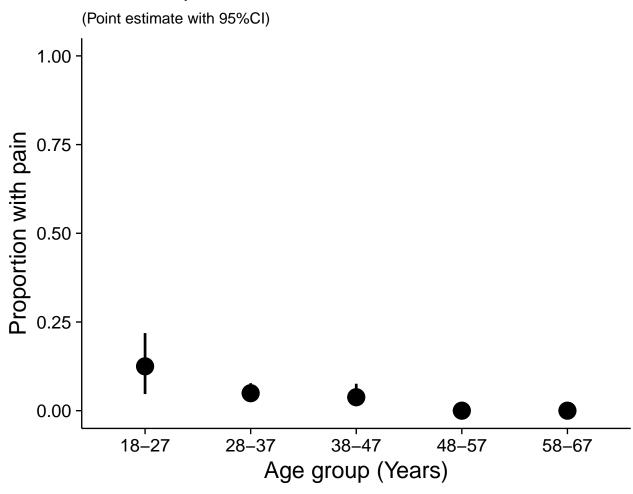
Age group 68-77 years removed because n = 2

## Abdomen



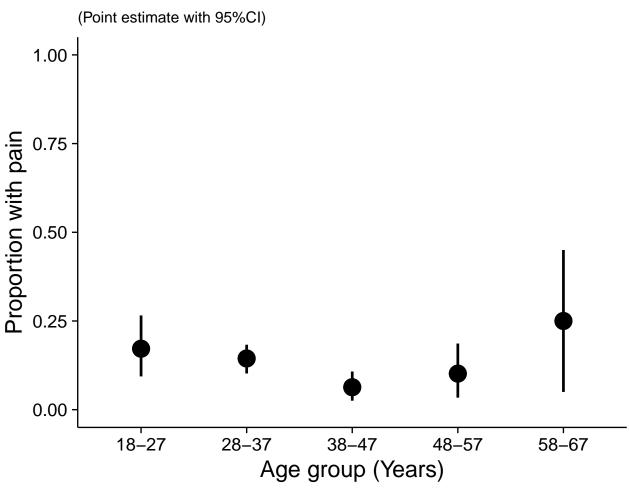
Age group 68-77 years removed because n = 2

# Cervical spine



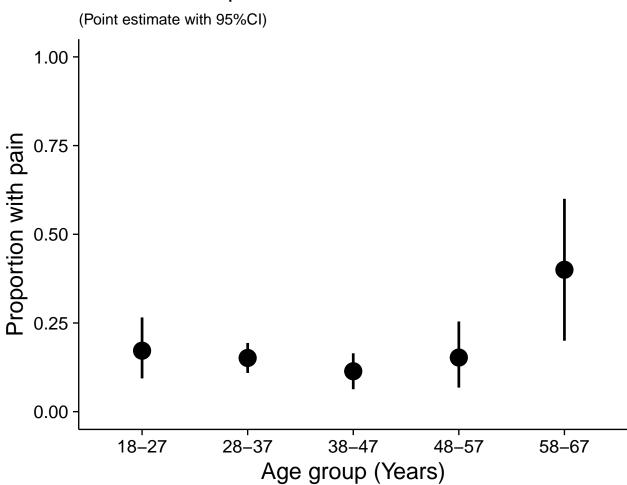
Age group 68-77 years removed because n = 2

# Thoracic spine



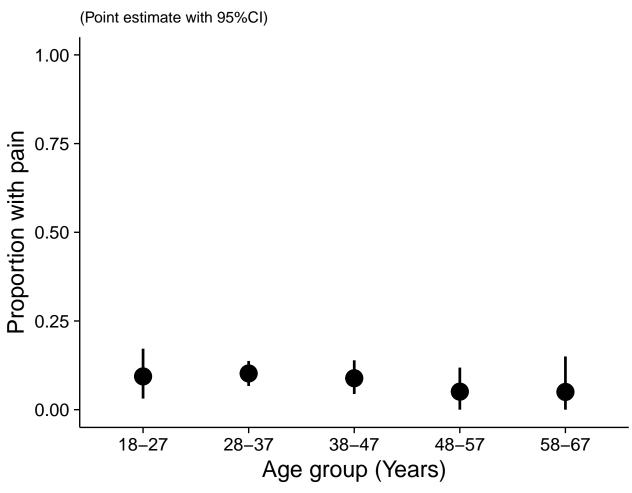
Age group 68-77 years removed because n = 2

# Lumbosacral spine

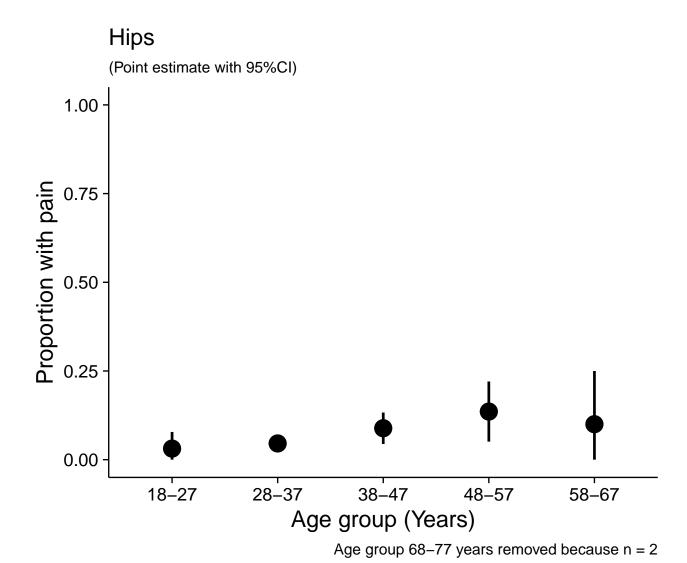


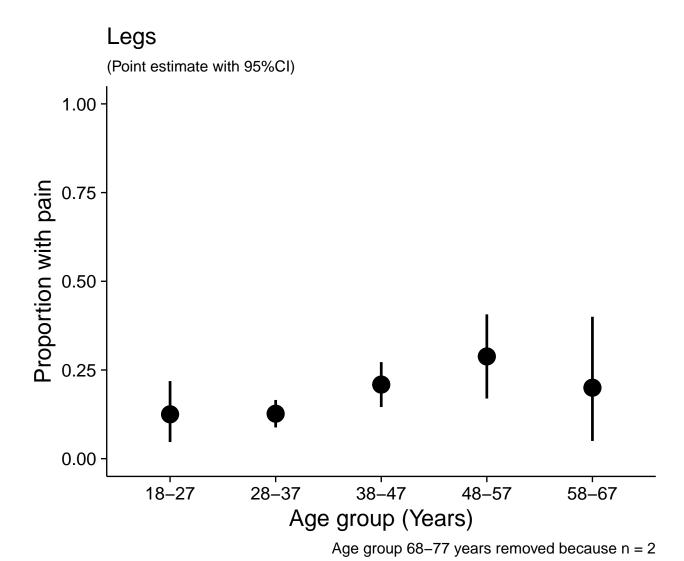
Age group 68-77 years removed because n = 2



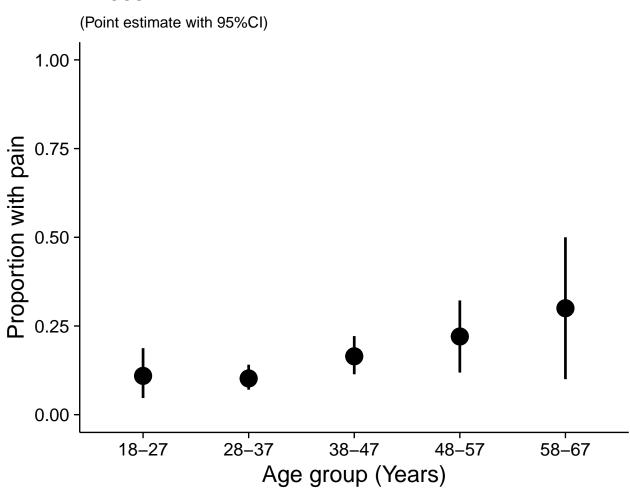


Age group 68-77 years removed because n = 2



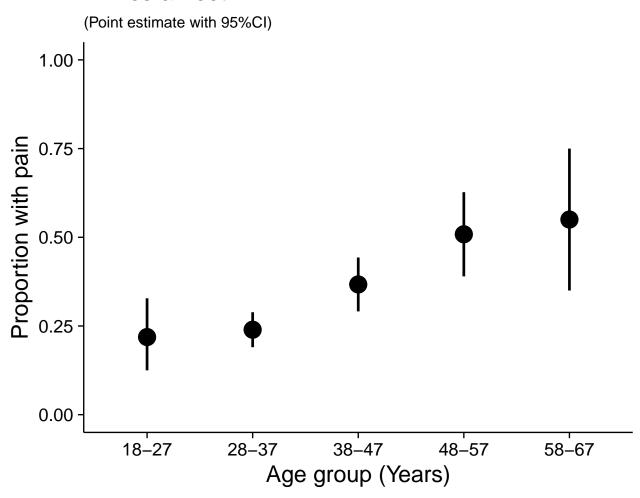


# Knees



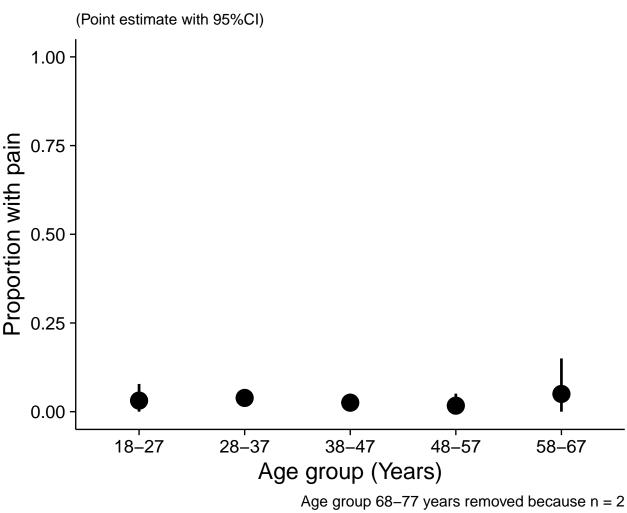
Age group 68-77 years removed because n = 2

## Ankles & Feet



Age group 68-77 years removed because n = 2





### By most recent CD4 T-cell count

### **Process data**

```
# Set seed
set.seed(2020)
# Select CD4 recent data
cd4 <- demo[, c('ID', 'CD4_recent')]</pre>
# Join to boot_data & remove ID and Upper_back (only one outcome -- no pain)
cd4 <- left_join(data, cd4) %>%
    select(-ID, -Upper_back)
# Get complete cases
```

```
cd4 <- cd4[complete.cases(cd4), ]</pre>
# Pivot and add CD4 recent group categories (counts of 100)
cd4_boot <- cd4 %>%
    # Pivot to long format
   pivot_longer(cols = -CD4_recent,
                 names_to = 'site',
                 values to = 'pain present') %>%
    # Add CD4 recent categories
   mutate(cd4_group = case_when(
       CD4_recent < 100 ~ '0-99',
        CD4_recent >= 100 & CD4_recent < 200 ~ '100-199',
        CD4_recent >= 200 & CD4_recent < 300 ~ '200-299',
        CD4_recent >= 300 & CD4_recent < 400 ~ '300-399',</pre>
        CD4_recent >= 400 & CD4_recent < 500 ~ '400-499',
        CD4_recent >= 500 ~ '500+'
   ))
# Print count per CD4 recent group
cd4_boot %>%
    group_by(site, cd4_group) %>%
   summarise(count = n()) %>%
   filter(site == 'Abdomen') %>%
   ungroup() %>%
   select(-site) %>%
   kable(caption = 'Participant count per CD4 group')
```

Table 35: Participant count per CD4 group

cd4_group	count
0-99	68
100-199	105
200-299	108
300-399	76
400-499	46
500+	93

```
# Get CI
   mutate(ci = map(.x = boot,
                    ~ boot.ci(.x, type = 'perc'))) %>%
    # Extract ci data
   mutate(point_est = map(.x = ci,
                           ~ .x$t0),
           lower_ci = map(.x = ci,
                         ~ .x$percent[[4]]),
           upper_ci = map(.x = ci,
                          ~ .x$percent[[5]])) %>%
    # Remove columns
    select(-data, -boot, -ci) %>%
    # Unnest
    unnest(cols = c(point est, lower ci, upper ci)) %>%
   ungroup()
# Re-nest by body region and generate figures and tables
cd4_boot2 <- cd4_boot2 %>%
    # Fix site labels
   mutate(site = str_replace_all(site,
                                  pattern = '_',
                                  replacement = ' '),
           site = str_replace_all(site,
                                  pattern = '\\.',
                                  replacement = ' & ')) %>%
    # Group and nest
   group by(site) %>%
   nest() %>%
    # Arrange CD4 recent groups
   # Plot data
   mutate(plots = map2(.x = data,
                        .y = site,
                       ~ .x %>%
                           ggplot(data = .) +
                           aes(x = cd4\_group,
                               y = point_est,
                               ymin = lower_ci,
                               ymax = upper_ci) +
                           geom_linerange(size = 1,
                                          colour = '#000000') +
                           geom_point(colour = '#000000',
                                      size = 6) +
                           labs(title = .y,
                                subtitle = '(Point estimate with 95%CI)',
                                x = expression('CD4 group (cells.mm'^-3*')'),
                                y = 'Proportion with pain') +
                           scale_y_continuous(limits = c(0, 1)) +
                           theme_minimal(base_size = 18) +
                           theme(plot.title = element_text(size = 18),
                                 plot.subtitle = element_text(size = 12),
                                 panel.grid = element_blank(),
                                 axis.text = element_text(colour = '#000000'),
                                 axis.line = element_line(size = 0.5),
```

### Tabulated proportions (with 95% Cls), by CD4 (recent) group and body site

```
walk(cd4_boot2$tables, ~ print(.x))
```

Table 36: Head

cd4_group	point_est	lower_ci	upper_ci
0-99	0.26	0.18	0.37
100-199	0.39	0.30	0.49
200-299	0.32	0.24	0.42
300-399	0.37	0.26	0.49
400-499	0.30	0.17	0.43
500+	0.25	0.17	0.33

Table 37: Throat

cd4_group	point_est	lower_ci	upper_ci
0-99	0.07	0.01	0.15
100-199	0.02	0.00	0.05
200-299	0.03	0.00	0.06
300-399	0.01	0.00	0.04
400-499	0.04	0.00	0.11
500+	0.00	0.00	0.00

Table 38: Shoulder

cd4_group	point_est	lower_ci	upper_ci
0-99	0.07	0.01	0.15
100-199	0.10	0.05	0.16
200-299	0.06	0.02	0.11
300-399	0.05	0.01	0.11
400-499	0.15	0.04	0.26
500+	0.09	0.03	0.15

Table 39: Arms

cd4_group	point_est	lower_ci	upper_ci
0-99	0.06	0.01	0.12
100-199	0.09	0.04	0.14
200-299	0.00	0.00	0.00
300-399	0.04	0.00	0.08
400-499	0.02	0.00	0.07
500+	0.05	0.02	0.11

Table 40: Elbows

cd4_group	point_est	lower_ci	upper_ci
0-99	0.01	0.00	0.04
100-199	0.05	0.01	0.10
200-299	0.02	0.00	0.06
300-399	0.04	0.00	0.09
400-499	0.07	0.00	0.15
500+	0.06	0.02	0.12

Table 41: Wrists & Hands

cd4_group	point_est	lower_ci	upper_ci
0-99	0.01	0.00	0.04
100-199	0.06	0.02	0.10
200-299	0.06	0.02	0.10
300-399	0.07	0.01	0.13
400-499	0.07	0.00	0.15
500+	0.14	0.08	0.22

Table 42: Chest

cd4_group	point_est	lower_ci	upper_ci
0-99	0.34	0.22	0.46
100-199	0.22	0.14	0.30
200-299	0.17	0.10	0.23
300-399	0.17	0.09	0.26
400-499	0.17	0.09	0.28
500+	0.11	0.04	0.17

Table 43: Lower back

cd4_group	point_est	lower_ci	upper_ci
0-99	0.06	0.01	0.12
100-199	0.08	0.03	0.13
200-299	0.06	0.02	0.10
300-399	0.07	0.01	0.13

cd4_group	point_est	lower_ci	upper_ci
400-499	0.11	0.02	0.20
500+	0.11	0.04	0.17

Table 44: Abdomen

cd4_group	point_est	lower_ci	upper_ci
0-99	0.34	0.22	0.46
100-199	0.30	0.21	0.38
200-299	0.27	0.19	0.36
300-399	0.22	0.13	0.32
400-499	0.30	0.17	0.46
500+	0.24	0.16	0.32

Table 45: Cervical spine

cd4_group	point_est	lower_ci	upper_ci
0-99	0.04	0.00	0.10
100-199	0.09	0.04	0.14
200-299	0.02	0.00	0.05
300-399	0.07	0.01	0.13
400-499	0.09	0.02	0.17
500+	0.03	0.00	0.08

Table 46: Thoracic spine

cd4_group	point_est	lower_ci	upper_ci
0-99	0.13	0.06	0.22
100-199	0.10	0.05	0.17
200-299	0.10	0.05	0.16
300-399	0.14	0.08	0.22
400-499	0.22	0.11	0.35
500+	0.15	0.09	0.23

Table 47: Lumbosacral spine

cd4_group	point_est	lower_ci	upper_ci
0-99	0.09	0.03	0.16
100-199	0.13	0.07	0.20
200-299	0.15	0.08	0.21
300-399	0.17	0.09	0.26
400-499	0.30	0.17	0.43
500+	0.20	0.13	0.29

Table 48: Groin

cd4_group	point_est	lower_ci	upper_ci
0-99	0.10	0.03	0.18
100-199	0.12	0.07	0.19
200-299	0.09	0.05	0.16
300-399	0.12	0.05	0.20
400-499	0.11	0.02	0.20
500+	0.08	0.02	0.13

Table 49: Hips

cd4_group	point_est	lower_ci	upper_ci
0-99	0.01	0.00	0.06
100-199	0.06	0.02	0.10
200-299	0.05	0.01	0.09
300-399	0.05	0.01	0.11
400-499	0.11	0.02	0.20
500+	0.13	0.06	0.20

Table 50: Legs

cd4_group	point_est	lower_ci	upper_ci
0-99	0.13	0.06	0.22
100-199	0.11	0.06	0.18
200-299	0.15	0.08	0.21
300-399	0.18	0.11	0.28
400-499	0.20	0.09	0.30
500+	0.22	0.13	0.31

Table 51: Knees

cd4_group	point_est	lower_ci	upper_ci
0-99	0.12	0.04	0.19
100-199	0.11	0.06	0.17
200-299	0.15	0.08	0.21
300-399	0.12	0.05	0.20
400-499	0.17	0.07	0.28
500+	0.20	0.13	0.29

Table 52: Ankles & Feet

cd4_group	point_est	lower_ci	upper_ci
0-99	0.26	0.16	0.37
100-199	0.30	0.22	0.39
200-299	0.31	0.23	0.41

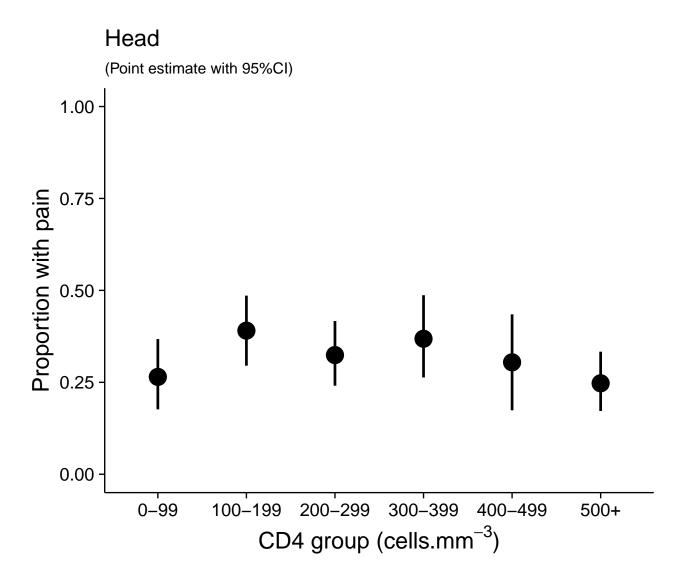
cd4_group	point_est	lower_ci	upper_ci
300-399	0.25	0.16	0.36
400-499	0.26	0.15	0.39
500+	0.33	0.24	0.43

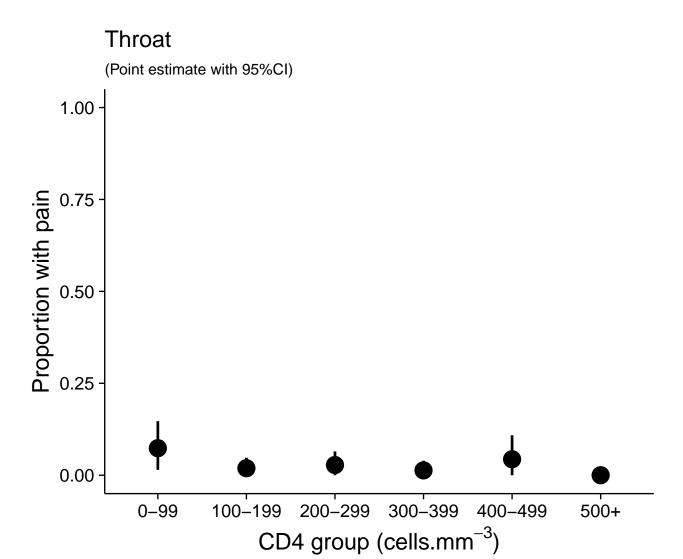
Table 53: Buttocks

cd4_group	point_est	lower_ci	upper_ci
0-99	0.04	0.00	0.10
100-199	0.01	0.00	0.03
200-299	0.03	0.00	0.06
300-399	0.05	0.01	0.12
400-499	0.04	0.00	0.11
500+	0.06	0.02	0.12

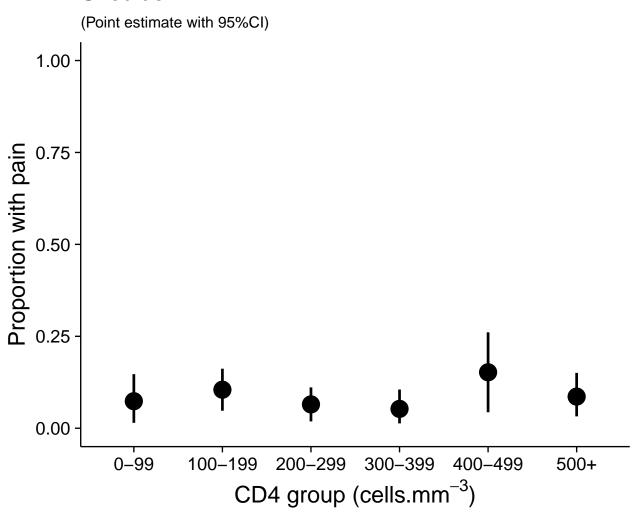
## Plotted proportions (with 95% Cls), by CD4 (recent) group and body site

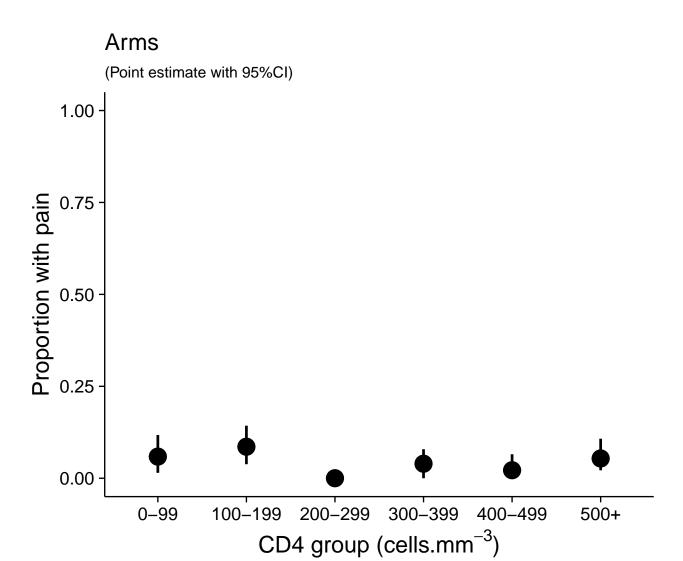
```
walk(cd4_boot2$plots, ~ print(.x))
```

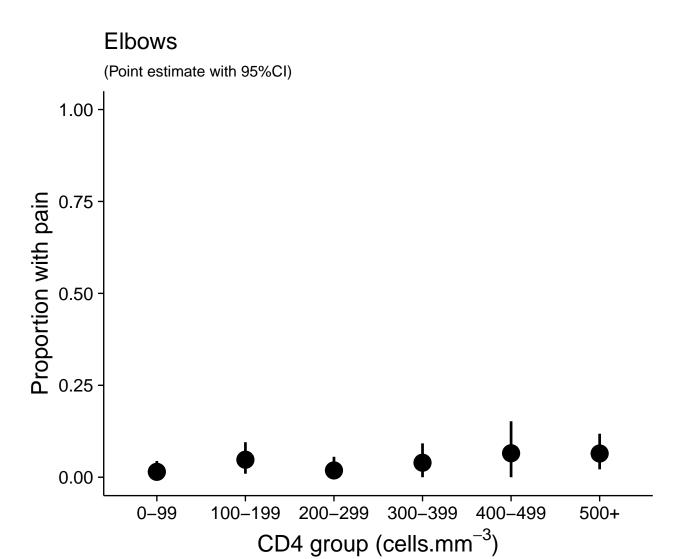




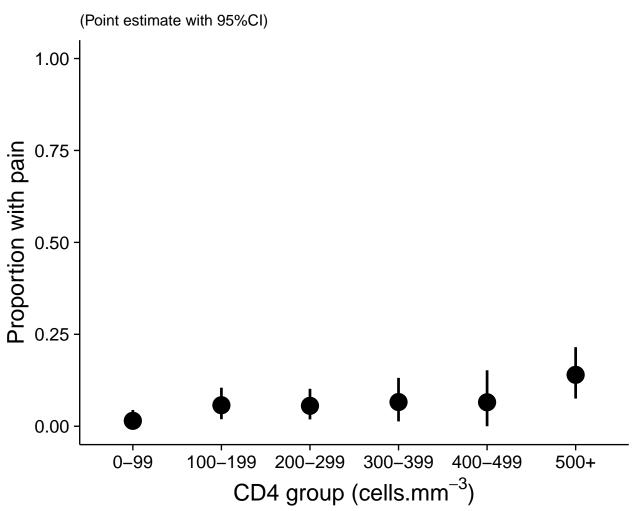


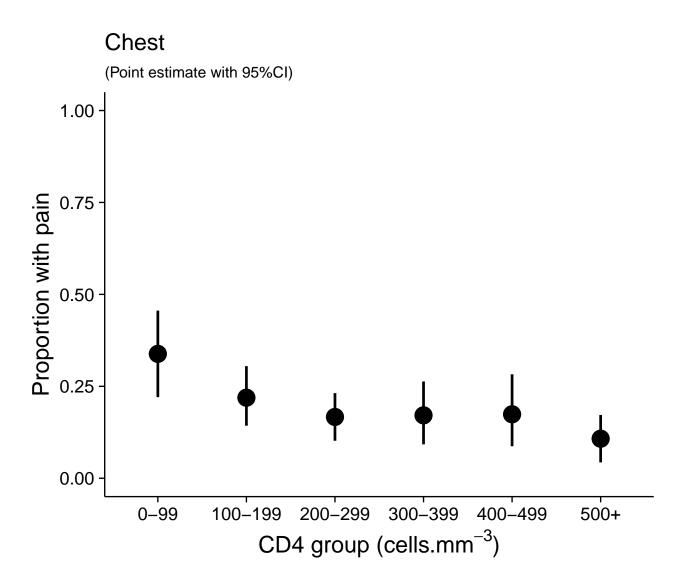




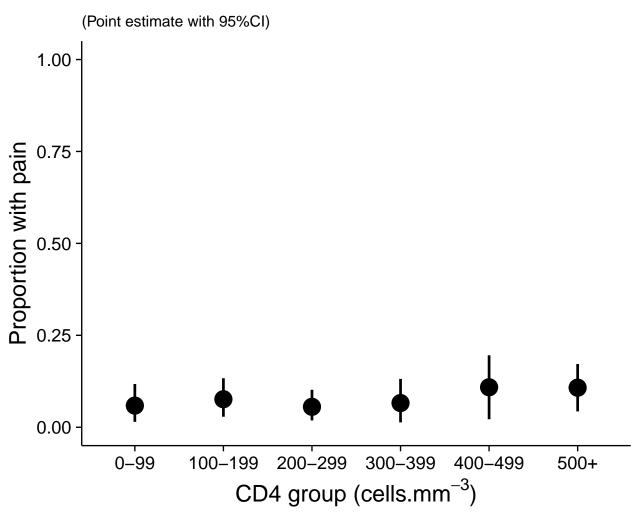


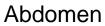


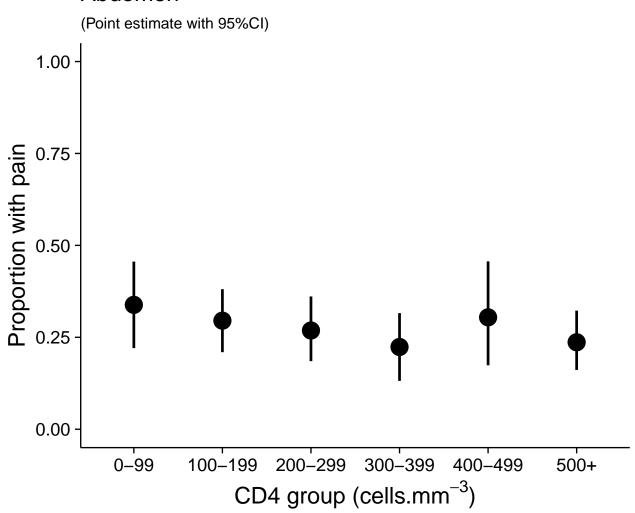




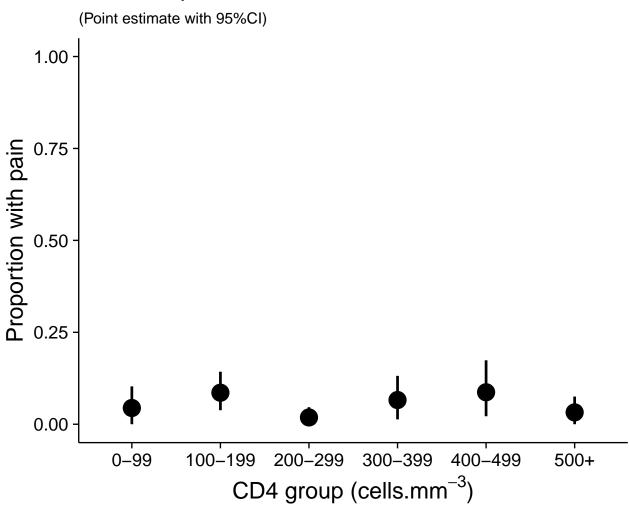




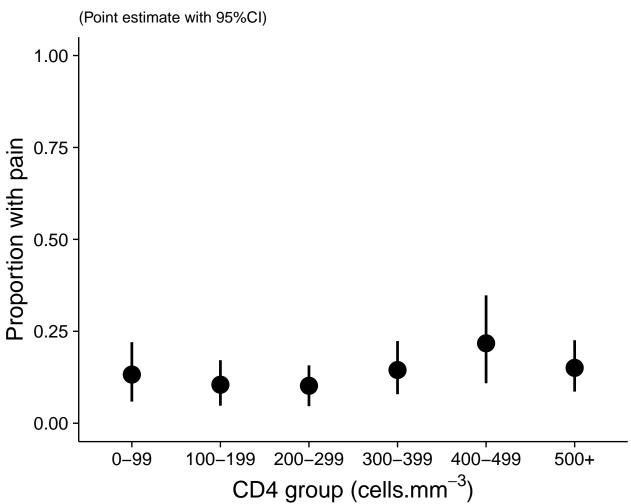




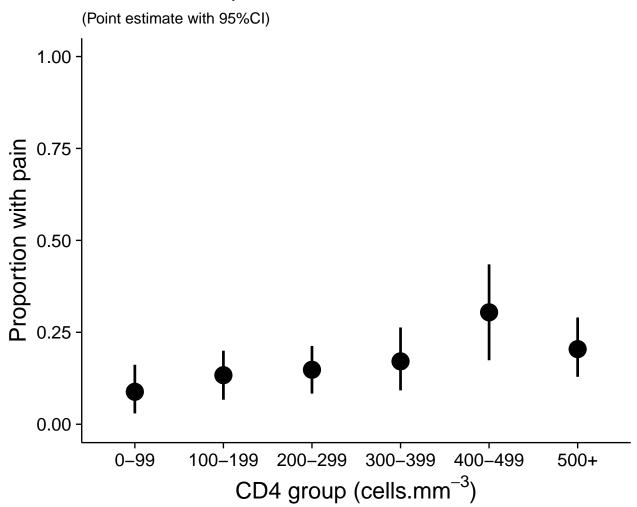


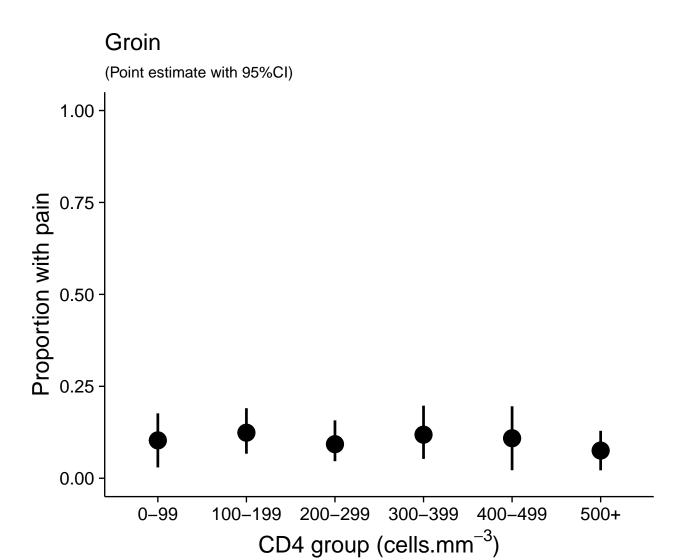


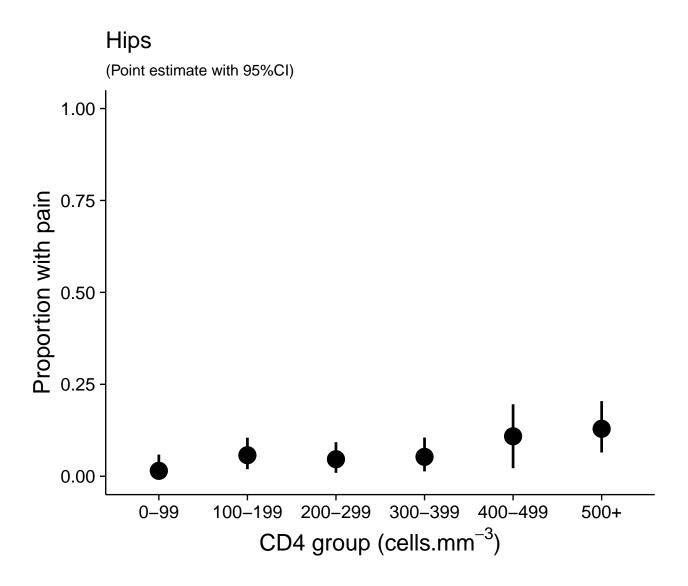


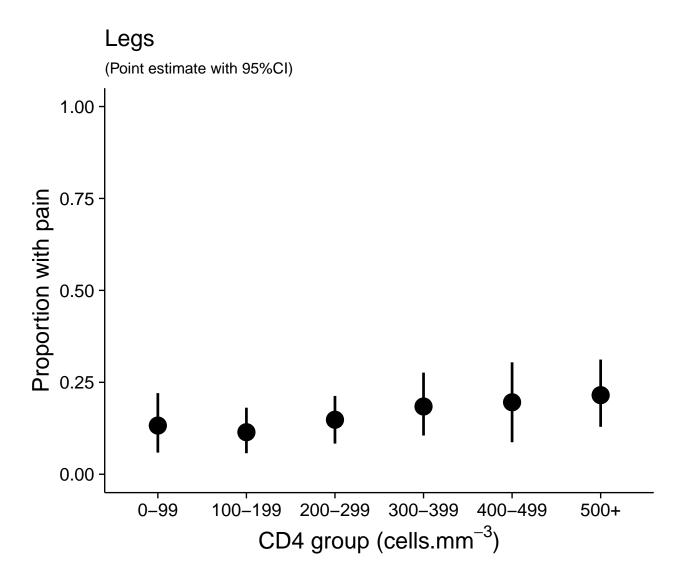


# Lumbosacral spine

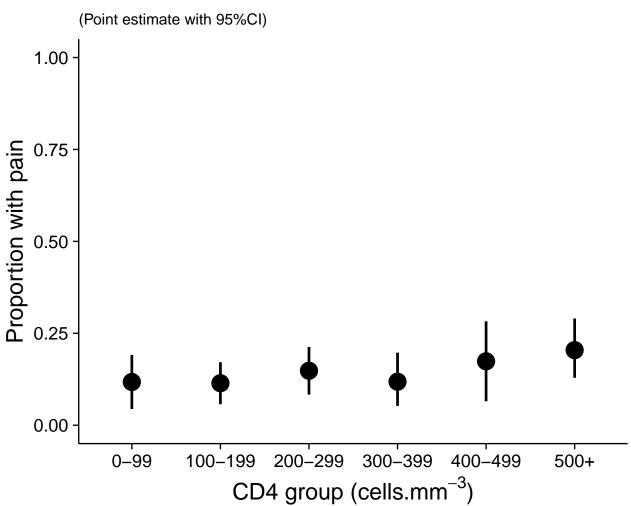


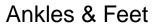


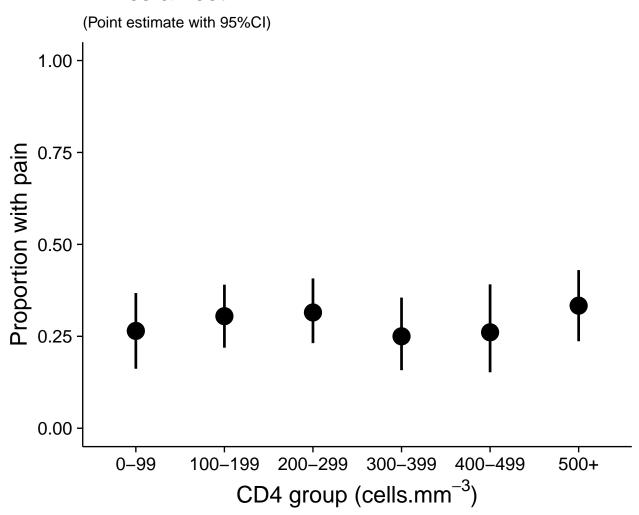




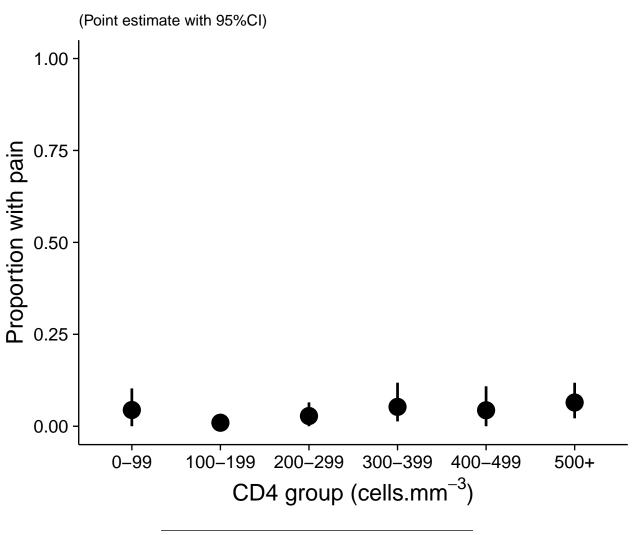












### Logistic regression (generalised linear mixed model)

Note: Age treated as a continuous variable, not like the categories used in the figures shwon above.

#### **Process data**

Extract cases with complete data for CD4 T-cell count, age, and sex.

```
# Process data and remove incomplete cases
data_glmm <- data %>%
    # Join with extracted demographics data (demo_reduced)
   left join(demo glmm) %>%
   # Remove upper back (only one outcome -- no pain) and ID
   select(-ID, -Upper_back) %>%
    # Extract complete cases
   filter(complete.cases(.))
# Convert data_glm to long format
data_glmm.long <- data_glmm %>%
   pivot_longer(cols = -c(CD4_recent, Age, Sex, Study_site),
                 names_to = 'Pain_site',
                 values_to = 'Pain_present')
# Extract vector of body sites where the proportion with pain was >= 10%
## Stops "Singular" fit warning when performing GLMM
high_prop <- data_glmm.long %>%
   group_by(Pain_site, Pain_present) %>%
   summarise(n = n()) \%
   mutate(total = sum(n),
           prop = n/total) %>%
   filter(Pain present == 'Yes') %>%
   filter(prop >= 0.10) %>%
    .$Pain site
# Filter for pain sites selected above
data_glmm.filtered <- data_glmm.long %>%
   filter(Pain_site %in% high_prop)
```

### Tabulate number of cases in original and GLM datasets

Table 54: Number of cases at each study site (original dataset)

Study site	Original	GLM
DD	60	42
NM	240	200
RESI	99	83
RPA	20	20
RPB	102	98
RPC	14	13
RPD	10	10
STIG	50	26

### Tabulate the pain sites removed

Sites removed because proportion of cases with pain at each site was < 0.1.

Table 55: Removed pain sites

```
Pain sites
Throat
Shoulder
Arms
Elbows
Wrists & Hands
Lower back
Cervical spine
Hips
Buttocks
```

#### **GLMM**

```
# Perform GLMM on each remaining body site glmm_mods <- data_glmm.filtered %>%
```

```
# Scale Age and CD4_recent
   mutate_if(is.numeric, scale) %>%
    # Dummy code outcome variable
   mutate(Pain_present = ifelse(Pain_present == 'No',
                                 yes = 0,
                                 no = 1)) %>%
    # Group and then nest by Pain_site
   group_by(Pain_site) %>%
   nest() %>%
    # Perform logistic regression
   mutate(logistic = map(.x = data,
                          ~ glmer(Pain_present ~
                                      Age +
                                      Sex +
                                      CD4_recent +
                                      (1|Study_site),
                                data = .x,
                                family = binomial(),
                                control = glmerControl(optimizer =
                                                            c('bobyqa',
                                                              'bobyqa'))))) %>%
    # Extract effect sizes and p-values
   mutate(OR = map(.x = logistic,
                    ~ exp(fixef(.x))),
           CI = map(.x = logistic,
                          ~ exp(confint.merMod(.x,
                                               method = 'Wald')[2:5, ])),
           p_value = map(.x = logistic,
                         ~ coef(summary(.x))[, 4])) %>%
    # Bind data into a dataframe and tabulate,
   mutate(df = pmap(.1 = list(OR, CI, p_value, Pain_site),
                     ~ cbind(..1, ..2, ..3) %>%
                         kable(caption = ..4,
                               col.names = c('OR', 'Wald lower 95%CI',
                                              'Wald upper 95%CI', 'p-value'))))
# Print OR for fixed effects with Wald-type 95% CIs and p-values
walk(glmm_mods$df, ~print(.x))
```

Table 56: Head

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.6025967	0.4111281	0.8832352	0.0094192
Age	0.9603824	0.7571667	1.2181392	0.7389474
SexMale	0.2483928	0.1168182	0.5281623	0.0002964
CD4_recent	0.7781325	0.5417471	1.1176622	0.1745119

Table 57: Chest

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.1758590	0.1089509	0.2838563	0.0000000
Age	1.0270898	0.7939823	1.3286360	0.8387355

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
SexMale	0.9685384	0.5232464	1.7927819	0.0.0000
CD4_recent	0.6066731	0.3740876	0.9838663	

Table 58: Abdomen

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.3934898	0.3105894	0.4985173	0.0000000
Age	0.8090008	0.6496259	1.0074758	0.0583007
SexMale	0.7543714	0.4217431	1.3493432	0.3420755
CD4_recent	1.0881886	0.8936920	1.3250140	0.4002194

Table 59: Thoracic\_spine

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.1734972	0.1289240	0.2334809	0.0000000
Age	0.9735667	0.7334718	1.2922543	0.8529032
SexMale	0.4232540	0.1665901	1.0753582	0.0707238
CD4_recent	1.1116459	0.8966374	1.3782122	0.3344899

Table 60: Lumbosacral\_spine

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.2151228	0.1511716	0.3061278	0.0000000
Age	1.0538056	0.8202668	1.3538355	0.6818089
SexMale	0.9184730	0.4740011	1.7797271	0.8010620
CD4_recent	1.0644237	0.8679756	1.3053338	0.5486620

Table 61: Groin

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.0487589	0.0179773	0.1322464	0.0000000
Age	0.9692042	0.7021360	1.3378559	0.8491598
SexMale	0.5147107	0.2176283	1.2173374	0.1304876
CD4_recent	1.0812723	0.8215002	1.4231888	0.5772630

Table 62: Legs

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.1715287	0.1164429	0.2526742	0.0000000
Age	1.2124724	0.9419946	1.5606134	0.1346604
SexMale	1.8610046	1.0339894	3.3494909	0.0383181
CD4_recent	1.0060419	0.8130468	1.2448486	0.9557953

Table 63: Knees

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.1580291	0.1025172	0.2436002	0.0000000
Age	1.3641246	1.0557327	1.7626015	0.0175612
SexMale	1.5342243	0.8263945	2.8483301	0.1751272
CD4_recent	0.9808591	0.7710007	1.2478388	0.8749729

Table 64: Ankles.Feet

	OR	Wald lower 95%CI	Wald upper 95%CI	p-value
(Intercept)	0.3801689	0.2550638	0.566636	0.0000020
Age	1.3849495	1.1025712	1.739647	0.0051216
SexMale	1.4774529	0.8761858	2.491329	0.1431536
CD4_recent	0.8825650	0.6731476	1.157132	0.3660350

### **Session information**

#### sessionInfo()

```
## R version 3.6.3 (2020-02-29)
## Platform: x86_64-apple-darwin15.6.0 (64-bit)
## Running under: macOS Catalina 10.15.4
##
## Matrix products: default
## BLAS: /Library/Frameworks/R.framework/Versions/3.6/Resources/lib/libRblas.0.dylib
## LAPACK: /Library/Frameworks/R.framework/Versions/3.6/Resources/lib/libRlapack.dylib
##
## locale:
## [1] en_US.UTF-8/en_US.UTF-8/en_US.UTF-8/C/en_US.UTF-8/en_US.UTF-8
## attached base packages:
## [1] stats
                graphics grDevices utils
                                              datasets methods
                                                                  base
##
## other attached packages:
## [1] lme4 1.1-21
                          Matrix 1.2-18
                                             boot 1.3-24
                                                                knitr 1.28
## [5] skimr_2.1
                          forcats_0.5.0
                                             stringr_1.4.0
                                                                dplyr_0.8.5
## [9] purrr_0.3.3
                          readr_1.3.1
                                             tidyr_1.0.2
                                                                tibble_3.0.0
## [13] ggplot2_3.3.0.9000 tidyverse_1.3.0
## loaded via a namespace (and not attached):
## [1] Rcpp_1.0.4
                        lubridate_1.7.4 lattice_0.20-38 utf8_1.1.4
## [5] assertthat_0.2.1 digest_0.6.25
                                         R6_2.4.1
                                                          cellranger_1.1.0
## [9] repr_1.1.0
                        backports_1.1.5 reprex_0.3.0
                                                          evaluate_0.14
## [13] highr_0.8
                        httr_1.4.1
                                         pillar_1.4.3
                                                          rlang_0.4.5
## [17] readxl 1.3.1
                        rstudioapi_0.11 minqa_1.2.4
                                                          nloptr 1.2.2.1
## [21] rmarkdown_2.1
                        labeling_0.3
                                          splines_3.6.3
                                                          munsell_0.5.0
```

## ## ## ## ##	[29] [33] [37] [41] [45] [49]	broom_0.5.5 pkgconfig_2.0.3 fansi_0.4.1 MASS_7.3-51.5 gtable_0.3.0 scales_1.1.0 fs_1.3.1	compiler_3.6.3 base64enc_0.1-3 crayon_1.3.4 grid_3.6.3 lifecycle_0.2.0 cli_2.0.2 xml2_1.3.0	modelr_0.1.6 htmltools_0.4.0 dbplyr_1.4.2 nlme_3.1-145 DBI_1.1.0 stringi_1.4.6 ellipsis_0.3.0	xfun_0.12 tidyselect_1.0.0 withr_2.1.2 jsonlite_1.6.1 magrittr_1.5 farver_2.0.3 generics_0.0.2
## ##	[53] [57]	vctrs_0.2.4 parallel_3.6.3 haven_2.2.0	tools_3.6.3 yaml_2.2.1	glue_1.3.2 colorspace_1.4-1	hms_0.5.3