

## Supplemental Material

First trimester maternal exposures to endocrine disrupting chemicals and metals and fetal size in the Michigan Mother Infant Pairs study

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Table S1. Sex-Stratified Anthropometric Measures

|                                  | <b>Females (n=29)</b> |               | <b>Males (n=27)</b> |               |
|----------------------------------|-----------------------|---------------|---------------------|---------------|
|                                  | <i>Median</i>         | <i>Range</i>  | <i>Median</i>       | <i>Range</i>  |
| Birthweight (g)                  | 3410                  | (2805 - 4070) | 3640                | (2860 - 4685) |
| Fenton z-score % tile            | 54.4                  | (6.6 - 85.2)  | 62.7                | (12.8 - 95.1) |
| Fetal Anthropometry <sup>a</sup> |                       |               |                     |               |
| Abdominal circ.                  | 137                   | (111 – 161)   | 140                 | (127 – 242)   |
| Biparietal diameter              | 43.4                  | (36 – 48)     | 44                  | (40 – 70.4)   |
| Femur length                     | 29.6                  | (23 – 36)     | 29.4                | (24 – 51.4)   |
| Head circ. <sup>b</sup>          | 161                   | (135 – 182)   | 164                 | (155 – 265)   |

<sup>a</sup> Units= mm; <sup>b</sup> Circ: circumference

Table S2. Correlations of phthalate metabolites in first trimester maternal urine samples (n=56)

| Phthalate | MnBP   | MBzP        | mCINP       | MCOMHP      | MCPP        | MeCPP       | MEHHP       | MEHP        | MEOHP       | MEP         | MiBP        |
|-----------|--------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
| MnBP      |        | <b>0.79</b> | <b>0.65</b> | <b>0.79</b> | <b>0.68</b> | <b>0.78</b> | <b>0.77</b> | <b>0.74</b> | <b>0.79</b> | <b>0.54</b> | <b>0.81</b> |
| MBzP      | <0.001 |             | <b>0.59</b> | <b>1.00</b> | <b>0.58</b> | <b>0.73</b> | <b>0.72</b> | <b>0.70</b> | <b>0.71</b> | <b>0.29</b> | <b>0.72</b> |
| mCINP     | <0.001 | <0.001      |             | <b>0.59</b> | <b>0.88</b> | <b>0.77</b> | <b>0.66</b> | <b>0.69</b> | <b>0.67</b> | <b>0.32</b> | <b>0.60</b> |
| MCOMHP    | <0.001 | <0.001      | <0.001      |             | <b>0.58</b> | <b>0.73</b> | <b>0.72</b> | <b>0.70</b> | <b>0.71</b> | <b>0.29</b> | <b>0.71</b> |
| MCPP      | <0.001 | <0.001      | <0.001      | <0.001      |             | <b>0.76</b> | <b>0.66</b> | <b>0.67</b> | <b>0.68</b> | <b>0.38</b> | <b>0.60</b> |
| MeCPP     | <0.001 | <0.001      | <0.001      | <0.001      | <0.001      |             | <b>0.92</b> | <b>0.85</b> | <b>0.92</b> | <b>0.40</b> | <b>0.76</b> |
| MEHHP     | <0.001 | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      |             | <b>0.91</b> | <b>0.99</b> | <b>0.45</b> | <b>0.75</b> |
| MEHP      | <0.001 | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      |             | <b>0.91</b> | <b>0.34</b> | <b>0.68</b> |
| MEOHP     | <0.001 | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      |             | <b>0.48</b> | <b>0.76</b> |
| MEP       | <0.001 | 0.03        | 0.02        | 0.03        | 0.004       | 0.002       | 0.001       | 0.01        | <0.001      |             | <b>0.48</b> |
| MiBP      | <0.001 | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | <0.001      | 0.001       |             |

Above diagonal denotes spearman correlation; Below diagonal denotes p-value associated with correlation value

Table S3. Correlations of phenols in first trimester maternal urine samples (n=56)

| Phenol  | 2,4-DCP | 2,5-DCP     | BP-3        | BPA         | BPF         | BPS         | BuPB  | EtPb        | MePB        | PrPB        | TCS         |
|---------|---------|-------------|-------------|-------------|-------------|-------------|-------|-------------|-------------|-------------|-------------|
| 2,4-DCP |         | <b>0.64</b> | <b>0.43</b> | <b>0.65</b> | <b>0.30</b> | <b>0.5</b>  | 0.04  | <b>0.50</b> | <b>0.46</b> | 0.19        | <b>0.60</b> |
| 2,5-DCP | <0.001  |             | <b>0.29</b> | <b>0.55</b> | 0.20        | <b>0.33</b> | 0.04  | <b>0.31</b> | <b>0.31</b> | 0.04        | <b>0.44</b> |
| BP-3    | 0.001   | 0.03        |             | <b>0.30</b> | <b>0.30</b> | <b>0.26</b> | 0.06  | <b>0.44</b> | <b>0.43</b> | <b>0.33</b> | <b>0.44</b> |
| BPA     | <0.001  | <0.001      | 0.02        |             | <b>0.31</b> | <b>0.46</b> | 0.08  | <b>0.41</b> | <b>0.36</b> | 0.13        | 0.40        |
| BPF     | 0.03    | 0.14        | 0.03        | 0.02        |             | 0.12        | -0.06 | <b>0.40</b> | 0.19        | 0.24        | 0.19        |
| BPS     | <0.001  | 0.01        | 0.05        | <0.001      | 0.39        |             | 0.11  | 0.17        | <b>0.51</b> | <b>0.35</b> | <b>0.32</b> |
| BuPB    | 0.79    | 0.75        | 0.67        | 0.55        | 0.64        | 0.41        |       | 0.15        | 0.05        | 0.07        | -0.09       |
| EtPB    | <0.001  | 0.02        | 0.001       | 0.002       | 0.003       | 0.22        | 0.28  |             | <b>0.44</b> | <b>0.44</b> | <b>0.37</b> |
| MePB    | 0.003   | 0.02        | 0.001       | 0.01        | 0.15        | <0.001      | 0.72  | 0.001       |             | 0.81        | 0.45        |
| PrPB    | 0.16    | 0.79        | 0.01        | 0.32        | 0.08        | 0.01        | 0.59  | 0.001       | <0.001      |             | 0.21        |
| TCS     | <0.001  | 0.001       | 0.001       | 0.002       | 0.15        | 0.02        | 0.53  | 0.01        | 0.001       | 0.12        |             |

Above diagonal denotes spearman correlation; Below diagonal denotes p-value associated with correlation value



Table S5. Distribution of summed chemicals among MMIP women (n=56)

| $\Sigma$ score      | N <sup>a</sup> | Mean  | SD    | Min | P10 | P25 | P50  | P75 | P95 | P99 | Max |
|---------------------|----------------|-------|-------|-----|-----|-----|------|-----|-----|-----|-----|
| Phthalates          | 12             | 12.18 | 7.38  | 1   | 1   | 6   | 11   | 19  | 23  | 23  | 23  |
| Phenols             | 12             | 12.45 | 5.31  | 3   | 6   | 8   | 12   | 17  | 22  | 22  | 22  |
| Metals <sup>b</sup> | 17             | 17.15 | 8.58  | 3   | 6   | 10  | 18   | 25  | 30  | 32  | 32  |
| Total <sup>b</sup>  | 41             | 41.46 | 19.71 | 7   | 16  | 24  | 39.5 | 62  | 71  | 75  | 75  |

<sup>a</sup>N=number of chemicals/elements measured; <sup>b</sup>n=55 women

Concentration ranks assigned were 0, 1, and 2; Maximum possible score for phthalates and metals = 24, metals = 34, and total = 82

Table S6. Bivariate associations of phthalate metabolites and survey responses among MMIP women

|                   | Metabolite   |   |       |   |        |   |              |   |             |   |             |   |       |   |      |   |       |   |        |             |
|-------------------|--------------|---|-------|---|--------|---|--------------|---|-------------|---|-------------|---|-------|---|------|---|-------|---|--------|-------------|
|                   | MnBP         |   | MBzP  |   | MCOMHP |   | mCINP        |   | MCPP        |   | meCPP       |   | MEHHP |   | MEHP |   | MEOHP |   | MEP    |             |
|                   | M            | p | M     | p | M      | p | M            | p | M           | p | M           | p | M     | p | M    | p | M     | p | M      | p           |
| P&C <sup>a</sup>  |              |   |       |   |        |   |              |   |             |   |             |   |       |   |      |   |       |   |        |             |
|                   | 0.14         |   | 0.76  |   | 0.75   |   | 0.90         |   | 0.50        |   | 0.50        |   | 0.26  |   | 0.23 |   | 0.18  |   | 0.10   |             |
| No                | 1.55         |   | 7.52  |   | 7.51   |   | 2.70         |   | 4.07        |   | 8.94        |   | 8.27  |   | 2.01 |   | 3.92  |   | 22.46  |             |
| Yes               | 2.04         |   | 15.99 |   | 15.99  |   | 1.90         |   | 4.55        |   | 8.87        |   | 9.33  |   | 3.18 |   | 5.20  |   | 83.71  |             |
| Hair <sup>a</sup> | 0.42         |   | 0.06  |   | 0.06   |   | 0.31         |   | 0.09        |   | 0.12        |   | 0.08  |   | 0.08 |   | 0.08  |   | 0.42   |             |
| No                | 11.22        |   | 13.10 |   | 13.10  |   | 2.18         |   | 6.66        |   | 10.99       |   | 12.25 |   | 3.96 |   | 7.15  |   | 166.74 |             |
| Yes               | 11.60        |   | 14.31 |   | 14.30  |   | 2.07         |   | 3.77        |   | 8.24        |   | 8.12  |   | 2.58 |   | 4.22  |   | 40.09  |             |
| FF <sup>b</sup>   | <b>0.004</b> |   | 0.10  |   | 0.10   |   | <b>0.002</b> |   | <b>0.04</b> |   | <b>0.02</b> |   | 0.20  |   | 0.17 |   | 0.14  |   | 0.17   | <b>0.01</b> |
| No                | 17.51        |   | 15.90 |   | 15.90  |   | 1.44         |   | 2.51        |   | 6.15        |   | 7.49  |   | 2.50 |   | 3.80  |   | 79.30  |             |
| Yes               | 8.19         |   | 10.66 |   | 10.66  |   | 3.24         |   | 7.91        |   | 13.81       |   | 11.95 |   | 3.63 |   | 6.68  |   | 51.84  |             |
|                   |              |   |       |   |        |   |              |   |             |   |             |   |       |   |      |   |       |   |        |             |

<sup>a</sup> no (n=13) yes (n=43); <sup>b</sup> no (n=36) yes (n=20)

M: Median; p: p-value; P&amp;C: Perfume and Cosmetics; Hair: Haircare Products; FF: Fast Food; FV: Fresh Vegetables

Table S7. Bivariate associations for phenols and survey responses among MMIP women

|                    | Phenols |      |         |             |       |             |      |      |      |             |      |      |
|--------------------|---------|------|---------|-------------|-------|-------------|------|------|------|-------------|------|------|
|                    | 2,4-DCP |      | 2,5-DCP |             | BP-3  |             | BPA  |      | BPF  |             | BPS  |      |
|                    | M       | P    | M       | P           | M     | P           | M    | P    | M    | P           | M    | P    |
| Perfume & Cosmetic |         | 0.31 |         | 0.30        |       | 0.80        |      | 0.63 |      | 0.84        |      | 0.06 |
| No                 | 0.23    |      | 0.20    |             | 58.22 |             | 0.72 |      | 0.14 |             | 0.14 |      |
| Yes                | 0.33    |      | 0.29    |             | 49.64 |             | 0.83 |      | 0.14 |             | 0.14 |      |
| Hair products      |         | 0.57 |         | 0.35        |       | <b>0.01</b> |      | 0.34 |      | 0.75        |      | 0.82 |
| No                 | 0.24    |      | 0.22    |             | 51.24 |             | 0.70 |      | 0.14 |             | 0.14 |      |
| Yes                | 0.74    |      | 0.61    |             | 47.12 |             | 1.06 |      | 0.14 |             | 0.14 |      |
| Canned Food        |         | 0.66 |         | 0.50        |       | 0.60        |      | 0.57 |      | <b>0.03</b> |      | 0.76 |
| No                 | 0.25    |      | 0.22    |             | 39.52 |             | 0.83 |      | 0.14 |             | 0.14 |      |
| Yes                | 0.34    |      | 0.31    |             | 58.22 |             | 0.64 |      | 0.14 |             | 0.14 |      |
| Fast food          |         | 0.11 |         | <b>0.03</b> |       | 0.75        |      | 0.40 |      | 0.50        |      | 0.37 |
| No                 | 0.24    |      | 0.22    |             | 51.24 |             | 0.70 |      | 0.14 |             | 0.14 |      |
| Yes                | 0.74    |      | 0.61    |             | 47.12 |             | 1.06 |      | 0.14 |             | 0.14 |      |

Note: Equivalent medians may result from low detection rates

Table S7 cont'd. Bivariate associations for phenols and survey responses among MMIP women

|                    |     | Phenols |   |      |   |       |   |             |   |       |   |
|--------------------|-----|---------|---|------|---|-------|---|-------------|---|-------|---|
|                    |     | BuPB    |   | EtPb |   | MePB  |   | PrPB        |   | TCS   |   |
|                    |     | M       | P | M    | P | M     | P | M           | P | M     | P |
| Perfume & Cosmetic | No  | 0.28    |   | 0.69 |   | 0.06  |   | 0.24        |   | 0.64  |   |
|                    | Yes | 0.14    |   | 0.73 |   | 15.55 |   | 6.88        |   | 13.20 |   |
| Hair products      | No  | 0.14    |   | 0.42 |   | 93.60 |   | 15.32       |   | 17.05 |   |
|                    | Yes | 0.14    |   | 0.64 |   | 0.86  |   | 0.21        |   | 0.35  |   |
| Canned Food        | No  | 0.14    |   | 0.28 |   | 25.27 |   | 7.51        |   | 11.18 |   |
|                    | Yes | 0.14    |   | 2.31 |   | 98.38 |   | 22.16       |   | 28.95 |   |
| Fast food          | No  | 0.14    |   | 1.00 |   | 75.87 |   | 7.96        |   | 15.47 |   |
|                    | Yes | 0.14    |   | 0.48 |   | 53.06 |   | 15.32       |   | 13.74 |   |
|                    | No  | 0.14    |   | 0.57 |   | 0.08  |   | <b>0.03</b> |   | 0.08  |   |
|                    | Yes | 0.14    |   | 0.28 |   | 25.27 |   | 7.51        |   | 11.18 |   |
|                    | No  | 0.14    |   | 2.31 |   | 98.38 |   | 22.16       |   | 28.95 |   |

Table S8. Bivariate associations with bisphenol A, F, and S and canned food consumption (n=56)

|                         | Bisphenol |         |      |             |      |         |
|-------------------------|-----------|---------|------|-------------|------|---------|
|                         | BPA       |         | BPF  |             | BPS  |         |
|                         | M         | p-value | M    | p-value     | M    | p-value |
| Canned food consumption |           | 0.57    |      | <b>0.03</b> |      | 0.76    |
| No                      | 0.83      |         | 0.14 |             | 0.14 |         |
| Yes                     | 0.64      |         | 0.14 |             | 0.14 |         |
| Canned food frequency   |           | 0.21    |      | <b>0.02</b> |      | 0.58    |
| None                    | 1.20      |         | 0.32 |             | 0.25 |         |
| ≤ 1 serving/day         | 1.34      |         | 1.39 |             | 0.32 |         |
| 2-3 servings/day        | 4.58      |         | 6.52 |             | 0.26 |         |
| ≥ 4 servings/day        | -         |         | -    |             | -    |         |

M: Median

Table S9. Bivariate associations for metals and survey responses among MMIP women

|                    | Metals |      |    |      |    |      |    |      |    |       |    |      |    |      |
|--------------------|--------|------|----|------|----|------|----|------|----|-------|----|------|----|------|
|                    | As     |      | Ba |      | Cd |      | Cr |      | Cu |       | Hg |      | Mn |      |
|                    | M      | P    | M  | P    | M  | P    | M  | P    | M  | P     | M  | P    | M  | P    |
| Perfume & Cosmetic |        | 0.26 |    | 0.82 |    | 0.14 |    | 0.37 |    | 0.93  |    | 0.33 |    | 0.27 |
|                    | No     | 2.75 |    | 5.96 |    | 0.04 |    | 0.59 |    | 7.76  |    | 0.09 |    | 1.24 |
|                    | Yes    | 4.26 |    | 4.60 |    | 0.06 |    | 0.72 |    | 9.36  |    | 0.04 |    | 1.67 |
| Hair products      |        | 0.46 |    | 0.39 |    | 0.49 |    | 0.75 |    | 0.49  |    | 0.70 |    | 0.09 |
|                    | No     | 4.57 |    | 2.95 |    | 0.07 |    | 0.71 |    | 9.69  |    | 0.07 |    | 1.97 |
|                    | Yes    | 3.68 |    | 4.91 |    | 0.04 |    | 0.71 |    | 9.13  |    | 0.06 |    | 1.40 |
| Canned Food        |        | 0.21 |    | 0.48 |    | 0.41 |    | 0.57 |    | 0.79  |    | 0.08 |    | 0.85 |
|                    | No     | 3.66 |    | 4.60 |    | 0.04 |    | 0.69 |    | 7.66  |    | 0.04 |    | 1.54 |
|                    | Yes    | 4.47 |    | 4.73 |    | 0.51 |    | 0.72 |    | 10.40 |    | 0.08 |    | 1.54 |
| Fast food          |        | 0.40 |    | 0.28 |    | 0.02 |    | 0.34 |    | 0.07  |    | 0.64 |    | 0.15 |
|                    | No     | 3.68 |    | 4.10 |    | 0.04 |    | 0.68 |    | 7.33  |    | 0.04 |    | 1.28 |
|                    | Yes    | 4.47 |    | 4.82 |    | 0.13 |    | 0.83 |    | 11.83 |    | 0.06 |    | 1.74 |

M: Median; P: P-value

Table S9 cont'd. Bivariate associations for metals and survey responses among MMIP women

|                  | Metals |       |    |      |    |      |    |             |    |             |    |      |    |             |
|------------------|--------|-------|----|------|----|------|----|-------------|----|-------------|----|------|----|-------------|
|                  | Mo     |       | Ni |      | Pb |      | Se |             | Sn |             | Tl |      | Zn |             |
|                  | M      | P     | M  | P    | M  | P    | M  | P           | M  | P           | M  | P    | M  | P           |
| Perf. & Cosmetic |        | 0.71  |    | 0.23 |    | 0.21 |    | 0.28        |    | 0.28        |    | 0.28 |    | 0.61        |
|                  | No     | 50.13 |    | 2.73 |    | 0.13 |    | 33.82       |    | 0.19        |    | 0.15 |    | 171.04      |
|                  | Yes    | 47.27 |    | 4.80 |    | 0.25 |    | 51.85       |    | 0.38        |    | 0.19 |    | 235.48      |
| Hair products    |        | 0.13  |    | 0.55 |    | 0.42 |    | 0.29        |    | 0.58        |    | 0.11 |    | 0.52        |
|                  | No     | 68.30 |    | 4.94 |    | 0.23 |    | 52.82       |    | 0.45        |    | 0.20 |    | 200.82      |
|                  | Yes    | 40.47 |    | 3.91 |    | 0.21 |    | 44.51       |    | 0.27        |    | 0.17 |    | 178.03      |
| Canned Food      |        | 0.06  |    | 0.48 |    | 0.11 |    | 0.30        |    | 0.09        |    | 0.60 |    | 0.32        |
|                  | No     | 41.25 |    | 3.91 |    | 0.17 |    | 42.13       |    | 0.29        |    | 0.15 |    | 169.82      |
|                  | Yes    | 61.00 |    | 5.01 |    | 0.28 |    | 50.98       |    | 0.39        |    | 0.20 |    | 254.41      |
| Fast food        |        | 0.07  |    | 0.06 |    | 0.31 |    | <b>0.01</b> |    | <b>0.02</b> |    | 0.22 |    | <b>0.02</b> |
|                  | No     | 40.47 |    | 3.16 |    | 0.17 |    | 36.05       |    | 0.18        |    | 0.15 |    | 170.65      |
|                  | Yes    | 81.86 |    | 4.92 |    | 0.28 |    | 66.06       |    | 0.50        |    | 0.19 |    | 332.71      |

M: Median; P: P-value

Table S10. Associations between Bisphenol F and S dichotomized by detected vs. not detected and birth outcomes (n=56)

| Phenol | Birth outcomes                |                          |             |                             |                       |             |
|--------|-------------------------------|--------------------------|-------------|-----------------------------|-----------------------|-------------|
|        | Birth weight (g) <sup>a</sup> |                          |             | Fenton z-score <sup>b</sup> |                       |             |
|        | $\beta$                       | (95% CI)                 | p-value     | B                           | (95% CI)              | p-value     |
| BPF    | -7.88                         | (-270.32, 254.56)        | 0.95        | -0.18                       | (-0.73, 0.37)         | 0.52        |
| BPS    | <b>-220.10</b>                | <b>(-424.94, -15.26)</b> | <b>0.04</b> | <b>-0.51</b>                | <b>(-0.94, -0.07)</b> | <b>0.02</b> |

<sup>a</sup> Adjusted for Specific gravity, gestational age, and gender; <sup>b</sup> Adjusted for Specific gravity;