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An ASA a day when a baby's on the way?

CLINICAL QUESTION

Is acetylsalicylic acid (ASA) effective in preventing complications in pregnant women at risk of preeclampsia?

BOTTOM LINE

In women at risk for preeclampsia at ~12-28 weeks gestation, low-dose ASA (50-150mg) reduces risk of preeclampsia by an absolute ~2%, perinatal death by ~0.5%, and preterm birth by ~2% compared to placebo. The risk of postpartum hemorrhage is increased by up to ~1%.

EVIDENCE

- 7 systematic reviews (17-77 randomized controlled trials [RCTs]; 26,952-46,568 patients) from the last 5 years comparing ASA to placebo in pregnant women at varying preeclampsia risk.¹⁻⁷ ASA usually initiated ~12-28 weeks, continued until delivery. Results statistically significant unless indicated.
 - Maternal Outcomes:
 - Preeclampsia: 5 systematic reviews (16-60 RCTs):¹⁻⁵
 - 4.5-9.6% versus 5.8-11.8% (placebo), number needed to treat (NNT)=31-72.
 - Placental abruption: 3 systematic reviews (9-29 RCTs):^{1,3,4}
 - 0.9-1.3% versus 0.7-1.2% (placebo) (not statistically different).
 - Postpartum hemorrhage (>500-1000mL blood loss): 4 systematic reviews (9-19 RCTs):^{1,3,4,6}

- 3.7-15.2% versus 3.3-14.3% (placebo), number needed to harm (NNH)=97-239 (1/4 systematic reviews not statistically different).⁴
- o Fetal outcomes:
 - Perinatal death: 3 systematic reviews (11-52 RCTs):^{1,3,4}
 - 2.1-3.1% versus 2.7-3.5% (placebo), NNT=179-239.
 - Preterm delivery/birth: 2 systematic reviews with comprehensive data (18-47 RCTs):^{1,3}
 - 15.9-16.6% versus 17.5-18.5% (placebo), NNT=54-64.
 - Fetal intracranial bleed: 1 systematic review (6 RCTs):⁴
 - Not statistically different.
- Limitations: Inconsistent definitions of patients at risk for preeclampsia; infrequent reporting of serious maternal outcomes (examples: eclampsia, death); some large RCTs not included in all systematic reviews.

CONTEXT

- No clear difference in outcomes between 50-150 mg daily.^{1,3-5,7}
- Earlier initiation (<16-20 weeks) may enhance preeclampsia benefit based on subgroup analyses. No consistent trends for other outcomes.^{1-4,7}
- Sensitivity of clinical risk factors for predicting pre-eclampsia is <40%.⁸
- Guidelines vary:
 - Common recommendations among guidelines for ASA use include, but not limited to:
 - Any high-risk factors (examples: prior preeclampsia, chronic hypertension, renal or autoimmune disease, diabetes) or,
 - At least 2 moderate-risk factors (examples: nulliparity, age >35-40, previous adverse pregnancy outcome).
 - o Canadian: ASA 81-162mg daily preferably before 16 weeks until 36 weeks gestation.⁸
 - American: ASA 81mg daily initiated between 12-28 weeks gestation (optimally before 16 weeks) until delivery.⁹

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