

Methodology of Disproportionality Analysis

	Drug-related ADEs	Non-drug-related ADEs	Total
Drug	a	b	a + b
Non-drug	c	d	c + d
Total	a + c	b + d	N = a + b + c + d

Method	Formula	Threshold
ROR	$ROR = \frac{a / c}{b / d}$	$a \geq 3$ $ROR \geq 3$ 95%CI (lower limit) > 1
	$SE(\ln ROR) = \sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}$	
	$95\%CI = e^{\ln(ROR) \pm 1.96se}$	
PRR	$PRR = \frac{a / (a + b)}{c / (c + d)}$	$a \geq 3$ $PRR \geq 2$ 95%CI (lower limit) > 1
	$SE(\ln PRR) = \sqrt{\frac{1}{a} - \frac{1}{a + b} + \frac{1}{c} - \frac{1}{c + d}}$	
	$95\%CI = e^{\ln(PRR) \pm 1.96se}$	
BCPNN	$IC = \log_2 \frac{p(x, y)}{p(x)p(y)} = \log_2 \frac{a(a + b + c + d)}{(a + b)(a + c)}$	IC025>0
	$E(IC)$ $= \log_2 \frac{(a + \gamma 11)(a + b + c + d + \alpha)(a + b + c + d + \beta)}{(a + b + c + d + \gamma)(a + b + \alpha 1)(a + c + \beta 1)}$	
	$V(IC) = \frac{1}{(\ln 2)^2} \left[\frac{(a + b + c + d) - a + \gamma - \gamma 11}{(a + \gamma 11)(1 + a + b + c + d + \gamma)} \right.$ $\quad + \frac{(a + b + c + d) - (a + b) + a - \alpha 1}{(a + b + \alpha 1)(1 + a + b + c + d + \alpha)}$ $\quad \left. + \frac{(a + b + c + d + \alpha) - (a + c) + \beta - \beta 1}{(a + b + \beta 1)(1 + a + b + c + d + \beta)} \right]$	
	$\gamma = \gamma 11 \frac{(a + b + c + d + \alpha)(a + b + c + d + \beta)}{(a + b + \alpha 1)(a + c + \beta 1)}$	
	$IC - 2SD = E(IC) - 2\sqrt{V(IC)}$	
EBGM	$EBGM = \frac{a(a + b + c + d)}{(a + c)(a + b)}$	EBGM05>2
	$SE(\ln EBGM) = \sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}}$	
	$95\%CI = e^{\ln(EBGM) \pm 1.96se}$	