

## ABSTRAK

**Latar belakang:** Sebanyak 930.000 kematian kanker kolorektal sering disebabkan oleh terlambatnya diagnosis. Akibatnya, pilihan terapi terbatas, resistensi dan rekurensi meningkat. Sel punca kanker kolorektal (SP-KKR) berkontribusi dalam kemoresistensi yang ditandai oleh CD133 dan CD44, sehingga diperlukan pendekatan terapi alami seperti *Phyllanthus niruri linn* (PNL) sebagai imunomodulator dalam mengeradikasi SP-KKR yang dapat dianalisis melalui jalur perantara TLR, NF- $\kappa$ B, dan CTL.

**Metode:** Penelitian *true experimental* dengan *posttest only control group design* ini menggunakan tikus *Sprague-Dawley* jantan yang terbagi menjadi lima kelompok yaitu kontrol normal (Kn), kontrol negatif (K-), kelompok perlakuan *capecitabine* (K+), PNL (X1), dan kombinasi PNL dan *capecitabine* (X2). PNL adalah variabel bebas, variabel terikat adalah CD133 dan CD44, dan variabel perantara adalah TLR, NF- $\kappa$ B, dan CTL pada kolon. Data dianalisis menggunakan SPSS v22.

**Hasil:** Rasio berat kolon sekum dan berat tikus memiliki perbedaan signifikan ( $p < 0,001$ ); K- dan X2 memiliki rasio lebih rendah dibanding Kn. Uji *Kruskal Wallis* menunjukkan ekspresi CD133 ( $p = 0,008$ ), CD44 ( $p = 0,004$ ), CTL ( $p = 0,01$ ), TLR ( $p = 0,212$ ) dan NF- $\kappa$ B ( $p = 0,361$ ). Ekspresi CD133 terendah pada X2. CD44 lebih rendah pada X1 dan X2. CTL lebih tinggi pada X1 dan X2.

**Simpulan:** Ekstrak PNL baik tunggal maupun kombinasi secara sinergis terbukti lebih efektif mengeradikasi SP-KKR dibanding *capecitabine* tunggal.

**Kata Kunci:** kanker kolorektal, sel punca kanker, CD133, CD44, TLR, NF- $\kappa$ B, CTL, *phyllanthus niruri linn*, *capecitabine*.

## ABSTRACT

**Background:** Up to 930,000 colorectal cancer deaths are caused by a late diagnosis, which limits therapeutic options and increases resistance and recurrence. CD133 and CD44-associated chemoresistance is facilitated by colorectal cancer stem cells (CRC-SCs). *Phyllanthus niruri* linn (PNL), a natural therapeutic approach, is an immunomodulator that can be analyzed through TLR, NF- $\kappa$ B, and CTL pathways, potentially eradicating CRC-SCs.

**Method:** The study was used a true experimental method with a posttest only control group design. Male Sprague-Dawley mice were separated into five groups normal control (Kn), negative control (K-), capecitabine treatment group (K+), PNL (X1), and a combination of PNL and capecitabine (X2). PNL was an independent variable, whereas CD133 and CD44 were dependent variables. The intermediate variables included in this study were TLR, NF- $\kappa$ B, and CTL, which were specifically related to the colon. The data was analyzed using SPSS version 22

**Results:** A significant difference ( $p < 0.001$ ) in colon cecum weight and mouse weight ratio; K- and X2 were have a lower ratio than Kn. The Kruskal-Wallis test was revealed significant expression of CD133 ( $p = 0.008$ ), CD44 ( $p = 0.004$ ), CTL ( $p = 0.01$ ), TLR ( $p = 0.212$ ), and NF- $\kappa$ B ( $p = 0.361$ ). CD133 expression was lowest in X2. CD44 was lower in X1 and X2. CTL was higher in X1 and X2.

**Conclusion:** PNL extract, in a synergistic combination or on its own, is more effective than capecitabine in eliminating CRC-SCs.

**Keywords:** colorectal cancer, cancer stem cells, CD133, CD44, TLR, NF- $\kappa$ B, CTL, *Phyllanthus niruri* Linn, capecitabine.