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## (12) United States Patent

Papadopoulos et al.

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### (54) MODIFIED CHIMERIC POLYPEPTIDES WITH IMPROVED PHARMACOKINETIC PROPERTIES

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(\*) Notice: Subject to any disclaimer, the term of this

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§ 371 (c)(1),

(2), (4) Date: Dec. 6, 2001

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  C07H 21/04 (2006.01)

  C07K 16/00 (2006.01)

  C07K 14/00 (2006.01)

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### (57) ABSTRACT

Modified chimeric polypeptides with improved pharmacokinetics are disclosed. Specifically, modified chimeric Flt1 receptor polypeptides that have been modified in such a way as to improve their pharmacokinetic profile are disclosed. Also disclosed are methods of making and using the modified polypeptides including but not limited to using the modified polypeptides to decrease or inhibit plasma leakage and/or vascular permeability in a mammal.

15 Claims, 55 Drawing Sheets



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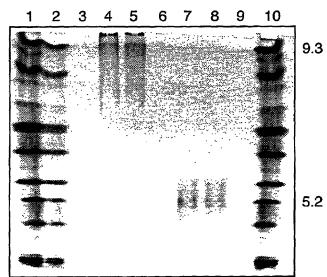
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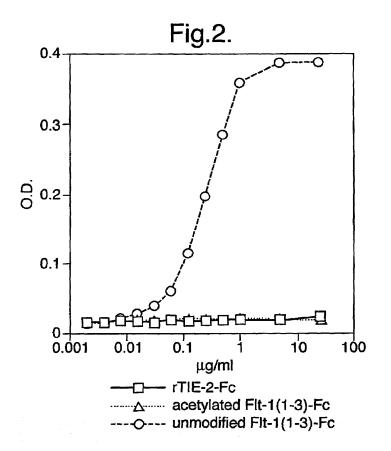


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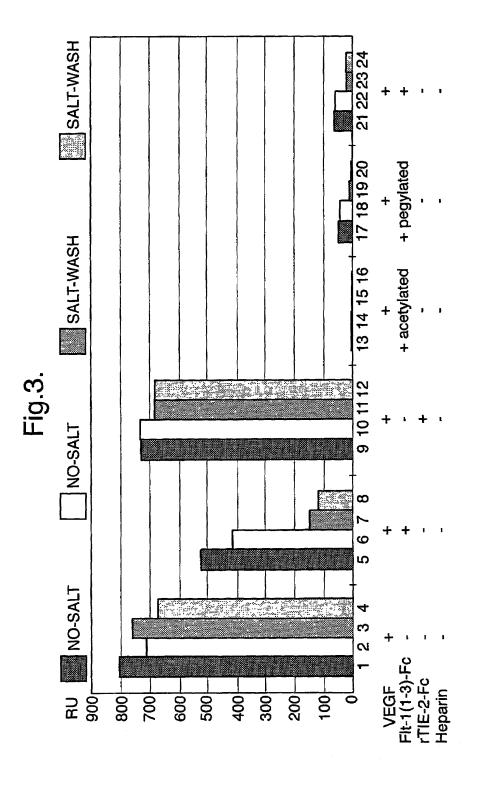
Fig.1.

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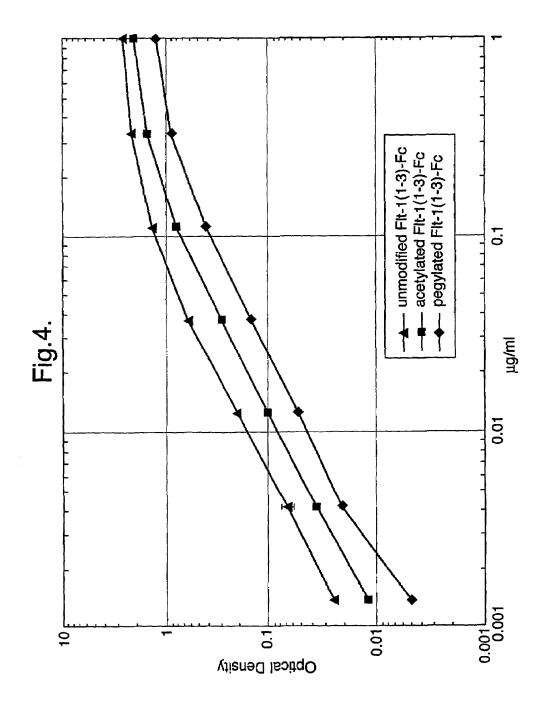


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