

group, a substituted or unsubstituted heterocyclic group and a halogen atom and  $P(O)(R_{13})_2$ ,  $S(O)R_{13}$ ,  $S(O_2)R_{13}$  and  $C(O)R_{13}$ , wherein  $R_{13}$  is a halogen atom.

26. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein the group other than a leaving group represented by  $R_6$ ,  $R_7$ ,  $R_8$  and  $R_9$  is selected from the group consisting of a hydrogen atom, a  $C_1$ - $C_{12}$  alkyl group, an aryl group, and a heterocyclic group.

27. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 26, wherein  $R_6$ ,  $R_7$ ,  $R_8$  and  $R_9$  are selected from the group consisting of  $C_2H_5$ , a phenyl group, or an imidazolyl group.

28. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein the leaving group represented by  $R_1$ ,  $R_2$ ,  $R_3$ ,  $R_4$ ,  $R_6$ ,  $R_7$ ,  $R_8$  and  $R_9$  is selected from the group consisting of a halogen atom and  $OR_{11}$ , wherein  $R_{11}$  is selected from the group consisting of a hydrogen atom, a substituted or unsubstituted  $C_1$ - $C_{12}$  alkyl group, a substituted or unsubstituted mono- or bicyclic aryl group and a substituted or unsubstituted heterocyclic group.

29. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 28, wherein  $R_{11}$  is selected from the group consisting of  $CH_3$ ,  $C_2H_5$ ,  $CH_2C_6H_5$ , and  $C_6H_5$ .

30. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein  $X$  is selected from the group consisting of  $SO_2$ ,  $SO$ ,  $P(O)(OR_{11})$ ,  $CO$ ,  $Se$ ,  $Te$  and  $Sr$ , wherein  $R_{11}$  is selected from the group consisting of a hydrogen atom, a substituted or unsubstituted  $C_1$ - $C_{12}$  alkyl group, a substituted or unsubstituted mono- or bicyclic aryl group and a substituted or unsubstituted heterocyclic group.

31. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 30, wherein  $X$  is selected from the group consisting of  $SO_2$ ,  $SO$ ,  $P(O)(OR_{11})$  and  $CO$ , wherein  $R_{11}$  is as defined above.

32. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein  $n=0$  to 2,  $n=0$  to 2, and  $l=0$  to 2.

33. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said reagent is selected from the group consisting of ethyl-2,2'-sulfonyl-bis-cyanoacetate, diethyl-2,2'-sulfonyl-bis-malonate, ethyl-2,2'-sulfonyl-2,2'-benzene sulfonyl-bis-acetate, 2,2'-sulfonyl-2,2'-imidazole-N-sulfonyl-bis-acetonitrile, methyl 2,2'-sulfonyl-bis-cyanoacetate, 2,2'-methoxymethylene-2,2'-sulfonyl-bis-acetonitrile and dimethyl 2,2'-methoxyphosphinylidene-2,2'-methoxymethylene-bis-methanephosphonate.

34. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said stroma-free hemoglobin is selected from the group consisting of human, equine, porcine, ovine, bovine, simian and fish hemoglobin.

35. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said stroma-free hemoglobin is bovine hemoglobin.

36. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said stroma-free hemoglobin is human hemoglobin.

37. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said cross-linking is carried out in the presence of oxygen.

38. The method of producing cross-linked stroma-free hemoglobin as claimed in claim 20, wherein said cross-linking is carried out in the absence of oxygen.

39. The method of producing cross-linked hemoglobin as claimed in claim 20, wherein said cross-linking is carried out at about  $10^\circ C.$  to  $37^\circ C.$

40. The method of producing cross-linked hemoglobin as claimed in claim 39 wherein said cross-linking is carried out at about  $25^\circ C.$  to  $30^\circ C.$

41. The method of producing cross-linked hemoglobin as claimed in claim 20, wherein the molar ratio of said reagent to said hemoglobin is 1:1 to 1:10.

42. The method of producing cross-linked hemoglobin as claimed in claim 41, wherein the molar ratio of said reagent to said hemoglobin is 1:2 to 1:6.

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