

[0026] For instance, in some examples, transferring the acidified pool from the first vessel to the second vessel based at least in part on receiving a signal indicating that the second vessel is empty.

[0027] Additionally, in some examples, filling the second vessel with the equilibration buffer is based at least in part on a receiving a signal indicating that the second vessel is empty.

[0028] Moreover, in some examples, the automated method of low pH viral inactivation may further include transferring the pool from the filter to a third vessel.

[0029] For instance, in some examples, transferring the pool from the filter to the third vessel is based at least in part on a receiving a signal indicating that the third vessel is empty.

[0030] Additionally, in some examples, the automated method of low pH viral inactivation may further include recalibrating the first pH probe responsive to the first alert. Similarly, in some examples, the automated method of low pH viral inactivation may further include recalibrating the second pH probe responsive to the second alert.

[0031] ~~In still another aspect, There is disclosed~~ a method for inactivating enveloped viruses during purification of a recombinant protein of interest is provided, the method comprising: obtaining a fluid known or suspected to contain at least one enveloped virus; subjecting the fluid to one or more of the following steps at a concentration and for a time sufficient to cause viral inactivation: adding the fluid to a first vessel; adding acid to the first vessel; measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel; ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel; transferring the fluid from the first vessel to a second vessel; filling the first vessel with an equilibration buffer having a known pH value; measuring, by the first pH probe, a second pH value associated with the first vessel; comparing the second measured pH value associated with the first vessel to the known pH value of the equilibration buffer; determining whether the second measured pH value associated with the first vessel is different from the known pH value of the equilibration buffer by greater than a threshold pH value; and generating a first alert responsive to the second measured pH value associated with the first vessel being different from the known pH value of the equilibration buffer by greater than the threshold pH value; and subjecting the neutralized viral inactivated fluid to at least one unit operation which includes at least a filtration step or a chromatography step.

[0032] In some examples, adding the fluid to the first vessel is based in part on receiving a signal indicating that the first vessel is empty.

[0033] Additionally, in some examples, transferring the fluid from the first vessel to the second vessel is based in part on receiving a signal indicating that the second vessel is empty.

[0034] Moreover, in some examples, filling the first vessel with the equilibration buffer is based in part on receiving a signal indicating that the first vessel is empty.

[0035] Furthermore, in some examples, the fluid comprises a recombinant protein of interest. Moreover, in some examples, the fluid is harvested host cell culture fluid. Additionally, in some examples, the fluid is from an effluent stream, eluate, pool, storage or hold from a unit operation comprising a harvest step, a filtration step or a chromatography step. Furthermore, in some examples, the fluid is eluate collected from depth filtration, microfiltration, affinity chromatography, ion exchange chromatography, multimodal

[0043] Furthermore, in some examples, the automated system for low pH viral inactivation may further include one or more additional pH probes associated with the first vessel and configured to measure the pH of contents of the first vessel. Similarly, in some examples, the automated system for low pH viral inactivation may further include one or more additional pH probes associated with the second vessel and configured to measure the pH of contents of the second vessel.

[0044] Moreover, in some examples, the acid is selected from formic acid, acetic acid, citric acid, and phosphoric acid at concentrations suitable to ensure viral inactivation. Furthermore, in some examples, the threshold pH for viral inactivation is from pH 2 to 4. Additionally, in some examples, the chromatography elution pool is exposed to the acid for less than 30 minutes prior to neutralization. Moreover, in some examples, the base is Tris base at a concentration of 2M. Furthermore, in some examples, the threshold range of neutral pH values is from pH 4.5 to 6. Additionally, in some examples, the low pH viral inactivation is conducted at a temperature of 5 to 25 °C.

[0045] Furthermore, in some examples, neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a holding vessel. For instance, in some examples, the neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a depth filter. Additionally, in some examples, following depth filtration, the neutralized viral inactivated eluate is transferred to a sterile filter. Moreover, in some examples, the neutralized viral inactivated chromatography elution pool from the second vessel is transferred a first polish chromatography column.

[0046] ~~In still another aspect, There is disclosed~~ an automated method of low pH viral inactivation is provided, the method comprising: adding a pool to a first vessel; adding acid to the first vessel; measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel; ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel; transferring the pool from the first vessel to a second vessel; adding base to the second vessel after an elapsed time after the transfer of the pool to the second vessel exceeds a threshold amount of time for reducing a concentration of virus in the pool to a predetermined safe level; measuring, by a second pH probe associated with the second vessel, a first pH value associated with the second vessel; ceasing, based on the first measured pH value associated with the second vessel being within a threshold range of neutral pH values, the addition of base to the second vessel; and transferring the pool from the second vessel to a filter for treatment of the neutralized viral inactivated pool.

[0047] In some examples, transferring the pool into the first vessel is based at least in part on receiving a signal indicating that the first vessel is empty.

[0048] Furthermore, in some examples, transferring the acidified pool from the first vessel to the second vessel based at least in part on receiving a signal indicating that the second vessel is empty.

[0049] Moreover, in some examples, the automated method of low pH viral inactivation may further include transferring the pool from the filter to a third vessel.

[0050] For instance, in some examples, transferring the pool from the filter to the third vessel is based at least in part on a receiving a signal indicating that the third vessel is empty.

[0051] Additionally, in some examples, the automated method of low pH viral inactivation may further include recalibrating the first pH probe responsive to the first alert. Similarly, in some examples, the automated method of low pH viral inactivation may further include recalibrating the second pH probe responsive to the second alert.

[0052] ~~In another aspect, There is disclosed~~ a method for inactivating enveloped viruses during purification of a recombinant protein of interest is provided, comprising: obtaining a fluid known or suspected to contain at least one enveloped virus; subjecting the fluid to one or more of the following steps at a concentration and for a time sufficient to cause viral inactivation: adding the fluid to a first vessel; adding acid to the first vessel; measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel; ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel; transferring the fluid from the first vessel to a second vessel; adding base to the second vessel; measuring, by a second pH probe associated with the first vessel, a second pH value associated with the second vessel; ceasing, based on the second measured pH value associated with the second vessel being within a tolerance band of a target pH value for neutrality, the addition of base to the second; and subjecting the neutralized viral inactivated fluid to at least one unit operation which includes at least a filtration step or a chromatography step.

[0053] In some examples, adding the fluid to the first vessel is based in part on receiving a signal indicating that the first vessel is empty.

[0054] Additionally, in some examples, transferring the fluid from the first vessel to the second vessel is based in part on receiving a signal indicating that the second vessel is empty.

[0055] Furthermore, in some examples, the fluid comprises a recombinant protein of interest. Moreover, in some examples, the fluid is harvested host cell culture fluid. Additionally, in some examples, the fluid is from an effluent stream, eluate, pool, storage or hold from a unit operation comprising a harvest step, a filtration step or a chromatography step. Furthermore, in some examples, the fluid is eluate collected from depth filtration, microfiltration, affinity chromatography, ion exchange chromatography, multimodal chromatography, hydrophobic interaction chromatography or hydroxyapatite chromatography. Additionally, in some examples, the fluid is a pool containing harvested cell culture fluid, eluate from depth filtration, eluate from microfiltration, eluate from affinity chromatography, eluate from ion exchange chromatography, eluate from multimodal chromatography, eluate from hydrophobic interaction chromatography, or eluate from hydroxyapatite chromatography. Furthermore, in some examples, the fluid is harvested host cell culture fluid and the unit operation includes depth filtration. Additionally, in some examples, the fluid is harvested host cell culture fluid and the unit operation includes microfiltration. Moreover, in some examples, the fluid is harvested host cell culture fluid and the unit operation includes Protein A affinity chromatography. Furthermore, in some examples, the fluid is Protein A eluant and the unit operation includes depth filtration.

[0056] Moreover, in some examples, the affinity chromatography is Protein A, Protein G, Protein A/G, or Protein L chromatography. Additionally, in some examples, the chromatography is selected from affinity chromatography, Protein A chromatography, ion exchange chromatography, anion exchange 20 chromatography, cation exchange chromatography; hydrophobic interaction chromatography; mixed modal or multimodal chromatography, or hydroxyapatite chromatography.

[0057] Additionally, in some examples, the unit operation includes depth filtration. Furthermore, in some examples, the unit operation includes microfiltration.

What is Claimed is:

1. An automated system for low pH viral inactivation, comprising:
 - a first vessel;
 - a second vessel;
 - a first pH probe associated with the first vessel and configured to measure the pH of contents of the first vessel;
 - a source of a fluid known or suspected to contain at least one enveloped virus to be transferred to the first vessel;
 - an acid pump configured to pump acid into the first vessel after the fluid is transferred into the first vessel and configured to cease pumping acid into the first vessel responsive to the first pH probe measuring a first pH value that is within a tolerance band of a target pH value for viral inactivation;
 - a transfer pump configured to pump the acidified pool from the first vessel to the second vessel responsive to the first pH probe measuring the first pH value that is below the threshold pH value for viral inactivation, and responsive to the acid pump ceasing to pump acid into the first vessel;
 - a first buffer pump configured to pump a first equilibration buffer, having a first known pH value, into the first vessel responsive to the entire acidified pool being pumped out of the first vessel; and
 - an alert generator configured to:
 - compare a second pH value, measured by the first pH probe after the first equilibration buffer is pumped into the first vessel, to the first known pH value of the first equilibration buffer;
 - determine whether the second pH value measured by the first pH probe is different from the first known pH value of the first equilibration buffer by greater than a threshold pH value; and
 - generate a first alert responsive to the second pH value measured by the first pH probe being different from the first known pH of the first equilibration buffer by greater than the threshold pH value.
2. The automated system for low pH viral inactivation of claim 1, further comprising a source pump configured to pump the fluid into the first vessel from the source based at least in part on a signal indicating that the first vessel is empty.
3. The automated system for low pH viral inactivation of claim 1, wherein the first buffer pump is configured to pump the first equilibration buffer into the first vessel based at least in part on a signal indicating that the first vessel is empty.
4. The automated system for low pH viral inactivation of claim 1, further comprising:
 - a second pH probe associated with the second vessel and configured to measure the pH of contents of the second vessel;
 - a base pump configured to pump base into the second vessel responsive to an elapsed time, from the entire acidified pool being pumped into the second vessel, exceeding a threshold amount of time for reducing a concentration of virus in the acidified pool to a predetermined safe level, and configured to cease pumping base into the second vessel responsive to the second pH probe measuring a first pH value that is within a threshold range of neutral pH values;
 - a discharge pump configured to pump the neutralized viral inactivated pool from the second vessel into a filter for treatment of the neutralized viral inactivated pool;
 - a second buffer pump configured to pump a second equilibration buffer, having a second known pH value, into the second vessel responsive to the entire pool being pumped out of the second vessel; and

wherein the alert generator is further configured to:

compare a second pH value, measured by the second pH probe after the first equilibration buffer is pumped into the second vessel, to the second known pH value of the second equilibration buffer;

determine whether the second pH value measured by the second pH probe is different from the second known pH value of the second equilibration buffer by greater than the threshold pH value; and

generate a second alert responsive to the second pH value measured by the second pH probe being different from the second known pH of the second equilibration buffer by greater than the threshold pH value.

5. The automated system for low pH viral inactivation of claim 4, wherein the first equilibration buffer and the second equilibration buffer are the same equilibration buffer.

6. The automated system for low pH viral inactivation of claim 4, wherein the first equilibration buffer and the second equilibration buffer are distinct equilibration buffers.

7. The automated system for low pH viral inactivation of claim 4, wherein the transfer pump is configured to pump the acidified pool from the first vessel to the second vessel based at least in part on a signal indicating that the second vessel is empty.

8. The automated system for low pH viral inactivation of claim 4, wherein the second buffer pump is configured to pump the second equilibration buffer into the second vessel based at least in part on a signal indicating that the second vessel is empty.

9. The automated system for low pH viral inactivation of claim 4, further comprising:
a third vessel; and
a collection pump configured to pump the filtered pool from the filter to the third vessel.

10. The automated system for low pH viral inactivation of claim 9, wherein the collection pump is configured to pump the filtered pool from the second vessel to the third vessel based at least in part on a signal indicating that the third vessel is empty.

11. The automated system for low pH viral inactivation of claim 1, further comprising:
a first pH probe recalibrator configured to automatically recalibrate the first pH probe responsive to the first alert.

12. The automated system for low pH viral inactivation of claim 1, further comprising one or more additional pH probes associated with the first vessel and configured to measure the pH of contents of the first vessel.

13. The automated system for low pH viral inactivation of claim 4, further comprising one or more additional pH probes associated with the second vessel and configured to measure the pH of contents of the second vessel.

14. The automated system for low pH viral inactivation of claim 4, further comprising:
a second pH probe recalibrator configured to automatically recalibrate the second pH probe responsive to the second alert.

15. The automated system for low pH viral inactivation of claim 4, further comprising:
an operator display configured to display one or more of the first alert or the second alert to an operator associated with the system.

~~16. The automated system of low pH viral inactivation of claim 1, wherein the acid is selected from formic acid, acetic acid, citric acid, and phosphoric acid at concentrations suitable to ensure viral inactivation.~~

~~17. The automated system of low pH viral inactivation of claim 1, wherein the threshold pH for viral inactivation is from pH 2 to 4.~~

~~18. The automated system of low pH viral inactivation of claim 1, wherein the chromatography elution pool is exposed to the acid for less than 30 minutes prior to neutralization.~~

~~19. The automated system of low pH viral inactivation of claim 1, wherein the base is Tris base at a concentration of 2M.~~

~~20. The automated system of low pH viral inactivation of claim 1, wherein the threshold range of neutral pH values is from pH 4.5 to 6.~~

~~21. The automated system of low pH viral inactivation of claim 1, wherein the low pH viral inactivation is conducted at a temperature of 5 to 25 °C.~~

~~22~~16. The automated system of low pH viral inactivation of claim 1, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a holding vessel.

~~23~~17. The automated system of low pH viral inactivation of claim 1, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a depth filter.

~~24~~18. The automated system of low pH viral inactivation of claim ~~23~~17, wherein following depth filtration, the neutralized viral inactivated eluate is transferred to a sterile filter.

~~25~~19. The automated system of low pH viral inactivation of claim 1, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred a first polish chromatography column.

~~26~~20. An automated method of low pH viral inactivation, the method comprising:
adding a pool to a first vessel;
adding acid to the first vessel;
measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel;
ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel;
transferring the pool from the first vessel to a second vessel;

filling the first vessel with an equilibration buffer having a known pH value;
measuring, by the first pH probe, a second pH value associated with the first vessel;
comparing the second measured pH value associated with the first vessel to the known pH value of the equilibration buffer;
determining whether the second measured pH value associated with the first vessel is different from the known pH value of the equilibration buffer by greater than a threshold pH value; and
generating a first alert responsive to the second measured pH value associated with the first vessel being different from the known pH value of the equilibration buffer by greater than the threshold pH value.

~~27. The automated method of low pH viral inactivation of claim 26, wherein transferring the pool into the first vessel is based at least in part on receiving a signal indicating that the first vessel is empty.~~

~~28. The automated system for low pH viral inactivation of claim 26, wherein filling the first vessel with the equilibration buffer is based at least in part on receiving a signal indicating that the first vessel is empty.~~

~~29. The automated method of low pH viral inactivation of claim 26, further comprising:
adding base to the second vessel after an elapsed time after the transfer of the pool to the second vessel exceeds a threshold amount of time for reducing a concentration of virus in the pool to a predetermined safe level;
measuring, by a second pH probe associated with the second vessel, a first pH value associated with the second vessel;
ceasing, based on the first measured pH value associated with the second vessel being within a threshold range of neutral pH values, the addition of base to the second vessel;
transferring the pool from the second vessel to a filter for treatment of the neutralized viral inactivated pool;
filling the second vessel with the equilibration buffer having the known pH value;
measuring, by a second pH probe associated with the second vessel, a second pH value associated with the second vessel;
comparing the second measured pH value associated with the second vessel to the known pH value of the equilibration buffer;
determining whether the second measured pH value associated with the second vessel is different from the known pH value of the equilibration buffer by greater than a threshold pH value; and
generating a second alert responsive to the second measured pH value associated with the second vessel being different from the known pH value of the equilibration buffer by greater than the threshold pH value.~~

~~30. The automated method of low pH viral inactivation of claim 29, wherein transferring the acidified pool from the first vessel to the second vessel based at least in part on receiving a signal indicating that the second vessel is empty.~~

~~31. The automated method of low pH viral inactivation of claim 29, wherein filling the second vessel with the equilibration buffer is based at least in part on a receiving a signal indicating that the second vessel is empty.~~

~~32. The automated method of low pH viral inactivation of claim 29, further comprising:
transferring the pool from the filter to a third vessel.~~

33. ~~The automated method for low pH viral inactivation of claim 32, wherein transferring the pool from the filter to the third vessel is based at least in part on a receiving a signal indicating that the third vessel is empty.~~

34. ~~The automated method of low pH viral inactivation of claim 26, further comprising:
recalibrating the first pH probe responsive to the first alert.~~

35. ~~The automated method of low pH viral inactivation of claim 34, wherein the recalibrating is an automatic recalibrating.~~

36. ~~The automated method of low pH viral inactivation of claim 34, wherein the recalibrating is a manual recalibrating.~~

37. ~~The automated method of low pH viral inactivation of claim 29, further comprising:
recalibrating the second pH probe responsive to the second alert.~~

38. ~~A method for inactivating enveloped viruses during purification of a recombinant protein of interest, comprising:
obtaining a fluid known or suspected to contain at least one enveloped virus;
subjecting the fluid to one or more of the following steps at a concentration and for a time sufficient to cause viral inactivation:
adding the fluid to a first vessel;
adding acid to the first vessel;
measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel;
ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel;
transferring the fluid from the first vessel to a second vessel;
filling the first vessel with an equilibration buffer having a known pH value;
measuring, by the first pH probe, a second pH value associated with the first vessel;
comparing the second measured pH value associated with the first vessel to the known pH value of the equilibration buffer;
determining whether the second measured pH value associated with the first vessel is different from the known pH value of the equilibration buffer by greater than a threshold pH value; and
generating a first alert responsive to the second measured pH value associated with the first vessel being different from the known pH value of the equilibration buffer by greater than the threshold pH value; and
subjecting the neutralized viral inactivated fluid to at least one unit operation which includes at least a filtration step or a chromatography step.~~

39. ~~The method according to claim 38, wherein adding the fluid to the first vessel is based in part on receiving a signal indicating that the first vessel is empty.~~

40. ~~The method according to claim 38, wherein transferring the fluid from the first vessel to the second vessel is based in part on receiving a signal indicating that the second vessel is empty.~~

41. ~~The method according to claim 38, wherein filling the first vessel with the equilibration buffer is based in part on receiving a signal indicating that the first vessel is empty.~~

42. ~~The method according to claim 38, wherein the fluid comprises a recombinant protein of interest.~~

43. ~~The method according to claim 38, wherein the fluid is harvested host cell culture fluid.~~

44. ~~The method according to claim 38, wherein the fluid is from an effluent stream, eluate, pool, storage or hold from a unit operation comprising a harvest step, a filtration step or a chromatography step.~~

45. ~~The method according to claim 44, wherein the fluid is eluate collected from depth filtration, microfiltration, affinity chromatography, ion exchange chromatography, multimodal chromatography, hydrophobic interaction chromatography or hydroxyapatite chromatography.~~

46. ~~The method according to claim 44, wherein the fluid is a pool containing harvested cell culture fluid, eluate from depth filtration, eluate from microfiltration, eluate from affinity chromatography, eluate from ion exchange chromatography, eluate from multimodal chromatography, eluate from hydrophobic interaction chromatography, or eluate from hydroxyapatite chromatography.~~

47. ~~The method according to claim 46, wherein the affinity chromatography is Protein A, Protein G, Protein A/G, or Protein L chromatography.~~

48. ~~The method according to claim 38, wherein the chromatography is selected from affinity chromatography, Protein A chromatography, ion exchange chromatography, anion exchange chromatography, cation exchange chromatography; hydrophobic interaction chromatography; mixed modal or multimodal chromatography, or hydroxyapatite chromatography.~~

49. ~~The method according to claim 38, wherein the fluid is harvested host cell culture fluid and the unit operation includes depth filtration.~~

50. ~~The method according to claim 38, wherein the fluid is harvested host cell culture fluid and the unit operation includes microfiltration.~~

51. ~~The method according to claim 38 wherein the fluid is harvested host cell culture fluid and the unit operation includes Protein A affinity chromatography.~~

52. ~~The method according to claim 38, wherein the fluid is Protein A eluant and the unit operation includes depth filtration.~~

53. ~~The method according to claim 38, wherein the unit operation includes depth filtration.~~

54. ~~The method according to claim 38, wherein the unit operation includes microfiltration.~~

55.——An automated system for low pH viral inactivation, comprising:
a first vessel;
a second vessel;
a first pH probe associated with the first vessel and configured to measure the pH of contents of the first vessel;
a source of a fluid known or suspected to contain at least one enveloped virus to be transferred to the first vessel;
an acid pump configured to pump acid into the first vessel after the fluid is transferred into the first vessel and configured to cease pumping acid into the first vessel responsive to the first pH probe measuring a first pH value that is within a tolerance band of a target pH value for viral inactivation;
a transfer pump configured to pump the acidified pool from the first vessel to the second vessel responsive to the first pH probe measuring the first pH value that is below the threshold pH value for viral inactivation, and responsive to the acid pump ceasing to pump acid into the first vessel;
a second pH probe associated with the second vessel and configured to measure the pH of contents of the second vessel;
a base pump configured to pump base into the second vessel responsive to an elapsed time, from the entire acidified pool being pumped into the second vessel, exceeding a threshold amount of time for reducing a concentration of virus in the acidified pool to a predetermined safe level, and configured to cease pumping base into the second vessel responsive to the second pH probe measuring a first pH value that is within a threshold range of neutral pH values; and
a discharge pump configured to pump the neutralized viral inactivated pool from the second vessel into a filter for treatment of the neutralized viral inactivated pool.

56.——The automated system for low pH viral inactivation of claim 55, further comprising source pump configured to pump the fluid into the first vessel from the source based at least in part on a signal indicating that the first vessel is empty.

57.——The automated system for low pH viral inactivation of claim 55, wherein the transfer pump is configured to pump the acidified pool from the first vessel to the second vessel based at least in part on a signal indicating that the second vessel is empty.

58.——The automated system for low pH viral inactivation of claim 55, further comprising:
a third vessel; and
a collection pump configured to pump the filtered pool from the filter to the third vessel.

59.——The automated system for low pH viral inactivation of claim 58, wherein the collection pump is configured to pump the filtered pool from the second vessel to the third vessel based at least in part on a signal indicating that the third vessel is empty.

60.——The automated system for low pH viral inactivation of claim 55, further comprising one or more additional pH probes associated with the first vessel and configured to measure the pH of contents of the first vessel.

61.——The automated system for low pH viral inactivation of claim 55, further comprising one or more additional pH probes associated with the second vessel and configured to measure the pH of contents of the second vessel.

62. ~~The automated system of low pH viral inactivation of claim 55, wherein the acid is selected from formic acid, acetic acid, citric acid, and phosphoric acid at concentrations suitable to ensure viral inactivation.~~

63. ~~The automated system of low pH viral inactivation of claim 55, wherein the threshold pH for viral inactivation is from pH 2 to 4.~~

64. ~~The automated system of low pH viral inactivation of claim 55, wherein the chromatography elution pool is exposed to the acid for less than 30 minutes prior to neutralization.~~

65. ~~The automated system of low pH viral inactivation of claim 55, wherein the base is Tris base at a concentration of 2M.~~

66. ~~The automated system of low pH viral inactivation of claim 55, wherein the threshold range of neutral pH values is from pH 4.5 to 6.~~

67. ~~The automated system of low pH viral inactivation of claim 55, wherein the low pH viral inactivation is conducted at a temperature of 5 to 25 °C.~~

68. ~~The automated system of low pH viral inactivation of claim 55, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a holding vessel.~~

69. ~~The automated system of low pH viral inactivation of claim 55, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred to a depth filter.~~

70. ~~The automated system of low pH viral inactivation of claim 58, wherein following depth filtration, the neutralized viral inactivated eluate is transferred to a sterile filter.~~

71. ~~The automated system of low pH viral inactivation of claim 55, wherein neutralized viral inactivated chromatography elution pool from the second vessel is transferred a first polish chromatography column.~~

72. ~~An automated method of low pH viral inactivation, the method comprising:
adding a pool to a first vessel;
adding acid to the first vessel;
measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel;
ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel;
transferring the pool from the first vessel to a second vessel;
adding base to the second vessel after an elapsed time after the transfer of the pool to the second vessel exceeds a threshold amount of time for reducing a concentration of virus in the pool to a predetermined safe level;
measuring, by a second pH probe associated with the second vessel, a first pH value associated with the second vessel;~~

ceasing, based on the first measured pH value associated with the second vessel being within a threshold range of neutral pH values, the addition of base to the second vessel; and
transferring the pool from the second vessel to a filter for treatment of the neutralized viral inactivated pool.

73.——The automated method of low pH viral inactivation of claim 72, wherein transferring the pool into the first vessel is based at least in part on receiving a signal indicating that the first vessel is empty.

74.——The automated method of low pH viral inactivation of claim 72, wherein filling the second vessel with the equilibration buffer is based at least in part on a receiving a signal indicating that the second vessel is empty.

75.——The automated method of low pH viral inactivation of claim 72, further comprising:
transferring the pool from the filter to a third vessel.

76.——The automated method for low pH viral inactivation of claim 75, wherein transferring the pool from the filter to the third vessel is based at least in part on a receiving a signal indicating that the third vessel is empty.

77.——A method for inactivating enveloped viruses during purification of a recombinant protein of interest, comprising:
obtaining a fluid known or suspected to contain at least one enveloped virus;
subjecting the fluid to one or more of the following steps at a concentration and for a time sufficient to cause viral inactivation:
adding the fluid to a first vessel;
adding acid to the first vessel;
measuring, by a first pH probe associated with the first vessel, a first pH value associated with the first vessel;
ceasing, based on the first measured pH value associated with the first vessel being within a tolerance band of a target pH value for viral inactivation, the addition of acid to the first vessel;
transferring the fluid from the first vessel to a second vessel;
adding base to the second vessel;
measuring, by a second pH probe associated with the first vessel, a second pH value associated with the second vessel;
ceasing, based on the second measured pH value associated with the second vessel being within a tolerance band of a target pH value for neutrality, the addition of base to the second; and
subjecting the neutralized viral inactivated fluid to at least one unit operation which includes at least a filtration step or a chromatography step.

78.——The method according to claim 77, wherein adding the fluid to the first vessel is based in part on receiving a signal indicating that the first vessel is empty.

79.——The method according to claim 77, wherein transferring the fluid from the first vessel to the second vessel is based in part on receiving a signal indicating that the second vessel is empty.

~~80. The method according to claim 77, wherein filling the first vessel with the equilibration buffer is based in part on receiving a signal indicating that the first vessel is empty.~~

~~81. The method according to claim 77, wherein the fluid comprises a recombinant protein of interest.~~

~~82. The method according to claim 77, wherein the fluid is harvested host cell culture fluid.~~

~~83. The method according to claim 77, wherein the fluid is from an effluent stream, eluate, pool, storage or hold from a unit operation comprising a harvest step, a filtration step or a chromatography step.~~

~~84. The method according to claim 83, wherein the fluid is eluate collected from depth filtration, microfiltration, affinity chromatography, ion exchange chromatography, multimodal chromatography, hydrophobic interaction chromatography or hydroxyapatite chromatography.~~

~~85. The method according to claim 83, wherein the fluid is a pool containing harvested cell culture fluid, eluate from depth filtration, eluate from microfiltration, eluate from affinity chromatography, eluate from ion exchange chromatography, eluate from multimodal chromatography, eluate from hydrophobic interaction chromatography, or eluate from hydroxyapatite chromatography.~~

~~86. The method according to claim 77, wherein the affinity chromatography is Protein A, Protein G, Protein A/G, or Protein L chromatography.~~

~~87. The method according to claim 77, wherein the chromatography is selected from affinity chromatography, Protein A chromatography, ion exchange chromatography, anion exchange 20 chromatography, cation exchange chromatography; hydrophobic interaction chromatography; mixed modal or multimodal chromatography, or hydroxyapatite chromatography.~~

~~88. The method according to claim 77, wherein the fluid is harvested host cell culture fluid and the unit operation includes depth filtration.~~

~~89. The method according to claim 77, wherein the fluid is harvested host cell culture fluid and the unit operation includes microfiltration.~~

~~90. The method according to claim 77 wherein the fluid is harvested host cell culture fluid and the unit operation includes Protein A affinity chromatography.~~

~~91. The method according to claim 77, wherein the fluid is Protein A eluant and the unit operation includes depth filtration.~~

~~92. The method according to claim 77, wherein the unit operation includes depth filtration.~~

~~93. The method according to claim 77, wherein the unit operation includes microfiltration.~~