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Purity of the precipitate: co-precipitation and post precipitation, Estimation of barium sulphate,Basic Principles,methods and application of diazotisation titration. 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Acid base titration can be defined as a titration reaction which takes place between an acid and base in which we determine the concentration of an acid or base by completely neutralising it with an acid or a base with a known concentration. These reactions help us determine the concentration of an acid or a base in terms of normality, molarity, molality, etc. In the process, we also make use of the PH indicator (phenolphthalein) to monitor the process of the reaction by knowing the colour of the pH indicator. The pH indicator phenolphthalein has a pH range between 8 to 10, which makes it an overall good indicator for the process of titration.TitrationTitration requires solid or weak acid or a base for knowing the concentration or determining whether the given solution is a strong or weak acid or base. The experiment involves a titrant and an acid or a base.There can be different types of titrations like:1) strong acid strong base2) strong acid weak base3) weak acid weak base4) weak acid strong baseThere is an equivalence point at which the acid and base neutralise exactly, i.e. there are equal numbers of moles of hydrogen and hydroxides. Now accordingly, the experimentalist can draw the titration curve. There is a use of an indicator through which we can detect the change and tell whether the solution has been neutralised or not. The common indicators used are phenolphthalein and methyl orange, and sometimes even malachite green can also be used. But for pharmaceutical industries, electrodes of pH metres are immersed as an indicator.Experiment1) Place an unknown concentration of an acid or a base in a beaker and add an indicator or electrode of a pH metre.2) Fill the buret with a titrating solution of known concentration, and this is called the titrant or standard solution.3) Standard solution is added slowly by measurements in the solution in the beaker until the equivalence is attained. Now, by calculation using mathematical formulas, the concentration of the unknown solution can be obtained. Importance of titration in the pharmaceutical industryThere are many reasons for using acid-base titration in pharmaceuticals, for example, analysis, quality control and product development, content analysis by redox titration and purity analysis of the pharmaceutical active substances. Purity analysis is used for determining active ingredients in the products, or example, acetylsalicylic acid in aspirin or vitamin c in multivitamin tablets. It is also used for the purity control of the drug additives used for the synthesis of medicinal preparations. One example is purity control of ephedrine hydrochloride, which is used as a cough syrup for the treatment of bronchial asthma. There are many methods of titration in pharmaceuticals:1) Titration in non-aqueous solutions: It is for weak acids and bases a) Non-aq. titration of bases: for amines determinationRNH2 + HClO4 – R.NH3+ + ClO4-Direct titration of amines: Benzyl nicotinate used in rheumatismProtonated amine salts titration-Diphenylpyraline Hydrochloride 3 is an antihistamine.HClO4 in glacial acetic acid is used in non-aqueous titration of amines.Solvents used can be acetic acid, acetic anhydride, methanol and acetone.Titrants used can be perchloric acid, perchloric acid in glacial acetic acid, and perchloric acid in acetic anhydride.b) Non-aq. titration of acids: for aromatic acids determinationROH + (R')4NOH – RO- + N(CH3)4+ + H2ODirect titration of phenol and its derivatives-Phenol and Rutidosolvents used can be 1,2-Diaminoethane, dimethylformamide, and acetone.Titrants used can be TMAH, TBAH, and 2-propanol/methanol mixtures.2) Titration in aqueous solution: It is an analytic process in which water is used as a solvent for the determination of the amount of desired substance.Buffer capacityIt is a solution which is capable of keeping the pH value constant when a small amount of either acid or a base is added; its resistance to pH changes is known as buffer capacity. This type of solution can be prepared by combining a weak acid and its salt with a strong base and vice-versa. The buffer capacity usually depends on the amount of substance of given weak acid and its conjugate base in the buffer solution.ConclusionIn this article, we got to know about titration and its uses in the pharmaceutical industry. There are many other types of titration, but the objective of this article was to focus on acid-base titration and its importance in the pharmaceutical industry, the uses of acid bases titration in the pharmaceutical industry and its analysis. We also discussed different types of acid-base titration like the titration in a non-aqueous and aqueous solvent and their examples concerning diseases, and lastly, about the buffer solution. 2. ACID-BASE THEORY:- ARRHENIUS THEORY:- Acid is a substance which gives hydrogen ion (H+) in an aqueous medium. Base is substance which gives hydroxy ions (OH-) in an aqueous medium. These theory are only applicable for aqueous titration. LOWRY AND BRONSTED'S THEORY:- Acid is a substance which accept a proton (hydrogen ion) base is a substance which donate a proton. LEWIS'S THEORY:- Acid is a substance which accept electron pair. Base is a substance which donate electron pair. ACID BASE TITRATION 3. USANOVICH THEORY:- Acid is a chemical species that reacts with a base thereby donating cations or accepting anions or electrons. Base is a chemical species that reacts with an acid thereby donating anions or electrons or accepting cations. LUX-FLOOD CONCEPT:- Acid is a substance which accept oxide-ion. Base is the substance which donate oxide ions. 4. LAW OF MASS ACTION The rate of a chemical reaction is proportional to the active masses of the reacting substance. A + B + C + D According to law of mass action V_f = k1[A][B] V_b = k2[C][D] V_f = Velocity of forward reaction V_b = Velocity of backward reaction [A], [B], [C], [D] is equal to molar concentration of A, B, C & D K1 & K2 are constant At equilibrium state, velocity of forward reaction is equal to velocity of backward reaction V_f = V_b 5. k1[A][B] = k2[C][D] k1 and k2 are constant hence, where k is the equilibrium constant of the reaction. where, a,b,c.....and p,q,r are the no. of molecules of reacting species. 7. Dissociation concept of acid and base:- A strong acid dissociate (or ionizes) completely in aqueous solution to form hydronium ions (H3O+) HCl + H2O H3O + Cl- A weak acid does not dissociate (or ionizes) completely in aqueous solution to form hydronium ions (H3O+) CH3COOH + H2O CH3COOH + H3O+ A strong base dissociate completely in aqueous solution to form hydronium ions (OH-) NaOH Na+ + OH- A weak base does not dissociate completely in aqueous solution to form hydronium ions (OH-) NHAOH NH4+ + OH- 8. EXAMPLES OF WEAK/STRONG ACID:- TYPE EXAMPLES Strong acids hydrochloric acid (HCl), sulfuric acid (H2SO4), nitric acid (HNO3) Weak Acids acetic acid (CH3COOH), hydrofluoric acid (HF), oxalic acid (COOH)2 Strong Bases sodium hydroxide (NaOH), potassium hydroxide (KOH), lithium hydroxide (LiOH) Weak Bases Liquid ammonia(NH3) 9. ACID BASE TITRATION An acid-base titration is a quantitative analysis of acid and bases by which known concentration of an acid or base i.e. titrant neutralizes the unknown concentration of an acid or base i.e. titrand. It is based on the neutralization reaction between acid and bases that form water by combination of H3O+ ions, therefore known as water-forming reaction. HA(ACID) + H2O H3O+ + OH- H2O The estimation of an alkaline solution by using a standard acid solution is known as Alkalimetry. The titration progress can be monitored by visual indicators, PH electrodes or both. 10. TYPE OF TITRATION & NEUTRALIZATION CURVE:- The mechanism of neutralization can be understood by studying the change in the hydrogen ion concentration during the course of appropriate titration. 1) The equivalence point of an acid-base reaction (the point at which the amounts of acid and of base are just sufficient to cause complete neutralization). 2) The pH of the solution at equivalence point is dependent on the strength of the acid and strength of the base used in the titration. For strong acid-strong base titration, pH = 7 at equivalence point For weak acid-strong base titration, pH > 7 at equivalence point For strong acid-weak base titration, pH < 7 at equivalence point 11. The neutralization curve can be categorized into four classes, which enlist below:- 1) Titration of a strong acid with strong base 2) Titration of a weak acid with a strong base 3) Titration of a strong acid with a weak base 4) Titration of a weak base with a weak acid 12. 1) TITRATION OF A STRONG ACID WITH STRONG BASE:- Suppose analyte is hydrochloric acid HCl (strong acid) and the titrant is sodium hydroxide NaOH (strong base). If we start plotting the pH of the Titration curve of a strong acid with a strong base. 13. Point 1:- At point 1, No NaOH added yet, so the pH of the analyte is low (because of it predominantly contains H3O+ from dissociation of HCl). HCl + H2O H3O + Cl- As NaOH is added dropwise, H3O+ slowly starts getting consumed by OH- produced by dissociation of NaOH. Analyte is still acidic due to predominance of H3O+ ions. Point 2:- This is the pH recorded at a time point just before complete neutralization takes place. 14. Equivalent point Point 3:- This is the equivalence point where, moles of NaOH added is equal to moles of HCl in the analyte. At this point, OH- ions are completely neutralized by OH- ions. The solution only has salt (NaCl) and water and therefore the pH is neutral. pH 7. HCl + NaOH NaCl + H2O Equivalent point Point 4:- Addition of NaOH continues. Ws, pH starts becoming basic because HCl has been completely neutralized and now excess of OH- ions are present in the solution (from dissociation of NaOH). After this NaOH react with indicator and produce color that is end point. NaOH Na+ + OH- 15. 2) TITRATION OF A WEAK ACID WITH A STRONG BASE:- Let's assume our analyte is acetic acid CH3COOH (weak acid) and the titrant is sodium hydroxide NaOH (strong base). If we start plotting the pH of the analyte against the volume of NaOH that we are adding from the burette, we will get a titration curve as shown below. 16. Point 1:- No NaOH added yet, so the pH of the analyte is low (it predominantly contains H3O+ from dissociation of CH3COOH). CH3COOH + H2O CH3COOH + H3O+ As NaOH is added drop wise, H3O+ slowly starts getting consumed by OH- produced by dissociation of NaOH). But analyte is still acidic due to predominance of H3O+ ions. Point 2:- This is the pH recorded at a time point just before complete neutralization takes place. 17. Equivalent point Point 3:- This is the equivalence point. At this point, moles of NaOH added is equal to moles of CH3COOH in the analyte. The H3O+ are completely neutralized by OH-ions. The solution contains only CH3COONa salt and H2O. CH3COOH + NaOH CH3COONa + H2O+ Equivalent point In the case of a weak acid versus a strong base, the pH is not neutral at the equivalence point. The solution is basic (pH~9) at the equivalence point. Let's reason this out. 18. Point 4:- Addition of NaOH continues, pH starts becoming basic because CH3COOH has been completely neutralized and now excess of OH- ions are present in the solution (from dissociation of NaOH). After this NaOH react with indicator and produce color that is end point. 19. 3) TITRATION OF A STRONG ACID WITH A WEAK BASE:- Suppose our analyte is hydrochloric acid HCl (strong acid) and the titrant is ammonia NH3 (weak base). If we start plotting the pH of the analyte against the volume of NH3 are added from the burette, we will get a titration curve as shown below. 20. Point 1:- No NH3 added yet, so the pH of the analyte is low (it predominantly contains H3O+ from dissociation of HCl). HCl + H2O H3O + Cl- As NH3 is added drop wise, H3O+ slowly starts getting consumed by NH3. Analyte is still acidic due to predominance of H3O+ ions. NH3 + H3O NH4+ + H2O Point 2:- This is the pH recorded at a time point just before complete neutralization takes place. 21. Point 3:- This is the equivalence point. At this point, moles of NH3 added is equal to the moles of HCl in the analyte. The H3O+ ions are completely neutralized by NH3. In the case of a weak base versus a strong acid, the pH is not neutral at the equivalence point. The solution is in fact acidic (pH~5.5) at the equivalence point. At the equivalence point, the solution only has ammonium ions NH4 and chloride ions Cl-. Where ammonium ion NH4 is the conjugate acid of the weak base NH3. So NH4 is a relatively strong acid (weak base NH3 has a strong conjugate acid), and thus NH4 react with H2O to produce hydronium ions making the solution acidic. NH4+ + H2O NH3 + H3O+ Makes solution acidic at equivalent point 22. Point 4:- After the equivalence point, NH3 addition continues and is in excess, so the pH increases. NH3 is a weak base so the pH is above 7. After this NaOH react with indicator and produce color that is end point. 23. 4) TITRATION OF A WEAK BASE WITH A WEAK ACID:- Suppose our analyte is NH3 (weak base) and the titrant is acetic acid CH3COOH (weak acid). If we start plotting the pH of the analyte against the volume of acetic acid that we are adding from the burette, we will get a titration curve as shown below. 24. There isn't any steep bit in this plot. There is just because of a 'point of inflexion' at the equivalence point. Lack of any steep change in pH throughout the titration renders. Titration of a weak base versus a weak acid is difficult, and not much information can be extracted from such a curve. The chief feature of the curve is that change of PH near equivalent point and during whole titration is very gradual hence the end point cannot be detected by ordinary indicator so the desired accuracy and precision of the analysis. First Year B Pharm Notes, Syllabus, Books, PDF Subjectwise/TopicwiseSuggested readings: Titration is a process commonly used to quantify a substance in a solution. One of the more popular analytical techniques, you may remember performing titration experiments at school. But this simple process is precise enough to be used in a wide range of industrial applications. Most notably, titration is used in the pharmaceutical industry to manufacture products like medicines.Titration is a common technique used in analytical chemistry to determine the concentration of an unknown solution by gradually adding a solution with a known concentration. The reactant of known concentration is added bit by bit until neutralisation is achieved.Neutralisation is usually indicated by a change in the colour of the solution, especially when an indicator is used. A chemical indicator is added to the unknown analyte mixture and is used to signal the end of the titration. For example, when carrying out an acid-base titration, the indicator will change colour when the solution reaches a neutral pH. For a more precise titration process, a measuring device such as a pH meter can be used to determine the point of neutralisation.Most titration experiments are intended to accurately determine the molar proportions necessary for an acidic solution to neutralise an alkaline solution, or vice versa. That said, titrations can be classified into different categories based on the types of reactants and the end product:Acid-base is the most common and well-known type of titration. The reactants are acids and bases, and the final products are salts. The main goal here is to determine the concentration of either the base or the acid. The known reactants are in one solution, which is added to the unknown solution in order to figure out the concentration of the analyte.Visual cues, like colour change and the formation of precipitates, are used to indicate when neutralisation has been achieved. Indicators like litmus, methyl violet, and phenolphthalein are commonly used to determine the endpoint of acid-base titrations. Depending on the type of indicator, when bromothymol blue changes from yellow to blue, an acid-base titration has reached an endpoint. However, it must be noted that the endpoint titration isn't always exactly the same as the equivalence point. The latter is determined by the stoichiometry of the reaction.As the name implies, this type of titration involves an oxidation-reduction reaction. It involves the transfer of electrons from one chemical species to the other. The reducing agent donates electrons, while the oxidising chemical receives the electrons. One good example of a redox reaction is a thermite reaction. For example, this type of reaction happens when the iron atoms in a ferric oxide compound lose oxygen atoms to combine with aluminium atoms, forming Al2O3.To determine the endpoint of a redox reaction, a potentiometer, – or type of resistor – is used. If a potentiometer isn't available, colour changes in the analyte can also be used as reference. Similarly, in some redox titration tests, chemical indicators aren't necessary, but the reactants themselves may change colour. For example, if you use potassium permanganate as an oxidizing agent, a slight persisting colour is an indicator of titration endpoint. Similarly, if you use iodine as an oxidising agent, the endpoint is reached when the deep brown triiodide ion disappears.The colour changes in the reactants, however, may not be sufficient to indicate the endpoint. For instance, if the oxidising agent is potassium dichromate, the reactant may change from orange to green, but this isn't a definite sign that the titration has ended. An indicator like sodium diphenylamine must be added to yield a definite result.Wine is one of the most common commercial substances that requires analysis using the redox titration method. For instance, if you want to know if sulfur dioxide is present in wine, you can use iodine as an oxidising agent and starch as an indicator. Once a blue colour appears, caused by a starch-iodine complex formation, it signals the endpoint of titration.Many titration experiments are necessary in the pharmaceutical industry when analysing substances, including illicit drugs. 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