

1: SwePub - On Distribution Coefficients

In this thesis, different types of chemical and physical distribution coefficients are identified, examined and used to describe the fate and transport pathways of substances in aquatic systems.

Hence the hydrophobicity of a compound as measured by its distribution coefficient is a major determinant of how drug-like it is. More specifically, for a drug to be orally absorbed, it normally must first pass through lipid bilayers in the intestinal epithelium a process known as transcellular transport. For efficient transport, the drug must be hydrophobic enough to partition into the lipid bilayer, but not so hydrophobic, that once it is in the bilayer, it will not partition out again. Pharmacodynamics[edit] In the context of pharmacodynamics what a drug does to the body , the hydrophobic effect is the major driving force for the binding of drugs to their receptor targets. In some cases the metabolites may be chemically reactive. Hence it is advisable to make the drug as hydrophilic as possible while it still retains adequate binding affinity to the therapeutic protein target. Agrochemical research[edit] Hydrophobic insecticides and herbicides tend to be more active. Hydrophobic agrochemicals in general have longer half lives and therefore display increased risk of adverse environmental impact. It is a critical parameter for purification using zone melting , and determines how effectively an impurity can be removed using directional solidification , described by the Scheil equation. The log P of a solute can be determined by correlating its retention time with similar compounds with known log P values. Moreover, since the value of log P is determined by linear regression , several compounds with similar structures must have known log P values, and extrapolation from one chemical class to anotherâ€™”applying a regression equation derived from one chemical class to a second oneâ€™”may not be reliable, since each chemical classes will have its characteristic regression parameters. The method does, however, require the separate determination of the pKa value s of the substance. Electrochemical[edit] Polarized liquid interfaces have been used to examine the thermodynamics and kinetics of the transfer of charged species from one phase to another. Two main methods exist. Here a reaction at a triple interface between a conductive solid, droplets of a redox active liquid phase and an electrolyte solution have been used to determine the energy required to transfer a charged species across the interface. For example, tens of thousands of industrially manufactured chemicals are in common use, but only a small fraction have undergone rigorous toxicological evaluation. Hence there is a need to prioritize the remainder for testing. QSAR equations which in turn are based on calculated partition coefficients can be used to provide toxicity estimates. Other prediction methods rely on other experimental measurements such as solubility. The methods also differ in accuracy and whether they can be applied to all molecules, or only ones similar to molecules already studied. Atom-based Standard approaches of this type, using atomic contributions, have been named by those formulating them with a prefix letter: A conventional method for predicting log P through this type of method is to parameterize the distribution coefficient contributions of various atoms to the over-all molecular partition coefficient, which produces a parametric model. This parametric model can be estimated using constrained least-squares estimation , using a training set of compounds with experimentally measured partition coefficients. While this method is generally the least accurate, the advantage is that it is the most general, being able to provide at least a rough estimate for a wide variety of molecules. It has been shown that the log P of a compound can be determined by the sum of its non-overlapping molecular fragments defined as one or more atoms covalently bound to each other within the molecule. Fragmentary log P values have been determined in a statistical method analogous to the atomic methods least squares fitting to a training set. In addition, Hammett type corrections are included to account of electronic and steric effects. This method in general gives better results than atomic based methods, but cannot be used to predict partition coefficients for molecules containing unusual functional groups for which the method has not yet been parameterized most likely because of the lack of experimental data for molecules containing such functional groups. Molecule mining approaches apply a similarity matrix based prediction or an automatic fragmentation scheme into molecular substructures. Furthermore, there exist also approaches using maximum common subgraph searches or molecule kernels. Log D from log P and pKa For cases where the molecule is un-ionized:

2: Partition coefficient - Wikipedia

NUREG/CRO802 DISTRIBUTION COEFFICIENTS FOR RADIONUCLIDES IN AQUATIC ENVIRONMENTS. II. Studies on Marine and Freshwater Sediment Systems Including the Radionuclides Ru, Cs and Am.

Water demand is the driving force for the operation of municipal water systems. Because water demands are stochastic in nature, water system operation requires an understanding of the amount of water being used, where it is being used, and how this usage varies with time. For most water systems the ratio of the maximum day water demand to the average day water demand ranges from 1. Of course, these values are system specific, and seasonal variations may make these ratios even more extreme Walski et al. Demands may be classified as follows Clark et al. Baseline demands, which usually correspond to consumer demands and unaccounted-for-water associated with average day conditions. Seasonal variations in demand because water use typically varies over the course of the year with higher demands occurring in the warmer months. Fire demands, which may be the most important consideration for water system design. Diurnal variations due to the continuously varying demands which are inherent in water systems. There is a need for research that relates distribution system design to demand in a stochastic framework. Variations in demand have an important influence on water distribution system operation and in the determination of water age which in turn influences water quality, as discussed later in the chapter. From an infrastructure perspective, a water distribution system is an elaborate conveyance structure in which pumps move water through the system, control valves allow water pressure and flow direction to be regulated, and reservoirs smooth out the effects of fluctuating demands flow equalization and provide reserve capacity for fire suppression and other emergencies. All these distribution system components and their operations and complex interactions can Page Share Cite Suggested Citation: Drinking Water Distribution Systems: Assessing and Reducing Risks. The National Academies Press. This, in turn, may lead to serious water quality problems, some of which may threaten public health. One of the most critical components of hydraulic integrity is the maintenance of adequate pressure, defined in terms of the minimum and maximum design pressure supplied to customers under specific demand conditions. Low pressures, caused for example by failure of a pump or valve, may lead to inadequate supply and reduced fire suppression capability or, in the extreme, intrusion of potentially contaminated water. High pressures will intensify wear on valves and fittings and will increase leakage and may cause additional leaks or breaks with subsequent repercussions on water quality. High pressures will also increase external load on water heaters and other fixtures. Pipes and pumps must be sized to overcome the head loss caused by friction at the pipe walls and thus to provide acceptable pressure under specific demands, while sizing of control valves is based on the desired flow conditions, velocity, and pressure differential. A related need is to ensure that pressure fluctuations associated with surge conditions are kept below an acceptable limit. Excessive pressure surges generate high fluid velocity fluctuations and may cause resuspension of settled particles as well as biofilm detachment. A second element of hydraulic integrity is the reliability of supply, which refers to the ability of the system to maintain the desirable flow rate even when components are out of service e. Examples include looping of the pipe network and the development of backup sources to ensure multiple delivery points to all areas. Many water quality parameters change with length of time in the distribution system, a factor directly related to the hydraulic design of the system. For example, chlorine residuals decrease with the increasing age of water and may be completely lost, and trihalomethanes concentrations may increase with time. In addition, higher concentrations of substances may leach from pipe materials and linings if the contact time with the water is increased. Low velocities in pipes create long travel times, resulting in pipe sections where sediments can collect and accumulate and microbes can grow and be protected from disinfectants. Furthermore, sediment deposition will result in rougher pipes with reduced hydraulic capacity. If peak velocity is increased or flow reverses in these pipe sections due to any operational change or shock loading, such as tank filling or draining, valve opening or closing, pump going on- or off-line, unexpected higher system pressure, or hydrant flushing, there is a risk that deposits will be suspended and carried to consumers. Long detention times can also greatly reduce corrosion control

effectiveness by effecting phosphate inhibitors and pH management. Thus, reducing residence time is an important hydraulic issue both in pipes and in storage facilities. A final component of hydraulic integrity is maintaining sufficient mixing and turnover rates in storage facilities. This can lead to bacterial regrowth and other biological changes in the water, including nitrification and taste and odor problems. This chapter discusses the factors that can cause the loss of hydraulic integrity, the consequences of losing hydraulic integrity, how to detect loss of hydraulic integrity, techniques for maintaining hydraulic integrity, and how to recover system hydraulic integrity once it is lost. A loss of hydraulic integrity implies a loss of positive line pressures, flow reversals, rapid changes in velocity, a reduction in hydraulic capacity, a detrimental increase in water residence time, or a combination of these events. Factors causing a loss of system hydraulic integrity include 1 pipe leaks and breaks, 2 rapid changes in pressure and flow conditions, 3 planned maintenance activities and emergencies, 4 tuberculation and scale formation in pipes, and 5 improper operational control.

Pipe Deterioration Pipe deterioration resulting in leaks or breaks can lead to a loss of hydraulic integrity because adequate pressures can no longer be maintained. As discussed in detail in Chapter 4, all pipe materials are vulnerable to some kind of chemical or physical deterioration, and all water mains will require rehabilitation and eventual replacement. Aging pipe infrastructure and chronic water main breaks are a common problem for many water utilities. Analysis of water industry data showed that on average, main breaks occur times per day in the United States Cromwell et al. The condition of distribution system pipes is influenced by material type and age, line pressure, type of soil, installation procedures, and many other factors, making it difficult to predict where breaks and leaks will occur. Chapter 4 discusses the roles of leak detection and condition assessment in determining the current condition of distribution system infrastructure.

Pressure Transients and Changes in Flow Regime Rapid changes in pressure and flow caused by events such as rapid valve closures or pump stoppages and hydrant flushing can create pressure surges of excessive magnitude. High-flow velocities can remove protective scale and tubercles, which will increase the rate of corrosion. Uncontrolled pump shutdown can lead to the undesirable occurrence of water-column separation, which can result in catastrophic pipeline failures due to severe pressure rises following the collapse of the vapor cavities. Vacuum conditions can create high stresses and strains that are much greater than those occurring during normal operating regimes. They can cause the collapse of thin-walled pipes or reinforced concrete sections, particularly if these sections were not designed to withstand such strains. In less drastic cases, strong pressure surges may cause cracks in internal lining, damage connections between pipe sections, and destroy or cause deformation to equipment such as pipeline valves, air valves, or other surge protection devices. Sometimes the damage is not realized at the time, but may cause the pipeline to collapse in the future, especially if combined with repeated transients. Transient pressure and flow regimes are inevitable. All systems will, at some time, be started up, switched off, or undergo rapid flow changes such as those caused by hydrant flushing, and they will likely experience the effects of human errors, equipment breakdowns, earthquakes, or other risky disturbances Wood et al. Figure illustrates typical hydraulic events following a pump trip. Page Share Cite Suggested Citation: There is also evidence that pressure transients can lead to the intrusion of contaminants into the distribution system. Most were caused by the sudden shutdown of pumps at a pump station because of either unintentional e. Locations with the highest potential for intrusion were sites experiencing leaks and breaks, areas of high water table, and flooded air-vacuum valve vaults. The system is pumping drinking water to an elevated storage tank while serving the intermediate customers with adequate pressures. Due to an unexpected power failure, the pump quickly runs down loses speed. This will create a negative pressure wave downsurge that will propagate into the distribution system, putting the customers at a potential intrusion risk due to negative pressures. In addition, it is possible that the pressure drops to the point that a vapor pocket forms adjacent to the pump. Subsequently, this cavity will collapse and produce a large pressure spike that can damage the pipeline and the seals which will make the system even more vulnerable to low pressure events. Planned maintenance activities include supplies going off line e. Examples of emergency situations include earthquakes, hurricanes, power failures, equipment failures, or transmission main failures. All these activities can result in a reduction in system capacity and supply pressure and changes to the flow paths of water within the distribution system.

Tuberculation and Scale The hydraulic capacity of distribution systems can be

compromised by deposits on the internal surface of the pipelines. The deposition of corrosion products in the form of tubercles and other types of scales on the interior of the pipes can seriously clog water lines and thus restrict the flow of water. Scales may also form because metal salts such as calcium carbonate, aluminum silicate, etc. Excessive pressure may be necessary to deliver the required flow of water in pipes with tuberculation and scales, further weakening aging pipes. The reduction in hydraulic capacity is caused by the increases in head loss due to the roughness of the deposits and to the decrease in pipe diameter that they cause. Inadequate Operational Control Historically, utilities have focused on the quality of water leaving the treatment plant, because of regulatory drivers, and on the quantity of water supplied by the distribution system, because of their mission to satisfy water demand and maintain system pressure. Thus, it is not surprising that distribution system operations at many utilities and their associated professionals designers, builders, plumbers, inspectors, etc. There is now greater recognition of the water quality effects of how long water is retained in the various elements of the distribution system. Retention time or water age is strongly related to the characteristics of the system and its operation. For example pipe roughness, which affects water flow and residence time, may be modified by repair or rehabilitation. A particularly important issue that demonstrates the interaction of system operation and water quality is the ability or inability of utilities to ensure adequate mixing intensity and time in storage tanks to minimize short circuiting and to limit residence times to be within acceptable limits. Interestingly, the design of tanks to ensure adequate turnover is required in only 15 of 34 states that responded to a survey of drinking water programs conducted by the Association of State Drinking Water Administrators in March see Table Dealing with these issues is discussed in the context of system operation later in this chapter. Each of these has attendant water quality implications, as described below. The most well documented contamination events are backflow and direct contamination at breaks and repair sites, discussed in Chapter 4. A specific type of backflow event related to a loss of hydraulic integrity is called intrusion, which refers to the entrance of contamination into the water distribution system through leaks caused by corroded areas, cracks, and loose joints because of sustained low or negative pressures or a pressure transient. When a section of the distribution system is depressurized due to a normal shutdown, failure of a main or a pump, routine flushing, or emergency fire-fighting water drawdown, contaminated water can be pulled into the main. For example, during a large fire, a pump is connected to a hydrant. High flows pumped out of the distribution system can result in a significantly reduced water pressure around the withdrawal point. A partial vacuum is created in the system, which can cause suction of contaminated water into the potable water system through nearby leaks. During such conditions, it is possible for water to be withdrawn from nonpotable sources into the distribution system and subsequently distributed to homes and buildings located near the fire. The same conditions can be caused by a water main break. Sustained low pressure events and transient pressure events that lead to intrusion of contaminated water have the potential for substantial water quality and health implications. The potential for intrusion of contaminated groundwater into pipes with leaky joints or cracks seems greatest in systems with pipes below the water table and where pathogens or chemicals are in close proximity to the pipe. As discussed in Chapter 4 , two recent studies Kirmeyer et al. In the event of a large intrusion of pathogens, the disinfectant residual normally sustained in drinking water distribution systems may be insufficient to neutralize contaminated water see Chapter 6 discussion on Adequate Disinfectant Residual. Transient events can also generate high intensities of fluid shear and may cause resuspension of settled particles as well as biofilm detachment. Sedimentation When water is moving slowly through a pipe, particles suspended in the water may settle out into the pipe. They also serve as a food source for bacteria and create a hospitable environment for microbial growth. If not removed these materials may cause water quality deterioration, taste and odor problems, or discoloration of the water. This is particularly evident if the sediments are disturbed stirred up by changes in the flow of water, such as when a main break occurs, a service Page Share Cite Suggested Citation: The normal flow of water through the system will reduce some but not all sediment accumulation over time, and supplemental measures are periodically needed to clear out the system.

3: Extraction and Determination of a Distribution Coefficient (K_d) | MendelSet

KURT C, PICEL et al. through correlations of K_D with water solubility, S_w, and with K_{ow}, the distribution coefficient in the octanol/water system.

Environmental Protection Agency, have been grouped into nine series. These nine broad categories were established to facilitate further development and application of environmental technology. Elimination of traditional grouping was consciously planned to foster technology transfer and a maximum interface in related fields. The nine series are: Environmental Protection Technology 3. Interagency Energy-Environment Research and Development 8. Karickhoff and David S. Environmental Protection Agency, Athens, Georgia, and approved for publication. Mention of trade names or commercial products does not constitute endorsement or recommendation for use. This report discusses specific problems associated with each determination and suggests techniques that can eliminate potential sources of error. In addition, these determinations are prerequisites for many chemical or biological process studies. In response to the accepted need for this type of data and in light of the widespread discrepancies in reported data, this report sets forth conventional methods of measurement and identifies potential sources of measurement error for these three physical properties. Water Solubility 7 4. It is not surprising, then, that these three physical properties are highly correlated and have been interrelated quantitatively. Chiou, ; Leo, ; Briggs, ; Karickhoff, For hydrophobic solutes, these determinations are beset with difficulties that produce widespread discrepancies in studies by different investigators. Contributing sources of error include: No general procedure for measuring these three physical properties is universally adaptable for all organic solutes. Numerous methods are available from the literature, but in most cases, they are tailored to meet time constraints or specific conditions of measurement or to suit specific compounds or sorbents. This document discusses specific problems associated with each determination for hydrophobic solutes that is, water solubility of a few parts per million or less and suggests techniques that can eliminate or circumvent each of the difficulties or potential sources of error. By convention, the ratio of concentrations is expressed as octanol over water. The concentrations are volume referenced that is, mass or moles of chemical per unit volume of liquid with the same units chosen for both phases. The following are extensions or refinements of this method for hydrophobic solutes. Reagent grade octanol is extracted once with 0. The solutes are prepared in octanol at or near saturation concentrations for the solids or approximately 0. A small volume of this octanol solution 1 to 5 milliliters is equilibrated with variable volumes of water determined by the amount of compound required for analysis and thus the analytical sensitivity and water solubility of the compound. Equilibration is achieved by gentle shaking of the sample for approximately 15 minutes. For most solutes, the mixing is done in stainless steel centrifuge tubes with sealable caps. This minimizes solute losses by volatilization and handling that is, sorption to vessels, etc. This phase sampling process is critical. A subsample of the octanol phase usually one half or less of the total phase is withdrawn by pipet and added directly into an analysis cell or diluting solvent suitable for analysis. The remainder of the octanol phase and interfacial solute crystals if present is removed prior to aqueous phase sampling. The extraction solvent should not be allowed to contact the exterior of the pipet stem. These solutes are phase-equilibrated in separatory funnels, and transferred to centrifuge tubes, which have been prerinsed with a portion of the aqueous phase, for the final phase separation. After centrifugation, aliquots are withdrawn from each tube as in the procedure described previously and recombined for extraction and analysis. This is accomplished by dilution of the stock octanol solutions prior to mixing with water. Generally, the distribution coefficient is relatively independent of the solute concentration. If the K_{ow} differs significantly more than one standard deviation for replicate determinations at the two solute levels, an intraphase interfering equilibrium solute association or dissociation is suggested. Changes in speciation must be accounted for and experimental conditions controlled accordingly, such that the partition coefficient of individual chemical species can be determined. Leo. Acid-base "protolyses" equilibria are indicated by pH changes in the aqueous phase as a function of solute concentration. If association aggregation effects are suggested, the monomer K_{ow} can be determined by reducing the solute concentration below the aggregation

threshold concentration, which is experimentally determined by consecutive dilution. All determinations should be done in triplicate. The analysis method should be tailored to the compound in question; chromatographic methods are preferred because of their compound specificity in isolating the parent compound, free from interfering impurities. This problem is especially apparent when the analysis involves radiolabels or DV absorption spectroscopy which can be very nonspecific for many solutes, but becomes far less important when chromatographic analysis is used. Regardless, the presence of contaminants can be easily checked by subjecting the solute-containing octanol phase to consecutive extractions with water. The K_{ow} should remain constant under consecutive extraction. The presence of a more-water-soluble impurity is indicated by a significant increase in K_{ow} . The impurity can be removed by exhaustive extraction with water that is, until the K_{ow} is unchanged by further extractions. Also, chromatographic analysis can potentially isolate the parent compound and thus determine the K_{ow} of the parent species. Only in rare cases would one expect the presence of an impurity to affect significantly the true K_{ow} of a chemical species. The solubility of hydrophobic solutes is decreased substantially by the equilibrium water content. The quantity of crystallized solute produced in this way is more than the amount required to saturate the aqueous phase even at water-to-octanol volume ratios as large as 10. This three-phase system equilibrates rapidly 10 to 20 minutes. After centrifugation 10K, 30 minutes the crystalline phase is largely at the interface of the two liquid phases. The two liquid phases can be sampled free of crystalline contamination as described previously. If the saturation concentrations of the chemical do not exceed 0.1, aqueous phase solute concentrations are within a few percentage points of measured aqueous phase solubilities. Apparently the octanol dissolved in the aqueous phase is 0. For hydrophobic solutes that are unstable in water and therefore cannot be equilibrated with water for long periods, the method provides a good means of getting a water solubility. For water-stable solutes, it provides a quick estimate of the solubility. We have used these three-phase measurements only for polycyclic aromatic hydrocarbons. This technique has not been tested for other families of compounds and has not been published. Briefly, the technique utilizes a chromatographic column, the stationary phase of which contains functional groups either as a coating substance or chemically bound on the surface that simulate the octanol phase. In reverse phase liquid chromatography, the elution volume can be related to the partitioning of solute between the stationary and mobile phases. For solutes whose water solubilities are 1 ppm or greater, water saturated with octanol can be used as the mobile phase and octanol as the stationary phase coating, thereby providing a "direct" measurement of K_{ow} . Solute partitioning in this system can be related to K_{ow} through the use of calibration standards compounds whose structure is similar to the compound to be measured and whose K_{ow} is known. With the continuous evolution of more sensitive liquid chromatographs, this technique promises to become the most precise and reliable method of making partitioning measurements of this type. The technique potentially can circumvent the sources of error in K_{ow} determinations described previously. To date, however, this technique is not available to all who need partitioning information and will not likely be so in the near future. In reality, solubility of crystalline hydrophobic compounds proves to be a most difficult physical property to measure. Hydrophobic liquids generally pose no real problem; water-solute equilibration is rapid less than 8 hours under gentle shaking. Special measurement problems exist for polychlorinated biphenyls and polybrominated biphenyls because the components of these hydrophobic compounds may be crystalline in their pure form. The definition and measurement of water solubility for these compounds are discussed by Dexter, Paris, Schoor, and Haque. For hydrophobic crystalline compounds, the literature abounds with methods or recipes for measuring water solubilities, some of which go so far as detailing the effect on the measurement of size or shape of the container in which the compound is equilibrated Campbell, ; Haque, ; Wauchope, ; Bowman, ; Mackay, ; Mackay, ; Biggar, ; Schoor, A real problem in solubility measurement centers around getting the crystalline material dissolved and equilibrated with the solution, and subsequently, isolating the solution phase, free of microcrystals or large solute aggregates. Handling problems degradation, volatilization, sorption associated with aqueous solutions of these materials are similar to those encountered in K_{ow} determination and will not be redressed here. The time required for compound equilibration is primarily a function of the method used to introduce the crystals into the water. Some methods prescribe adding the crystalline compounds to water as large "chunks"

with subsequent stirring extending for days or even weeks; this is done specifically to minimize the formation of microcrystals that are difficult to remove effectively without perturbation of the solute concentration. Although this time-consuming process may prove necessary for some compounds, it is impractical for many applications and unsuitable for compounds that tend to be unstable in water over the required time frame. A much quicker and more facile procedure for introducing the crystalline material Haque, involves "coating" the compound in excess of the amount required to saturate the water out of a volatile solvent onto the walls of the container used for equilibration. Potential solvents include hexane, isooctane, methylene chloride, ethyl ether, etc. The vessel containing the compound dissolved in a minimal amount of solvent is rotated on its side, allowing the solvent to evaporate and thus plate the solute onto the container walls. The crystalline form varies with the choice of solvent. The best choice achieved by trial and error seems to be that which deposits the most translucent crystalline film on the container. Chalky films produce excessive microcrystal loading of the aqueous phase. After the crystalline compound has been "suitably" coated onto the container, membrane-filtered 0. After 24 hours of agitation, aliquots of aqueous phase from which microcrystals have been removed are analyzed periodically until a maximum solute concentration is achieved. Crystalline material can be removed from the aqueous phase by quiescent settling, centrifuging, or filtering. Quiescent settling requires no additional handling but is time consuming and may not be adequate for many compounds. Centrifugation should be in glass or stainless steel tubes that have been pre-rinsed with aliquots of the aqueous phase. Centrifugation time and speed are rather arbitrary; generally 30 minutes at 10K will suffice. It should be established that additional centrifugation does not remove appreciable compound from the "solution" phase. If filtration is used, consecutive aliquots of filtrate should be analyzed until a constant solute response is obtained, indicative that the solute is not being further sorbed by the filter material. Two or more filters in series may be required in some instances. If filtration or centrifugation are used, care must be taken to avoid appreciable handling-induced temperature fluctuations in the aqueous samples. For many crystalline solutes whose solubilities are a few parts per billion or less, these conventional techniques give at best a coefficient of error of 0. Measurement by two different techniques or in different laboratories commonly agree within a factor of 2. Although this technique remains untested for a wide variety of compounds, its speed and ease-of-use make it most attractive in lieu of the conventional alternatives. In this procedure, a saturated aqueous solution is prepared by passing water through a column filled with glass beads onto which the compound has been "plated" out of a volatile solvent similar to the procedure described in conventional method. A measured aliquot of the saturated solution is passed through a resin column suitable for trapping the compound. The collected compound is then eluted from the resin with an organic solvent or mixture of solvents directly into the chromatograph for solute analysis. This method is fast and potentially circumvents all the difficulties commonly encountered with classical methods. The HPLC technique, however, may well prove to be the most accurate and precise method for this difficult determination.

Ch. 13 On the Subject of Estimating Placer examination; principles and practice Brecht on theatre part one
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