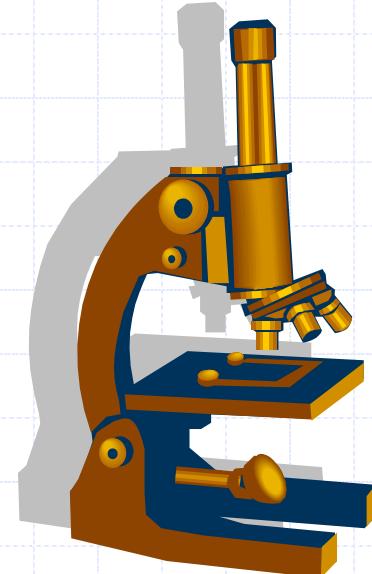


# Improved Combinatorial Group Testing for Real-World Problem Sizes

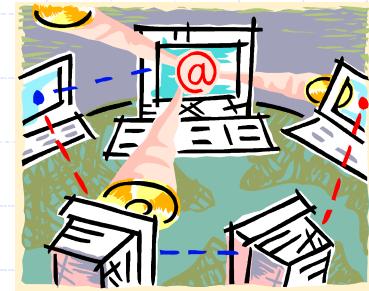


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joint w/ David Eppstein and Dan Hirschberg

# Group Testing



- *Input:*  $n$  items, numbered  $0, 1, \dots, n-1$ , at most  $d$  of which are **defective**.
- *Output:* the indices of all the defective items (or possibly an error condition indicating that more than  $d$  items are defective).
- Items can be grouped into arbitrary test subsets, which can be tested in whole to see they contain a defective item or not.

# 1<sup>st</sup> Motivation: Blood Testing

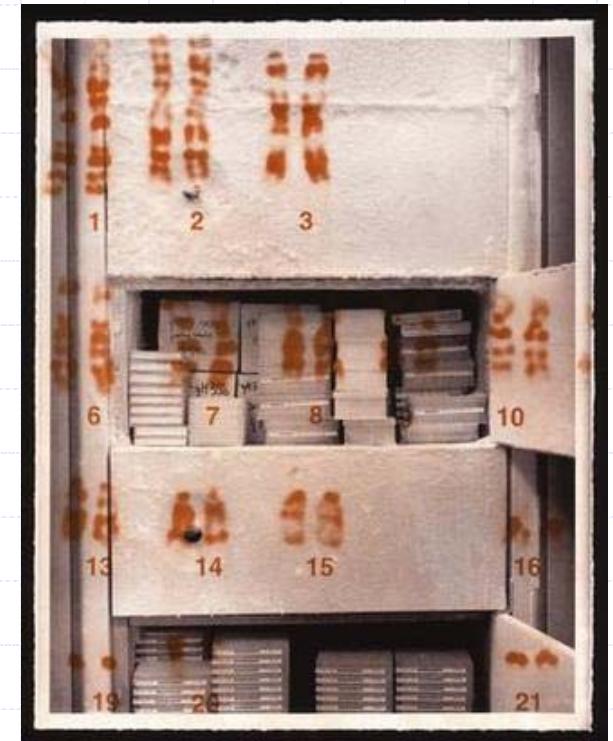
- Items are  $n$  **blood samples** (in the original problem, they were taken from WWII G.I.'s).
- Drops from different samples are **mixed together** and this mixture is tested for disease antigens.
- **Goal:** minimize the total number of tests



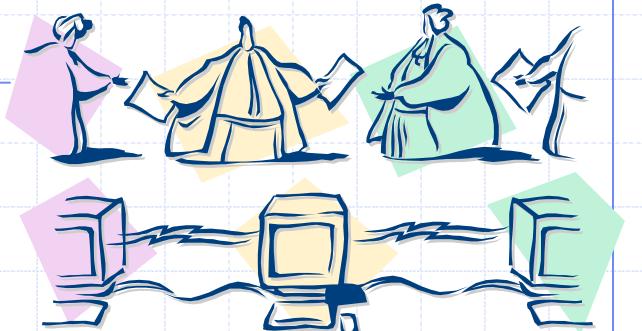
FIGURE 26.—U.S. Army cadets from Marquette University ready to give blood at the Milwaukee, Wis., donor center.

# Modern Applications

- Screening vaccines for contamination
- Filtering clone libraries of DNA sequences (identifying which ones contain a certain DNA sequence)
- Computer security – for data forensics
- Computer fault diagnosis



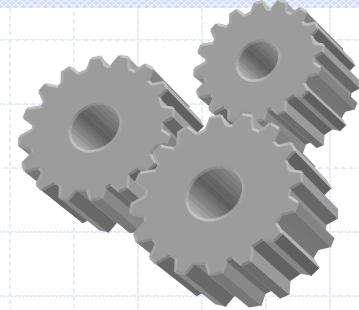
# Testing Regimens



- **Non-adaptive:** All tests must be done in parallel
- **Partially adaptive:** Tests can be done in rounds (e.g., 2 rounds), with the tests in each round done in parallel
- **Fully adaptive:** Tests can be done sequentially

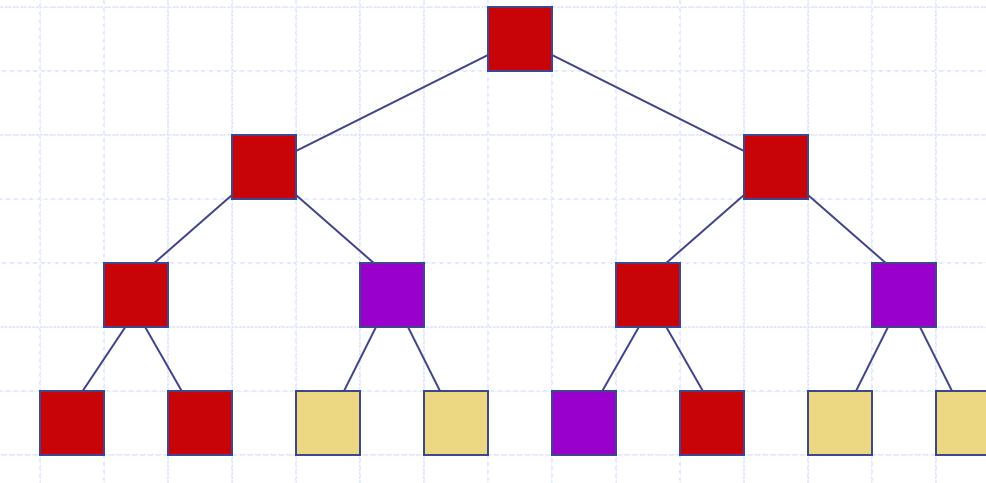
# Efficiency Measures

- $t(n,d)$  = number of tests to identify up to  $d$  defectives among  $n$  items.
  - $t(n,d)$  must be  $\Omega(\min\{n, d \log (n/d)\})$ .
- $A(n,t)$  = analysis time needed to determine which items are defective (after the tests are done).
  - time-optimal is  $A(n,t)$  is  $O(t)$ .
- $S(n,d)$  = sampling rate – the maximum number of tests in which any item may be included.
  - We would like  $S(n,d)$  to be  $O(t(n,d)/d)$

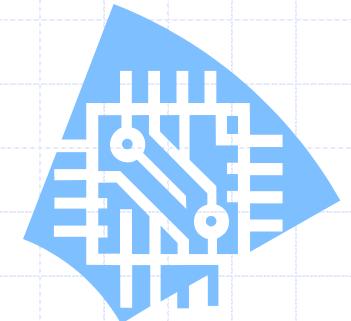


# A Simple Test Regimen

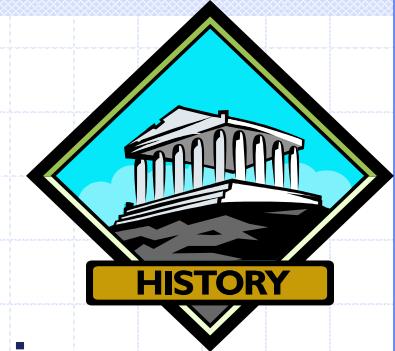
- For the fully-adaptive case:
  - Place a complete binary tree “on top of” the items
  - do a top-down search to defectives
  - will use  $O(d \log (n/d))$  tests



# Another Simple Regimen



- For non-adaptive case when  $d=1$ :
  - Consider item numbers in binary
  - Test  $i$  is set of items w/ bit  $i = 1$
  - Positive (defective) and negative (non-defective) tests identify the binary index of the defective item
  - $t(n,d)$  is  $O(\log n)$
  - $d=2$  and  $d=3$  cases are much harder...



# Previous Related Work

- [Du-Hwang, 00] achieve non-adaptive algorithm with  $t(n,d)$  being  $O(d^2 \log n)$ .
- For two-stage case, [Debonis et al., 03] achieve  $t(n,d) < 7.5 * (d \log (n/d))$
- For  $d=2$ , non-adaptive, [Kautz-Singleton, 64] achieve  $t=3^{q+1}$  for  $n=3^{2^q}$
- For  $d=2$ , non-adaptive, [Macula-Reuter, 98] achieve  $t=(q^2+3q)/2$  for  $n=2^q-1$
- For  $d=3$ , [Du-Hwang, 00] describe an approach that should achieve  $t=18q^2-6q$  for  $n=2^q-1$ .

# Our Results

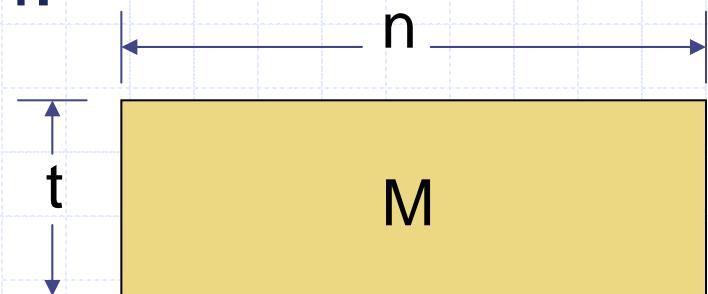
- Chinese Remainder Sieve: An improved non-adaptive test regimen for general  $d$  and  $n < 10^{80}$ 
  - also an improvement for  $n < 10^{57}$  and small  $d$  values
- Rake-and-Winnow: A 2-stage algorithm with a better constant factor (4).
- Improved (and time-optimal) algorithms for the  $d=2$  and  $d=3$  cases.



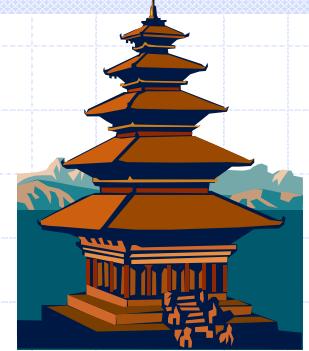
# Matrix View of Testing



- A non-adaptive testing regimen can be viewed as a  $t \times n$  binary matrix  $M$ :
  - $M[i,j] = 1$  if and only if test  $i$  includes item  $j$
- $M$  is **d-disjunct** if the Boolean sum of any  $d$  columns does not contain any other column.
  - An item is defective iff all its tests are positive
- $M$  is **d-separable** if the Boolean sums of each set of at most  $d$  columns are distinct (harder analysis algorithm)



# Chinese Remainder Sieve



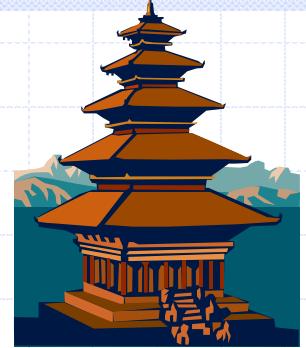
- Let  $\{p_1^{e_1}, p_2^{e_2}, \dots, p_k^{e_k}\}$  be a set of prime powers multiplying to at least  $n^d$ .
- Construct a  $t \times n$  matrix  $M$  as a concatenation of  $k$  submatrices, where  $M_j$  is  $t_j \times n$  matrix where  $t_j = p_j^{e_j}$ .
  - thus,  $t = \sum p_j^{e_j}$ .
- Each row  $q$  of  $M_j$  has a 1 in column  $m$  if  $m \bmod t_j = q$ .
  - if  $q=2$  and  $t_j=3^2=9$ , then row  $q$  has 1's in columns 2, 11, 20, ... .

# Why it Works



- If all tests are positive (defective) for column  $i$ , then  $i$  is defective.
  - For each (true) defective item  $h$ , let  $P_h$  be the product of moduli  $t_j$  associated with tests  $h$  has in common with  $i$ .
  - By a pigeon-hole argument, there is a (true) defective item  $h$  such that  $P_h$  is at least  $n$ .
  - By construction,  $i$  is congruent to the same values that  $h$  is contruent to, modulo each of the prime powers in  $P_h$ .
  - Thus, by Chinese Remainder Theorem,  $i$  is equal to  $h$  modulo a number that is at least  $n$ ; hence,  $i=h$ .

# Analysis



- The number of tests is the sum of the prime products (take each  $e_j=1$  for simplicity)
- We need a bound on the sum of primes whose product is at least  $n^d$ .
- We show that this sum is at most  $(1+o(1))*(2d \ln n)^2/2\ln (2d \ln n)$ .
  - Uses a new bound on the sum of primes, which may be of independent interest.

# Rake-and-Winnow



- Uses a randomized approach motivated by Bloom filtering.
- Also uses a matrix  $M$ , but in 2 rounds
- Given a set  $D$  of  $d$  columns in  $M$  and a column  $j$ , say  $j$  is **distinguishable** from  $D$  if there is a row  $i$  such that  $M[i,j]=1$  but  $M[i,j']=0$  for each  $j'$  in  $D$ .
- $M$  is **( $d,k$ )-resolvable** if, for any  $d$ -sized subset  $D$ , there are fewer than  $k$  columns that are not distinguishable from  $D$ .

# The 2-Round Scheme

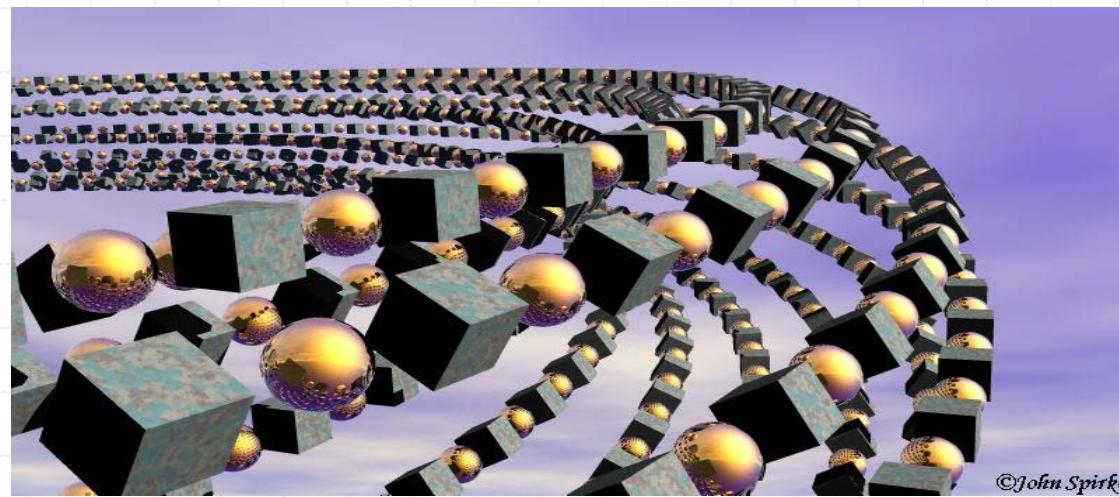


- Use a  $(d,k)$ -resolvable matrix  $M$  in the first round and make a test for each row.
- Discard all the items in negative (non-defective) tests.
- There are at most  $d+k$  remaining items.
- Test each remaining item individually.



# Constructing the Matrix

- Given  $t$  (set in the analysis), let  $M$  be a  $2t \times n$  matrix defined randomly:
  - For each column  $j$ , choose  $t/d$  rows of  $M$  at random and set these entries to 1.
  - that is, we “inject”  $j$  into those  $t/d$  tests



# Analysis



- We show that  $M$  will be completely  $(d,1)$ -resolvable for any particular choice of  $D$ , with high probability, provided  $t > 3.7183d \log n$ .
  - that is, in practice, this will be a single-round scheme with  $t$  being  $O(d \log n)$ .
- There are a lot of possible  $D$ 's however.
- Still, we show that if
$$t > 2d \log (en/d) + \log n,$$
then  $M$  will be  $(d,d)$ -resolvable with high probability.

# Conclusion and Questions

- We have presented improved algorithms for combinatorial group testing for real-world sizes.
- Open problem: design a single non-adaptive scheme that matches or improves our algorithms for small  $n$ , while being asymptotically optimal

